



Capacities/Research Potential  
FP7-REGPOT-2012-2013-1

Project No. 316254  
BASTION

*"From Basic to Translational Research in Oncology"*

Deliverable D3.2

Report on the research activities of 9 experienced researchers

|                          |            |
|--------------------------|------------|
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All reports are available on BASTION Webpage: [www.bastion.wum.edu.pl](http://www.bastion.wum.edu.pl)



## 1. Introduction

The BASTION project is envisioned to allow Medical University of Warsaw (MUW) to become a leading research and clinical oncology centre in Central Europe. One of the objectives realized in WP3 is to build human potential by attracting top-level scientists with international experience in basic and clinical oncology who can contribute to an increase in the quality of research. Deliverable D3.2 corresponds to the task T3.1 in WP3.

## 2. Reports on research activities of experienced researchers

### I. Malgorzata Firczuk, PhD (TEAM of Dominika Nowis)



| DATE (YEARS) | DEGREE/ EXPERIENCE | PLACE   | SUPERVISOR               |
|--------------|--------------------|---|--------------------------|
| 2002         | M.Sc.              | Department of Molecular Biology, Intercollegiate Faculty of Biotechnology, University of Gdańsk, Poland | Prof. Jarosław Marszałek |
| 2003         | M.Sc.              | Department of Microbiology, Faculty of Pharmacy, Medical University of Gdańsk, Poland                   | Prof. Władysław Werel    |
| 2007         | PhD                | Laboratory of Structural Biology, International Institute of Molecular and Cell Biology, Warsaw, Poland | Prof. Matthias Bochtler  |
| 2008-2009    | Postdoc            | Laboratory of Structural Biology, International Institute of Molecular and Cell Biology, Warsaw, Poland | Prof. Matthias Bochtler  |
| 2009-2012    | Postdoc            | Department of Immunology, Medical University of Warsaw, Poland  | Prof. Jakub Gołąb        |
| 2013-now     | Postdoc            | Department of Immunology, Medical University of Warsaw, Poland  | Dr hab. Dominika Nowis   |

#### A. Biosketch (provided by Malgorzata Firczuk)

I gained an extensive academic background in bio-medical sciences, studying biotechnology and pharmacy. In 2001-2002 I was working on my first research project, at the Laboratory of Molecular Biology, Intercollegiate Faculty of Biotechnology, University of Gdańsk, in the group of prof. Jarosław Marszałek. I had learned there basic methods of protein expression in yeasts, principles of protein purification, and earned a master degree. In 2003 I had also completed my studies at Medical University of Gdańsk, Faculty of Pharmacy. I worked on my master thesis project at the Laboratory of Microbiology under the supervision of prof. Władysław Werel. I was studying the interactions between bacterial RNA polymerase and its promoter. My work was awarded as the best master thesis of all Polish pharmacy faculties, presented on the competition organized by Polish Pharmaceutical Society in 2003.

Being more and more fascinated with how proteins work and how protein structure determines its function, for a PhD I moved to Warsaw to work under the supervision of prof. Matthias Bochtler, the head of the Laboratory of Structural Biology at the International Institute of Molecular and Cell Biology. My PhD work



concentrated around structural biology and macromolecular crystallography. I have learned how to produce, purify, crystallize proteins, protein-DNA complexes, and solve their three - dimensional structures by X-ray crystallography. Moreover, I understood how proteins work at the atomic level, what are their mechanisms of interactions, and how the structure influences protein function. My main PhD theme involved peptidoglycan amidases, prokaryotic enzymes that contribute to bacterial pathogenicity. I have managed to obtain the crystal structures for two of them, LytM and MepA. Based on the structures, we designed mutated protein variants to conclude about the mechanisms of action, and studied small molecule interactions in the protein's active site. Driven by the need to work on more medically-oriented research topics, I moved to the Department of Immunology at the Medical University of Warsaw, led by prof. Jakub Golab, working in the field of experimental oncology. I worked as a post-doctoral fellow in the TEAM project "Improvement of antitumor effectiveness of photodynamic therapy" financed by Foundation for Polish Science. I have learned molecular and cellular biology techniques, mammalian cell culture and *in vivo* mouse models. Importantly, I had an opportunity to supervise students, technicians, design whole projects and become more independent. I successfully applied for my own project funding. I am currently leading two research projects: "Improvement of photodynamic therapy by mobilization of dendritic cells", financed by Polish National Science Centre, and "Search for target proteins for the new compounds with antitumor activity", financed by Polish Ministry of Science and Higher Education within IUVENTUS program.

## B. Publications published during BASTION project

|          | <b>Authors, title, journal, year</b>   | <b>IF</b>    |
|----------|--|--------------|
| <b>1</b> | Siernicka M, Winiarska M, Bajor M, <b>Firczuk M</b> , Muchowicz A, Bobrowicz M, Fauriat C, Golab J, Olive D, Zagozdzon R. „Adenanthin, a new inhibitor of thiol-dependent antioxidant enzymes, impairs the effector functions of human natural killer cells.” <b>Immunology</b> . 2015 Jun 11. doi: 10.1111/imm.12494.   | <b>3,795</b> |
| <b>2</b> | Muchowicz A; <b>Firczuk M</b> ; Wachowska M; Kujawa M; Jankowska-Steifer E; Gabrysiak M; Pilch Z; Klossowski S; Ostaszewski R; Golab J. „SK053 triggers tumor cells apoptosis by oxidative stress-mediated endoplasmic reticulum stress”. <b>Biochem Pharmacol</b> . 2015 Feb 15;93(4):418-27. doi: 10.1016/j.bcp.2014.12.019.   | <b>5,009</b> |
| <b>3</b> | Nowis D, Malenda A, Furs K, Oleszczak B, Sadowski R, Chlebowska J, <b>Firczuk M</b> , Bujnicki JM, Staruch AD, Zagozdzon R, Glodkowska-Mrowka E, Szablewski L, Golab J. „Statins impair glucose uptake in human cells”, <b>BMJ Open Diabetes Research &amp; Care</b> , 2014, vol.2, doi:10.1136/bmjdr-2014-000017.   | <b>0</b>     |
| <b>4</b> | <b>Firczuk M</b> , Gabrysiak M, Gołab J. „GRP78-targeting subtilase cytotoxin sensitizes cancer cells to photodynamic therapy”, <b>book chapter 6</b> : “GRP78-targeting sensitizes cancer cells to cytotoxic effects of photodynamic therapy” in the book entitled: “ <b>Resistance to Photodynamic Therapy in Cancer</b> ”, Springer. 2014, DOI: 10.1007/978-3-319-12730-9.                                      | <b>0</b>     |
| <b>5</b> | Muchowicz A, <b>Firczuk M</b> , Chlebowska J, Nowis D, Stachura J, Barankiewicz J, Trzeciecka A, Klossowski S, Ostaszewski R, Zagozdzon R, Pu JX, Sun HD, Golab J. Adenanthin targets proteins involved in the regulation of disulphide bonds. <b>Biochem Pharmacol</b> . 2014 May 15;89(2):210-6. doi: 10.1016/j.bcp.2014.02.022.   | <b>5,009</b> |
| <b>6</b> | <b>Firczuk M</b> , Gabrysiak M, Barankiewicz J, Domagala A, Nowis D, Kujawa M, Jankowska-Steifer E, Wachowska M, Glodkowska-Mrowka E, Korsak B, Winiarska M, Golab J. GRP78-targeting subtilase cytotoxin sensitizes cancer cells to photodynamic therapy. <b>Cell Death Dis</b> . 2013 Jul 25;4:e741. doi: 10.1038/cddis.2013.265.  | <b>5,177</b> |
| <b>7</b> | Winiarska M, Nowis D, Bil J, Glodkowska-Mrowka E, Muchowicz A, Wanczyk M, Bojarczuk K, Dwojak M, <b>Firczuk M</b> , Wilczek E, Wachowska M, Roszczenko K, Miaczynska M, Chlebowska J, Basak GW, Golab J. 2012. Prenyltransferases Regulate CD20 Protein Levels and Influence Anti-CD20 Monoclonal Antibody-mediated Activation of Complement-dependent Cytotoxicity. <b>J Biol Chem</b> . Sep 14;287(38):31983-93. | <b>4,651</b> |



### C. Grant applications submitted during BASTION project

1. OPUS7, National Science Centre, “ The role for thiol-dependent antioxidant enzymes in estrogen receptor-positive breast cancer”, Key Investigator. 2014. Funding granted.
2. Diamond Grant, Ministry of Science and Higher Education, Project Leader: Antoni Domagała, “Role of autophagy in tumor cells response to photodynamic therapy”, 2014. Project supervisor. Funding granted.
3. OPUS 8, National Science Centre, „Studies of the role of peroxiredoxin 1 and other antioxidant enzymes in B cell acute lymphoblastic leukemia”, leader (PI). 2014. Not funded.
4. Horizon 2020, Call: H2020-TWINN-2015. European Commission. Proposal entitled: Strategies towards Excellence in Immuno-Oncology - „STREAM”. Role: work package deputy leader. June 2015. Under evaluation.

### D. Participation in grants during BASTION project

| Grant number         | Title   | Function           | Duration  | Funding Institution                                     |
|----------------------|---|--------------------|-----------|---|
| <b>N N401 037138</b> | Improvement of the efficacy of photodynamic therapy by the mobilization of dendritic cells                                  | Leader (PI)        | 2010-2014 | Ministry of Science and Higher Education                |
| <b>1M19/DG8</b>      | Investigation of the effects of EGF-SubA fusion protein on the efficacy of photodynamic therapy in vivo using mice models.  | Project supervisor | 2012-2014 | Diamond Grant, Ministry of Science and Higher Education |
| <b>IP1/2011/71</b>   | Search for target proteins for the new compounds with antitumor activity  | Leader (PI)        | 2012-2015 | Ministry of Science and Higher Education                |
| <b>1M19/DG8</b>      | Role of autophagy in tumor cells response to photodynamic therapy”  | Project supervisor | 2014-2016 | Diamond Grant, Ministry of Science and Higher Education |
| <b>1M19/PM13</b>     | Investigation of the role of peroxiredoxin 1 on the proliferation and survival of a human Burkitt’s lymphoma cell line Raji | Leader (PI)        | 2013-2014 | Medical University of Warsaw                            |
| <b>NZ5/01354</b>     | The role for thiol-dependent antioxidant enzymes in estrogen receptor-positive breast cancer                                | Key Investigator   | 2015-2018 | National Science Center                                 |

### E. Participation in the conferences during BASTION project

1. European Society for Photobiology 2013 Congress, Liege, Belgium, 2-6 September 2013.
2. 21<sup>st</sup> ECDO Euro conference on Apoptosis on “Cell death: a Biomedical paradigm”, Paris, France, 25-28 September, 2013.
3. 56<sup>th</sup> American Society of Hematology Annual Meeting, USA, San Francisco, 6-9 December 2014.
4. International Conference Translational Research in Oncology in New Member States Economies TRON, Warsaw, Poland, 21-22 May 2015.
5. 15th International Conference on Oxidative Stress Reduction, Redox Homeostasis & Antioxidants, France, Paris, Pasteur Institute, 22-24 June 2015.



#### **F. Oral presentation at the conferences**

1. European Society for Photobiology 2013 Congress, Liege, Belgium, 2-6 September 2013, lecture, title: "GRP78-targeting subtilase cytotoxin sensitizes cancer cells to photodynamic therapy".
2. 15th International Conference on Oxidative Stress Reduction, Redox Homeostasis & Antioxidants, France, Paris, Pasteur Institute, 22-24 June 2015. Oral presentation, title: "Thiol-reactive peptidomimetic SK053 targets dimeric peroxiredoxins in human lymphoma cell lines".

#### **G. Poster presentation at the conferences**

1. 21<sup>st</sup> ECDO Euro conference on Apoptosis on "Cell death: a Biomedical paradigm", 25-28 September, 2013, title "Photodynamic therapy combined with GRP78-targeting subtilase cytotoxin trigger atypical cell death in apoptosis-deficient prostate cancer cells".
2. 56<sup>th</sup> American Society of Hematology Annual Meeting, poster title: "Peroxiredoxins-1 and 2 Affect Proliferation and Survival of Lymphoma Cells" –presenting author, USA, San Francisco, 6-9 December 2014
3. International Conference Translational Research in Oncology in New Member States Economies TRON, Poster presentation, poster title: "Peroxiredoxins-1 and 2 Affect Proliferation and Survival of Lymphoma Cells", Warsaw, Poland, 21-22 May 2015.

#### **H. Participation in courses/trainings/workshops**

1. Research Team Management Workshop organized by Foundation for Polish Science within SKILLS program, 10-11 May 2013.
2. Scientific Writing Workshop organized by Foundation for Polish Science within SKILLS program, 10-12 June 2013.
3. "Cancer genetics for medical community" - workshop organized by the Medical University of Warsaw in the project BASTION, Warsaw, Poland, 17 June 2013.
4. "Application of flow cytometry in molecular oncology", workshop organized by BASTION, Medical University of Warsaw, 15 – 16 October 2014.
5. Workshop: "Genome-wide methods in cancer genetics", organized by BASTION, Medical University of Warsaw, 28 October 2014.
6. Coaching, training program organized by Foundation for Polish Science, within SKILLS project, October 2014-March 2015.
7. Workshop: "Molecular diagnostic in cancer" organized by BASTION, Medical University of Warsaw, 8 June 2015.

#### **I. Awards/fellowships obtained during BASTION project**

1<sup>st</sup> degree scientific reward from the Rector of the Medical University of Warsaw, for the participation in a publication series, diploma (2013).

#### **J. Students supervision**

Supervising three students, participants of student's scientific group at the Department of Immunology: Joanna Barankiewicz, Antoni Domagała and Anna Trzeciecka.

#### **K. Collaboration with other research teams started during BASTION project**

1. Prof. Eugene Jansen, National Institute for Public Health and Environment. Utrecht, Netherlands.





2. Prof. Wojciech Młynarski, dr Agata Pastorczak, Department of Pediatrics, Oncology, Hematology and Diabetology Medical University of Łódź, Poland.
3. Prof. Przemysław Juszczynski, mgr Anna Polak, Laboratory of Experimental Hematology, Institute of Hematology and Transfusion Medicine, Warsaw, Poland.
4. Prof. Ewa Lech-Marańda, lek med Elżbieta Patkowska, Department of Hematology, Institute of Hematology and Transfusion Medicine, Warsaw, Poland.
5. Prof. Monika Prochorec-Sobieszek, Department of Diagnostic Hematology, Institute of Hematology and Transfusion Medicine, Warsaw, Poland.
6. Prof. Matthias Bochtler, Laboratory of Structural Biology, International Institute of Molecular and Cell Biology, Warsaw, Poland.
7. Prof. Janusz Bujnicki, Dr Anna Czerwoniec, Laboratory of Bioinformatics and Protein Engineering, International Institute of Molecular and Cell Biology, Warsaw, Poland.
8. Prof. Michał Dadlez, Dr Agata Malinowska, Mass Spectrometry Laboratory, Institute of Biochemistry and Biophysics, Warsaw, Poland.

#### **L. International research visits during BASTION project**

University of Verona, Department of Neurology and Movement Sciences, 27 July 2013-10 August 2013

#### **M. Current research interests**

Recently, Malgorzata Firczuk has been trying to apply her knowledge of protein structure and structure-to-function relationships to the field of experimental oncology. She is particularly interested in disease-related proteins, which are involved in protein folding, redox homeostasis, and support tumour cell proliferation.

She is focused on investigating mechanisms of action of small molecule compounds, drug target selection and validation. Recently, she had identified peroxiredoxins as molecular targets for an electrophilic peptidomimetic compound, SK053, initially designed as thioredoxin / thioredoxin reductase system inhibitor. Using biotin-avidin affinity approach, she found two-cysteine dimeric peroxiredoxins as covalently bound to the biotin-labelled compound. She is now investigating the detailed mechanism of SK053 binding to peroxiredoxin-1, both in cells and with purified recombinant proteins. In addition, she is validating peroxiredoxins as potential targets in B lymphocyte-derived malignancies.

The second line of her scientific interest is focused on the role of endoplasmic reticulum resident chaperone, glucose regulated protein 78 (Grp78), in tumour cell survival and response to anti-tumour therapies. Grp78 is highly expressed in tumour cells and plays a cytoprotective role, supporting tumour growth. She had recently shown that Grp78 is up-regulated in response to photodynamic therapy, and contributes to the therapy resistance.

#### **N. Envisioned carrier paths after BASTION project**

Collaboration of Dr. Dominika Nowis with Dr. Malgorzata Firczuk ends with the finalization of the BASTION project as Dr. Firczuk joins the research team of Dr. Zagodzón. She will be employed for 18 months (half-time contract) within project funded by National Science Center: "The role for thiol-dependent antioxidant enzymes in estrogen receptor-positive breast cancer", NZ5/01354. Currently, Dr Firczuk is developing her expertise in the field of redox biology using in vitro and in vivo models. The results of Dr. Firczuk's projects should allow development of her research carrier in the field of tumor biology and therapy and establishment of her own research team. In the following months Dr. Firczuk will focus on gathering the preliminary results to support her future grant applications to the National Science Center in Poland.



## II. Anna Wojcicka (TEAM of Krystian Jazdzewski)



| DATE (YEARS) | DEGREE/EXPERIENCE | PLACE   | SUPERVISOR                             |
|--------------|-------------------|---|--|
| 1999-2006    | M.Sc.             | University of Warsaw, Faculty of Biology  | Agnieszka Dzikowska, PhD               |
| 2007-2012    | PhD               | Centre of Postgraduate Medical Education, Warsaw, Poland  | Prof. Alicja Nauman. Ph.D.             |
| 2012-present | Postdoc           | Genomic Medicine, Department of General, Transplant and Liver Surgery, Medical University of Warsaw | Prof. Krystian Jazdzewski, M.D., Ph.D. |

### A. Biosketch (provided by Anna Wojcicka)

I graduated from the Faculty of Biology at the Warsaw University. My Master's thesis, concerning the arginine catabolism in fungus *Aspergillus nidulans* was performed in the Department of Genetics. After graduation I was employed at the Department of Genetics, Institute of Biochemistry and Biophysics, Polish Academy of Sciences, where I investigated mechanisms of sister chromatid cohesion in the fungal model of *Saccharomyces cerevisiae*. In October, 2007 I commenced PhD studies at the Medical Centre of Postgraduate Education in Warsaw, Laboratory of Molecular Biology, under supervision of Professor Alicja Nauman whose scientific interest has been focused on the elucidation of the role of thyroid hormones in carcinogenesis. The research I conducted within the topic of my PhD thesis consisted of the analysis of the thyroid hormone receptor beta (*THRB*) gene methylation and miRNA-dependent regulation in clear cell renal cell carcinoma, as well as of the evaluation of the effect of thyroid hormones on expression of genes coding for DNA methyltransferases.

I also gained additional experience working in international laboratories and cooperating with other laboratories in Poland. During my Master studies I spent six months at the Department of Clinical Genetics, Vrije Universiteit in Amsterdam, investigating mutations that could be potentially involved in pathogenesis of multiple sclerosis. Furthermore, since May until September 2011 I was an occasional student at Imperial College London, Laboratory of Molecular Endocrinology, where I performed part of my PhD thesis, analyzing expression of DNA methyltransferases in tissues obtained from wild-type and mutant mice with disrupted T3 signalling. I also participated in other projects conducted in the Laboratory, analyzing in vivo phenotype of the developing and adult skeleton in murine and avian models. From October 2011 until February 2012 I worked at the Ohio State University Comprehensive Cancer Center. My research was focused on the analysis of the risk factors for thyroid cancer and included conduction of the genetic association study using the Sequenom technology and the analysis of microRNAs that are aberrantly expressed in thyroid cancer.

### B. Publications published during BASTION project

|   | Authors, title, journal, year  | IF    |
|---|--|-------|
| 1 | Lakshmanan A, Wojcicka A, Kotlarek M, Zhang X, Jazdzewski K, Jhiang SM. 2015 MicroRNA-339-5p modulates Na <sup>+</sup> /I <sup>-</sup> symporter-mediated radioiodide uptake. <i>Endocr Relat Cancer</i> | 4,805 |



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|-----------|---|--------------|
|           | Feb;22(1):11-21   |              |
| <b>2</b>  | Wojcicka A, Swierniak M, Kornasiewicz O, Gierlikowski W, Maciąg M, Kolanowska M, Kotlarek M, Gornicka B, Koperski L, Niewinski G, Krawczyk M, Jazdzewski K. 2014 Next generation sequencing reveals microRNA isoforms in liver cirrhosis and hepatocellular carcinoma. <i>Int J Biochem Cell Biol.</i> 53:208-17  | <b>4,046</b> |
| <b>3</b>  | Wojcicka A, Piekuelko-Witkowska A, Kedzierska H, Rybicka B, Poplawski P, Boguslawska J, Master A, Nauman A. 2014 Epigenetic regulation of thyroid hormone receptor Beta in renal cancer. <i>Plos One.</i> 2014 May 21;9(5):e97624   | <b>3,234</b> |
| <b>4</b>  | Wójcicka A, Czetwertyńska M, Świerniak M, Długosińska J, Maciąg M, Czajka A, Dymecka K, Kubiak A, Kot A, Płoski R, de la Chapelle A, Jazdzewski K 2014 Variants in the ATM-CHEK2-BRCA1 axis determine genetic predisposition and clinical presentation of papillary thyroid carcinoma. <i>Genes Chromosomes Cancer,</i> 53(6): 516-23   | <b>4,041</b> |
| <b>5</b>  | Wojcicka A, de la Chapelle A, Jazdzewski K. 2014 MicroRNA-related sequence variations in human cancers. <i>Human Genetics,</i> 133(4):463-9   | <b>4,824</b> |
| <b>6</b>  | Boguslawska J, Piekuelko-Witkowska A, Wojcicka A, Kędzierska H, Popławski P, Nauman A 2014 Regulatory feedback loop between T3 and microRNAs in renal cancer. <i>Mol Cell Endocrinol.</i> 25;384(1-2):61-70   | <b>4,405</b> |
| <b>7</b>  | Swierniak M, Wojcicka A, Czetwertynska M, Stachlewska E, Maciąg M, Wiechno W, Gornicka G, Bogdanska M, Koperski L, de la Chapelle A. Jazdzewski K. 2013 In-depth characterization of the microRNA transcriptome in normal thyroid and papillary thyroid carcinoma. <i>J Clin Endocrinol Metab,</i> 2013 Aug;98(8):E1401-9.  | <b>6,310</b> |
| <b>8</b>  | Liyanarachchi S, Wojcicka A, Li W, Czetwertynska M, Stachlewska E, Nagy R, Hoag K, Wen B, Ploski R, Ringel MD, Kozłowicz-Gudzinska I, Gierlikowski W, Jazdzewski K, He H and Albert de la Chapelle 2013 Cumulative Risk Impact of Five Genetic Variants Associated With Papillary Thyroid Carcinoma <i>Thyroid,</i> 23(12):1532-40  | <b>3,843</b> |
| <b>9</b>  | Wojcicka A, Bassett JH; Williams GR 2013 Mechanisms of action of thyroid hormones in the skeleton. <i>BBA - General Subjects, Special Issue: Thyroid hormone signalling.</i> <i>Biochim Biophys Acta Jul;</i> 1830(7):3979-86   | <b>3,829</b> |
| <b>10</b> | Piekuelko-Witkowska A, Kedzierska H, Poplawski P, Wojcicka A, Rybicka B, Maksymowicz M, Grajkowska W, Matyja E, Mandat T, Bonicki W, Nauman P. 2013 Alternative splicing of iodothyronine deiodinases in pituitary adenomas. Regulation by oncoprotein SF2/ASF <i>Biochim Biophys Acta Jun;</i> 1832(6):763-72  | <b>5,089</b> |
| <b>11</b> | He H, Li W, Wu D, Nagy R, Liyanarachchi S, Akagi K, Jendrzewski J, Jiao H, Hoag K, Wen B, Srinivas M, Waidyaratne G, Wang R, Wojcicka A, Stachlewska E, Czetwertynska M, Dlugosinska J, Gierlikowski W, Ploski R, Krawczyk M, Jazdzewski K, Kere J, Symer DE, Jin V, Wang Q, de la Chapelle A. 2013 Ultra-rare mutation in long-range enhancer predisposes to thyroid carcinoma with high penetrance, <i>PLoS One.</i> 2013; 8(5): e61920                     | <b>3,534</b> |
| <b>12</b> | He H, Bronisz A, Liyanarachchi S, Nagy R, Li W, Huang Y, Akagi K, Saji M, Kula D, Wojcicka A, Nihil S, Wen B, Puch Z, Kalemba M, Stachlewska E, Czetwertynska M, Dlugosinska J, Dymecka K, Ploski R, Krawczyk M, Morrison PJ, Ringel MD, Kloos RT, Jazdzewski K, Symer DE, Vieland VJ, Ostrowski M, Jarząb B, de la Chapelle A. 2013 SRGAP1 is a candidate gene for papillary thyroid carcinoma susceptibility. <i>J Clin Endocrinol Metab,</i> 98(5):973-980 | <b>6,310</b> |

### C. Patent applications submitted during BASTION project

1. World Intellectual Property Organization "Use Of A Micro-RNA Marker For Thyroid Tumor Diagnosis And A Diagnostic Kit Comprising Such Markers" PCT/IB2014/066057 (2014)
2. World Intellectual Property Organization "Use Of Micro-RNA Markers For Diagnosis Of Liver PCT/IB2014/065342 (2014)
3. Polish Patent Bureau "A new method for diagnosis of hepatocellular carcinoma, the use of microRNA markers in diagnostics of liver pathologies, in prediction of their progression and response to treatment,



together with a diagnostic panel based on the above markers” P30817PL00 (2013)

4. Polish Patent Bureau “A new method for diagnosis of thyroid carcinoma, the use of microRNA markers in diagnostics of thyroid pathologies, in prediction of their progression and response to treatment, together with a diagnostic panel based on the above markers”. P30859PL00 (2013)

#### **D. Participation in grants during BASTION project**

1. National Science Centre Grant Sonata: MicroRNA-dependent regulation of iodide transporters: NIS, AIT and Pendrin and aberrations of this process in papillary thyroid carcinoma 2012/07/D/NZ3/04149 (2012-2015) – Principal Investigator
2. National Centre for Research and Development Lider Grant: The use of next-generation sequencing for elucidation of a sensitive and specific molecular panel for diagnostics of thyroid cancers (2014-2017) – Principal Investigator
3. Ministry of Science and Higher Education Iuventus Plus Grant: Evaluation of the possibility of using microRNA inhibitors as adjuvant therapy for thyroid cancer (2015-2017) – Principal Investigator
4. Foundation For Polish Science Impuls Programme: Implementation of a molecular prognostic panel for thyroid cancer (2015) – Principal Investigator
5. Foundation For Polish Science TEAM Programme: In search of new pathways of tumorigenesis - genome-wide functional analysis of microRNAs deregulated in human cancers, financed by the European Union within the European Regional Development Fund (2014) – Investigator
6. Foundation for Polish Science FOCUS Programme: Role of microRNAs in thyroid carcinogenesis (2011-2012) – Investigator

#### **E. Participation in the conferences during BASTION project**

1. European human genetics conference (ESHG), Glasgow, UK, 6-9 June 2015
2. 16th International Congress of Endocrinology/The Endocrine Society's 96th Annual Meeting (ICE/ENDO 2014), Chicago, USA, 20-25 June 2014
3. European Congress of Endocrinology, Wroclaw, Poland, 3-7 May 2014

#### **F. Oral presentation at the conferences**

1. M Kolanowska, A Wojcicka, A Kubiak, M Swierniak, M Maciag, W Wiechno and K Jazdzewski Next-Generation Sequencing Reveals a Novel, Thyroglobulin-Embedded microRNA Gene Deregulated in Papillary Thyroid Carcinoma ICE/ENDO, 20-25 June 2014, Chicago, USA

#### **G. Poster presentation at the conferences**

1. Wojcicka, A. Kubiak, M. Kotlarek, A. Czajka, M. Czetwertynska, J. Dlugosinska, M. Swierniak, N. Fedoryszak-Kuska, B. Gornicka, K. Jazdzewski A polymorphism in miR-146a tailors genetic predisposition to differentiated thyroid cancer, modulates its clinical outcome and alters proliferation of tumor cells. European human genetics conference, Glasgow, UK, 6-9 June 2015
2. M. Swierniak, A. Wojcicka, M. Czetwertynska, J. Dlugosinska, E. Stachlewska, W. Gierlikowski, B. Gornicka, L. Koperski, M. Bogdanska, W. Wiechno, K. Jazdzewski Association between GWAS-derived rs966423 genetic variant and overall mortality in patients with differentiated thyroid cancer European human genetics conference, Glasgow, UK, 6-9 June 2015
3. Wójcicka A, Czetwertyńska M, Świerniak M, Długosińska J, Maciąg M, Czajka A, Dymecka K, Płoski R, de la Chapelle A, Jazdzewski K. Variants in the ATM-CHEK2-BRCA1 Axis Determine Genetic Predisposition and



Clinical Presentation of Papillary Thyroid Carcinoma; ICE/ENDO, Chicago, USA, Endocrine Society Presidential Award for the best poster in thyroidology, 20-25 June 2014.

4. Wójcicka A, Gierlikowski W, Kotlarek M, Bakuła-Zalewska E, Jażdżewski K. Apical iodide transporter (AIT) and its microRNA – induced silencing in thyroid malignancies ICE/ENDO, Chicago, USA, 20-25 June 2014.
5. The role of ATM-CHEK2-BRCA1 axis in determination of genetic predisposition and clinical presentation of papillary thyroid carcinoma. Anna Wójcicka, Małgorzata Czetwertyńska, Michał Świerniak, Joanna Długosińska, Monika Maciąg, Agnieszka Czajka, Kinga Dymecka, Adam Kot, Rafał Płoski, and Krystian Jażdżewski, European Congress of Endocrinology, Wrocław, Poland, 3-7 May 2014
6. The effect of allelic variants of the thyroid hormone receptor beta (THRB) gene on the incidence of papillary thyroid carcinoma. Anna Wójcicka, Marek Rosłon, Małgorzata Czetwertyńska, Michał Świerniak, Joanna Długosińska, Adam Kot, Rafał Płoski, Aneta Hromada-Judycka, Marta Świech and Krystian Jażdżewski, European Congress of Endocrinology, Wrocław, Poland, 3-7 May 2014

#### **H. Participation in courses/trainings/workshops**

1. 55<sup>th</sup> Annual Short Course of Medical and Experimental Mammalian Genetics, Jackson Laboratory, Bar Harbor, USA, 20 July-03 August 2014.

#### **I. Organization of the conferences**

1. Cancer Genetics for Medical Community, 17 June 2013, Warsaw

#### **J. Awards/fellowships obtained during BASTION project**

1. Minister of Science and Higher Education - Scholarship for outstanding young scientists (2014)
2. March of Dimes Foundation Scholarship towards participation in the 55<sup>th</sup> Annual Short Course of Medical and Experimental Mammalian Genetics, Jackson Laboratory, Bar Harbor, USA (2014)
3. Endocrine Society Presidential Poster Award for the best poster presentation in thyroidology (2014)
4. Fellowship within the Foundation for Polish Science Mentoring Programme (2013)
5. 1st degree Prize awarded by the Director of the Medical Centre of Postgraduate Education for a chapter in the "Clinical Endocrinology" textbook (2013)

#### **K. Students supervision**

Supervising two PhD students: Wojciech Gierlikowski and Marta Kotlarek

#### **L. Collaboration with other research teams started during BASTION project**

1. Group of Prof. Albert de la Chapelle, Department of Molecular Virology, Immunology and Medical Genetics, Ohio State University
2. Prof Sissy M. Jhiang, Ohio State University

#### **M. International research visits during BASTION project**

University of Ferrara, lab of Stefano Volinia, 24 June-17 July 2015 and 22 July-07 August 2015



#### **N. Current research interests**

Anna Wojcicka is a molecular biologist working in the field of molecular endocrinology and oncology. For the past several years her research has been focused on the molecular basis of thyroid hormones action: their involvement in the processes of cell division and proliferation as well as on the role of aberrances in thyroid hormone signaling in development and progression of human cancers.

The projects she is currently involved in aim at elucidation of the role of microRNAs in the pathology of human diseases. Increased expression of miRs, observed in cancers, leads to their enhanced binding with target mRNAs, causing severe downregulation of synthesis of proteins and resulting in deregulation of numerous cellular pathways. In her ongoing projects she employs next-generation sequencing to identify comprehensive miRNA profiles of human cancers, including papillary thyroid carcinoma and hepatocellular carcinoma. She seeks to identify novel, previously unknown microRNAs and their isoforms, and to elucidate their impact on the cellular transcriptome together with a potential linkage with pathogenesis of cancer. She is also attempting to propose specific, microRNA-based diagnostic panels for non-invasive diagnostics of thyroid and liver malignancies.

#### **O. Envisioned career paths in BASTION project**

Anna's scientific plans are focused on further studies on the pathology of thyroid cancer. Her long-term goals include elaboration of specific diagnostic panels allowing for non-invasive diagnostics of thyroid cancers. Moreover, she is currently initiating collaboration on the project aiming at elucidation of therapeutic tools for thyroid cancer, based on reestablishment of expression of genes coding for iodide transporters. Within the project, she will supervise the projects and theses of two PhD students. Anna is currently a PI in 4 on-going grant projects and will be employed at the Medical University of Warsaw and at the Centre of New Technologies, University of Warsaw.



### III. Malgorzata Czystowska-Kuzmicz (TEAM of Jakub Golab)



| DATE (YEARS)     | DEGREE/ EXPERIENCE | PLACE   | SUPERVISOR              |
|------------------|--------------------|---|-------------------------|
| 1998             | M.Sc.              | Heinrich-Heine University, Duesseldorf, Germany                     | Prof. Frank Wunderlich  |
| 2006             | PhD                | Heinrich-Heine University, Duesseldorf, Germany                     | Prof. Peter Dall        |
| 2006-2009        | Postdoc            | University of Pittsburgh Cancer Institute (UPMC), Pennsylvania, USA | Prof. Theresa Whiteside |
| 2009-2013        | Postdoc            | Maternity leave   |                         |
| since March 2013 | Postdoc            | BASTION   | Prof. Jakub Golab       |

#### A. Biosketch (provided by Malgorzata Czystowska-Kuzmicz)

I earned my MSc degree at Heinrich-Heine University in Dusseldorf, Germany in 1998. During my Masters studies I also worked as an undergraduate research assistant at the Institute of Transplant-Diagnostic and Cell Therapeutics of the Heinrich-Heine University, participating in HLA-class I and II typification of patients and donors for bone-marrow and stem-cell transplantations. For graduate studies I joined Dr. Dieter Niederacher's Laboratory of Molecular Genetics, part of the Department of Obstetrics and Gynecology of Heinrich-Heine University. My Ph.D. thesis was a part of research projects of the German Cancer Aid Study and the German Human Genome Project. Basing on bioinformatic analysis of EST-databases and microarray data we tried to identify novel genetic markers in gynecological tumors. For this purpose I designed and developed a specific strategy to validate these candidate genes, which included expression analysis, high throughput pre-screening for mutations and promoter methylation (LOH-analysis, DHPLC) and functional cell-based assays. I identified the insulin-like growth factor binding protein 4 (IGFBP-4) as a putative tumor suppressor in ovarian cancer. I showed that IGFBP-4 downregulation in ovarian tumors was due to allelic loss and promoter-hypermethylation and was ER-status dependent. IGFBP-4 showed IGF-I-dependent anti-proliferative and partly IGF-I-independent pro-apoptotic effects in OvCa cell lines. After completing my Ph.D. in 2006, I moved to Prof. Theresa Whiteside's laboratory, at University of Pittsburgh Cancer Institute (UPMC), Pennsylvania, USA. Her laboratory had been doing pioneering work in characterizing tumor-mediated escape mechanisms and identifying surrogate immunologic markers of prognosis and response to therapy. During this time I was investigating mechanisms responsible for tumor-induced suppression of immune effector cells – primarily in head and neck cancer (HNC), but also in ovarian cancer and acute myelogenous leukemia. My special attention was devoted to the role of tumor-derived microvesicles as mediators of immune suppression and disease progression. In this context I also investigated the exosome-driven Treg (regulatory T cells)- mediated death of effector cells in HNC patients. My second focus was the chemokine receptor signalling that regulates host response to tumors. I found that patients with cancer have an increased frequency of circulating apoptosis-sensitive CD8+ cells, which do not express the chemokine receptor CCR7, and few CD8(+)/CCR7(+) apoptosis-resistant T cells. Moreover, I showed that the CD8(+)/CCR7(+) T-cell frequency in HNSCC patients' blood tested at diagnosis can discriminate them from normal controls and predicts disease recurrence. Finally, I was involved in the development of cytokine therapies for cancer patients in cooperation with an industry partner, investigating



the molecular mechanisms of T-cell protection of a new developed cytokine-based immunotherapeutic. I also participated in the evaluation of a randomized phase II p53 vaccine trial in ovarian cancer patients.

#### B. Publications published during BASTION project

|          | <b>Authors, title, journal, year</b>   | <b>IF</b>    |
|----------|--|--------------|
| <b>1</b> | Czystowska M, Gooding W, Szczepanski MJ, Lopez-Abaitero A, Ferris RL, Johnson JT, Whiteside TL. The immune signature of CD8(+)/CCR7(+) T cells in the peripheral circulation associates with disease recurrence in patients with HNSCC. Clin Cancer Res. 2013 Feb 15;19(4):889-99. | <b>8,193</b> |

#### C. Grant applications submitted during BASTION project

1. OPUS6; Title: 'Elucidation of the role of tumor-derived and exosomal arginases in avoiding immune responses by ovarian cancer' – Principal investigator, Accepted for funding.

#### D. Participation in the conferences during BASTION project

1. Annual Meeting of the International Society of Extracellular Vesicles (ISEV), Washington DC, United States, 22-26 April 2015
2. BASTION conference "Translational Research in Oncology in New Member State Economies" 21-22 May 2015

#### E. Poster presentation at the conferences

1. Annual Meeting of the International Society of Extracellular Vesicles (ISEV), poster title: "The adenosine pathway in ovarian carcinoma: tumor cells and tumor-derived exosomes express CD39 and CD73 ectonucleotidases, produce adenosine and mediate immune suppression", Washington DC, United States, 22-26 April 2015.
2. BASTION conference "Translational Research in Oncology in New Member State Economies", poster title: "The adenosine pathway in ovarian carcinoma: tumor cells and tumor-derived exosomes express CD39 and CD73 ectonucleotidases, produce adenosine and mediate immune suppression", Warsaw, 21-22 May 2015.

#### F. Participation in courses/trainings/workshops

1. FNP "Project management" workshop (3-4 November 2014)
2. Training "Isolation and molecular characterisation of cancer-derived exosomes", University of Pittsburgh Cancer Institute in Pittsburgh, PA (United States), 13-19 July 2014
3. Nordic Immunohistochemistry Basic Course organized in DAKO's Glostrup/Kopenhagen facility, Denmark, 21-22 January 2015,
4. ISEV – NIH ERCC collaborative Education Day "RNA Diversity in Extracellular Vesicles", Washington DC, USA 22 April 2015
5. BASTION Molecular Diagnostics in Cancer Workshop, Warsaw 08 June 2015

#### G. Students supervision

Supervising student Anna Czekalska (participants of student's scientific group at the Department of Immunology), master student Karolina Soroczynska, title of master thesis "Role of exosome-derived arginase in tumor escape of ovarian cancer", end of master thesis: September 2015

#### H. Collaboration with other research teams started during BASTION project





1. Dr. Marta Szajnik-Szczepański from Poznań University of Medical Sciences, Department of Gynecologic Oncology,
2. Dr. Jacek Sieńko from Second Clinic of Obstetrics and Gynecology, Medical University of Warsaw

#### **I. Current research interests**

Recent research activities of Małgorzata focus on the understanding of the defensive strategies developed by tumors to protect against immune attack. This phenomenon is referred to as “tumor escape” and has been recently accepted as a major problem responsible for the tumor resistance to immune therapies and for the general lack of success in generation of clinical responses to vaccines in patients diagnosed with cancer. She and others have identified tumor-derived exosomes (TDE) as carriers for the delivery of defined signals from tumor site to distant organs, enabling the tumor to develop a systemic immune suppression. Recently, she identified on ovarian cancer exosomes two enzymes, i.e. arginase-1 and -2, that are involved in degradation of non-essential amino-acids and play a critical role in chronic inflammation and evasion of anti-tumor immunity. Thus, she hypothesizes that through the release of arginase-expressing exosomes which become systemically distributed through the bloodstream, tumor cells achieve a global L-Arg depletion leading to a systemic T-cell dysfunction. She plans to delineate the immunosuppressive role of these tumor-derived enzymes of the amino-acids metabolism. Taking also into account the recent development of inhibitors of amino acid metabolism, we also assume that the inhibition of the expression and enzymatic activity of the above-mentioned enzymes may tilt the balance from an immune-suppressive to an immune-active environment and should have a measurable impact on the disease outcome. Therefore, blocking arginase could be a target for novel anti-cancer strategies, especially in combination with existing molecularly targeted therapies, but also classical chemotherapy.

#### **J. Envisioned career paths in BASTION project**

Małgorzata Czystowska-Kuzmicz will be funded from the National Science Center grant (NCN; “Elucidation of the role of tumor-derived and exosomal arginases in avoiding immune responses by ovarian cancer”) until July 2017. She will focus on elucidating the role of tumor-derived exosomes, which contain immune response-suppressing enzymes. The results of this project should allow development of research carrier of Dr. Małgorzata Czystowska in the field of tumor immunobiology. We also expect that the results of the research will allow to prepare a habilitation (Ds.C.) thesis of Dr. Małgorzata Czystowska in 2018. Further employment is warranted pending successful acquisition of grants from NCN or other sources.



#### IV. Beata Pyrzynska (TEAM of Magdalena Winiarska)



| DATE (YEARS) | DEGREE/ Experience | PLACE   | SUPERVISOR                  |
|--------------|--------------------|---|-----------------------------|
| 1989-1994    | M.Sc.              | Faculty of Biology, Warsaw University, Poland                                       | Prof. A. K. Tarkowski       |
| 1995-1997    | Assistant          | Nencki Institute of Experimental Biology, Polish Academy of Science, Warsaw, Poland | Prof. A. Sobota             |
| 1998-2001    | PhD                | Nencki Institute of Experimental Biology, Polish Academy of Science, Warsaw, Poland | Prof. B. Kaminska-Kaczmarek |
| 2002-2006    | Postdoc            | Emory University, Atlanta, GA, USA  | Prof. E. G. van Meir        |
| 2007-2013    | Postdoc            | International Institute of Molecular and Cell Biology, Warsaw, Poland               | Prof. M. Miaczynska         |

##### A. Biosketch (provided by Beata Pyrzynska)

I was first time involved in the laboratory work as a M.Sc. student at the Department of Embryology, Warsaw University, studying development of the block against polyspermy in different stages of oocyte maturation. Later, as an assistant at the Nencki Institute of Experimental Biology in Warsaw I studied involvement of the cytoskeletal and signaling proteins in phagocytosis. During that time I also gained my first international research experience as a TEMPUS fellow at the Institute of General Pathology, Perugia, Italy. My interest in cancer biology started when I joined the Laboratory of Transcriptional Regulation at the Nencki Institute of Experimental Biology, where I conducted research on the molecular mechanisms leading to glioma cell death upon treatment with immunosuppressive drug cyclosporine A. I used the short-term fellowship from EMBO as an opportunity to collaborate with the National Center of Biotechnology (CNB) in Madrid. My Ph.D. thesis dissertation (2001) was awarded at the Nencki Institute of Experimental Biology and I was also recognized as a young outstanding scientist by the Foundation for Polish Science (FNP, START program).

In 2002 I was recruited to the Winship Cancer Institute, Emory University, Atlanta, USA, to work as a postdoctoral fellow in the Laboratory of Molecular Neuro-Oncology. I used the microarray approach to study the influence of tumor suppressors (such as p53 or p14ARF) status on the development and progression of glioblastoma. My work was awarded by the research fellowships from NATO and from the American Brain Tumor Association. In 2006 I returned to Poland to work at the International Institute of Molecular and Cell Biology in Warsaw. I brought the experience in cancer biology to study the signal transduction pathways originated at the endocytic compartments and influencing different aspects of tumor growth. Over the years I have conducted the research in cancer biology and cancer therapeutics fields. I studied the regulation of cellular signaling leading to changes in gene expression and tumorigenesis. Beside the basic methods of molecular and cell biology I had the opportunity to gain some experience in bioinformatics, microarray technique and proteomics. Recently, I was recruited under the BASTION program to work as an experienced scientist at the Department of Immunology, Medical University of Warsaw.



## B. Publications published during BASTION project

|          | <b>Authors, title, journal, year</b>   | <b>IF</b>     |
|----------|--|---------------|
| <b>1</b> | Zerrouqi A., Pyrzynska B., Brat D.J., Van Meir E.G. (2014). P14ARF suppresses tumor-induced thrombosis by regulating the tissue factor pathway <i>Cancer Res.</i> 74, 1371-8.1.  | <b>9,329</b>  |
| <b>2</b> | Pyrzynska B, Banach-Orlowska M, Teperek-Tkacz M, Miekus K, Drabik G, Majka M, Miaczynska M. Multifunctional protein APPL2 contributes to survival of human glioma cells. <i>Mol Oncol.</i> 2013 Feb;7(1):67-84.  | <b>5,935</b>  |
| <b>3</b> | Bojarczuk K., Siernicka M., Dwojak M., Bobrowicz M., Pyrzynska B., Gaj P., Karp M., Giannopoulos K., Efremov D.G., Golab J., Winiarska M. (2014). B-cell receptor pathway inhibitors affect CD20 levels and impair antitumor activity of anti-CD20 monoclonal antibodies. <i>Leukemia</i> 28, 1163-7.  | <b>10,431</b> |
| <b>4</b> | Winiarska M., Bojarczuk K., Pyrzynska B., Bil J., Siernicka M., Dwojak M., Bobrowicz M., Miazek N., Zapala P., Zagodzón A., Krol M., Syta A., Podszywalow-Bartnicka P., Pilch Z., Dabrowska-Iwanicka A., Juszczynski P., Efremov D.G., Slabicki M., Zenz T., Le Roy A., Olive D., Rygiel T.P., Leusen J., Golab J. (2014). Inhibitors of SRC kinases impair antitumor activity of anti-CD20 monoclonal antibodies. <i>mAbs</i> 6, 1300-13. | <b>4,558</b>  |
| <b>5</b> | Dwojak M., Bobrowicz M., Bil J., Bojarczuk K., Pyrzynska B., Siernicka M., Malenda A., Lech-Maranda E., Tomczak W., Giannopoulos K., Golab J., Winiarska M. (2015) Sorafenib improves rituximab and ofatumumab efficacy by decreasing the expression of complement regulatory proteins. <i>Blood Cancer J.</i> 5(E300):1-4.  | <b>3,467</b>  |

## C. Grant applications submitted during BASTION project

1. Grant application as principal investigator "Influence of AKT signaling pathway on CD20 expression and antitumor activity of therapeutic monoclonal antibodies." National Science Center (NCN, grant OPUS); December 2013;
2. Participation in grant application as supervisor of the student - Nina Miązek; Diamond Grant from the Ministry of Science and Higher Education "Impact of selected chemotherapeutic agents on the efficacy of anti-CD20 immunotherapy in B-cell lymphoma"; February 2015;
3. Participation in Twinning grant application, call: H2020-TWINN-2015, acronym: STREM; Principal Investigator – Prof. Jakub Golab; May 2015.

## D. Participation in grants during BASTION project

1. Principal Investigator in grant OPUS "Influence of AKT signaling pathway on CD20 expression and antitumor activity of therapeutic monoclonal antibodies." National Science Center (NCN; grant no: 2013/11/B/NZ5/03240; 1074759 PLN).

## E. Participation in the conferences during BASTION project

1. "Research-driven, multidisciplinary oncological care in Poland: sharing experiences to foster collaborations between MD Anderson Cancer Center Sister Institutions"; Warsaw, Poland, 23-24 June 2014.
2. "56th ASH Annual Meeting and Exposition", San Francisco, USA, 6-9 December 2014.
3. "Heart of Europe Zebrafish Meeting"; Warsaw, Poland, 17-19 September, 2014.
4. "Open Research Data: Implications for Science and Society"; Warsaw, Poland, 28-29 May 2015.

## F. Oral presentation at the conferences



1. Oral presentation of data during Second International Advisory Board (IAB) Meeting of BASTION project "Regulation of CD20 expression and its impact on the therapy."; 22 May 2014;
2. Oral presentation of data for evaluators of BASTION project "Regulation of CD20 expression and its impact on the therapy."; 30 January 2015;

#### **G. Poster presentation at the conferences**

1. 55<sup>th</sup> ASH Annual Meeting and Exposition , poster presentation "Inhibitors of SRC family and AKT regulate the activity of CD20 promoter" Pyrzynska B, Bojarczuk K, Winiarska M, Bil J, Miazek N, Zapala P, Bobrowicz M, Dwojak M, Siernicka M, Golab J, abstract published in *Blood*, 122 (21): abstract no.1838, New Orleans, USA, 7-10 December 2013;
2. AACR Annual Meeting, poster presentation "PTEN regulates the CD20 antigen expression and affects rituximab-based therapy of lymphoma malignancies." Pyrzynska B, Bojarczuk K, Siernicka M, Dwojak M, Bobrowicz M, Miazek N, Zapala P, Zagodzdon A., Bil J, Golab J, Winiarska M., late-breaking abstract no. LB-242/4, Philadelphia, USA, 18-22 April 2015.
3. Poster presentation "AKT and PTEN, two important players in the regulation of CD20 expression, affect the sensitivity of lymphoma malignancies to rituximab-based therapy." Pyrzynska B, Bojarczuk K, Dwojak M, Bobrowicz M, Siernicka M, Miazek N, Zapala P, Zagodzdon A, Bil J, Golab J, Winiarska M. Conference "TRON"; Warsaw, Poland, 21-22 May 2015.

#### **H. Participation in courses/trainings/workshops**

1. Training in scientific project management "Scientists of Tomorrow" organized by pm2pm; Warsaw, Poland; Feb-June 2013
2. Flow Cytometry Workshop "Apoptosis and Cell Signaling" organized by the Nencki Institute of Experimental Biology as part of BIO-IMAGINE project, Warsaw, Poland, 22 April 2013;
3. Workshop "Cancer genetics for medical community" organized by the Medical University of Warsaw as part of the BASTION project, Warsaw, Poland, 17 June, 2013;
4. Workshop "Commercialization of research results" organized by the Bio&Technology Innovations Platform of Biocentrum Ochota, Warsaw, Poland, 27 June 2013;
5. Workshop "Application of flow cytometry in Molecular Oncology"; Medical University of Warsaw, Poland, 15-16 October, 2014;
6. Workshop "Genome-wide methods in cancer genetics"; Medical University of Warsaw, Poland, 28 October 2014;
7. Info day "Horizon2020 – programs related to health care" ([www.kpk.gov.pl](http://www.kpk.gov.pl)); Warsaw, Poland, 18 November 2014,
8. Workshop "Marie Skłodowska-Curie Innovative Training Network" ([www.kpk.gov.pl](http://www.kpk.gov.pl)); Warsaw, Poland, 19 November 2014,
9. Workshop "Idea 2 Business"; Medical University of Warsaw, Poland, 26 November – 17 December 2014,
10. Workshop "Molecular Diagnostics in Cancer"; Medical University of Warsaw, Poland, 08 June 2015,
11. Workshop "Individual Fellowships - MSCA - how to apply" ([www.kpk.gov.pl](http://www.kpk.gov.pl)); Warsaw, Poland, 18 June 2015,

#### **I. Awards/fellowships obtained during BASTION project**

1. Mentoring Program awarded to Beata Pyrzynska by the Foundation for Polish Science - FNP (Mentor: Prof. M.A. Shipp, Dana-Farber Cancer Institute, Boston, USA) – June 2014- May 2015



#### **J. Students supervision**

Co-supervising PhD student: Michał Dwojak, supervising students (participants of student's scientific group at the Department of Immunology): Piotr Zapała and Nina Miązek

#### **K. Collaboration with other research teams started during BASTION project**

1. Prof. dr hab. Przemysław Juszczynski, Institute of Hematology and Transfusion Medicine, Warsaw, Poland;
2. Prof. Daniel Olive and Dr Cyril Fauriat, Cancer Research Center of Marseille (CRCM), University of Mediterranean, INSERM, Institut Paoli Calmettes, Marseille, France;

#### **L. International research visits during BASTION project**

1. Twinning visit to Cancer Research Center of Marseille (CRCM), University of Mediterranean, INSERM, Institut Paoli Calmettes, Marseille, France; 25 June – 24 July 2015.

#### **M. Current research interests**

Current research interest of Beata Pyrzynska is focused on the molecular mechanisms that regulate expression of CD20 antigen in malignant B-cells. Clinical management of B-cell tumors (particularly non-Hodgkin's lymphoma and chronic lymphocytic leukemia) includes treatment with monoclonal antibodies (such as rituximab, ofatumumab or GA-101) directed against CD20 antigen. Nevertheless, the resistance to this therapy is a frequent problem in the clinic. The resistance is often related to decreased levels of CD20 on the surface of malignant cells. Therefore, before investigating novel therapeutic combinations in cancer patients, the molecular mechanisms modulating the level of CD20 antigen, such as its transcriptional regulation, protein stability and its cellular localization should be taken into consideration. Looking for signaling pathways affecting these processes the research group of Dr. Winiarska has recently found that the BCR-SRC-AKT signaling is the key regulator of CD20 expression. Beata would like to extend the study mentioned above by elucidating the molecular mechanism acting downstream of AKT and leading to transcriptional repression of CD20 expression upon treatment with AKT inhibitors. She has already performed the detailed characterization of the effect of clinically used AKT inhibitors on CD20 expression and on the efficacy of anti-CD20 treatment using different cell lines as well as primary samples of B-cell tumors. Importantly, she would like to employ modern molecular approaches to find the signaling molecules acting downstream of AKT and contributing to the regulation of CD20 expression. She expects that the proteomic approaches and database searches will lead to the identification of proteins that recognize and bind to the region of CD20 promoter that we have recently identified to be critical for regulation by AKT.

#### **N. Envisioned career paths in BASTION project**

Beata Pyrzynska has managed to secure funding until August 7<sup>th</sup>, 2017 for research and her salary as principal investigator of the grant OPUS (NCN; 2013/11/B/NZ5/03240) „Influence of AKT signaling pathway on CD20 expression and antitumor activity of therapeutic monoclonal antibodies”. She would like to continue her studies in experimental hematology, in particular in anti-CD20 mAbs field. Moreover, we also expect that the results of Dr Pyrzynska project will allow her to develop scientific career in the field of monoclonal antibodies and establish her independent research team. In coming years she is also planning to present her results at the international conferences, publish the results of her project in peer-review journals and participate in other projects of Dr Winiarska group.



## V. Joanna Drzewinska-Chanko (TEAM of Tomasz Stoklosa)



| DATE (YEARS) | DEGREE/EXPERIENCE  | PLACE  | SUPERVISOR                 |
|--------------|--------------------|--|----------------------------|
| 2000-2005    | M.Sc.              | Department of Molecular Biophysics, University of Lodz       | Prof. dr hab. G. Bartosz   |
| 2005-2010    | Ph.D.              | Department of Molecular Biophysics, University of Lodz       | Prof. dr hab. M. Soszyński |
| 2011         | Research assistant | Department of Molecular Biophysics, University of Lodz       | Prof. dr hab. G. Bartosz   |
| 2012         | Postdoc            | Department of General Biophysics, University of Lodz         | Prof. dr hab. B. Klajnert  |
| 2013         | Postdoc            | BASTION, Department of Immunology, Warsaw Medical University | Dr T. Stoklosa             |

### A. Biosketch (provided by Joanna Drzewinska)

I obtained Master of Science degree at University of Lodz, Department of Molecular Biology in 2005. During my Master's studies, I worked on molecular cloning, recombinant expression and transcriptional regulation of human proteins ABCC1, ABCC2, ABCC3 by MAP kinases signal transduction pathways. Upon completion of my M.Sc., I started my Ph.D. project focused on characterization of transcriptional regulation of *DHCR24* gene, which encodes seladin-1 protein. I investigated the transcriptional activity of *DHCR24* promoter in various mammalian cell types in response to oxidative stress, overexpression of wide array of transcriptional factors and transcriptional factors' inducers. As a result of these studies I demonstrated that mechanisms of DNA methylation and histone acetylation are responsible for tissue specific expression of *DHCR24* gene (Drzewinska et al., 2011). Moreover I identified glucocorticoids as inducers of *DHCR24* expression acting by glucocorticoid receptor-mediated mechanism in lung cancer cells. I completed my Ph.D. in biological sciences in 2010 from the Department of Molecular Biophysics, University of Lodz. In 2011 I worked in the project "Role of multidrug transporters in pharmacokinetics and toxicology – in vitro tests in pharmaceutical and clinical practice" conducted at University of Lodz in the frames of Innovative Economy National Cohesion Strategy. During this time, I worked on molecular cloning and recombinant expression of ABC proteins responsible for multidrug resistance in cancer cells. In the year 2012 I joined the lab of Prof. Barbara Klajnert to work as Postdoctoral Researcher in the project "Biological properties and biomedical applications of dendrimers" conducted within the framework of the TEAM programme, University of Lodz. In these studies we showed that dendrimers (synthesized branched cationic polymers) form stable complexes with anti-HIV antisense oligonucleotides and effectively protect them from nucleolytic degradation. Furthermore, we demonstrated that modification of dendrimer's surface with carbohydrates improves dendrimer's capability to protect the oligonucleotides from digestion by serum nucleases or nuclease S1 (Drzewinska et al., 2012). We also studied interactions between



dendriplexes (complexes composed from dendrimers and oligonucleotides) and glucosaminoglycans (the main components of extracellular matrix) which may limit effectiveness of transfection. We were able to demonstrate that the effect of glucosaminoglycans on dendriplexes depends on the glucosaminoglycan type and the oligosaccharide serving as the surface group of the dendrimer (Szewczyk et al., 2012). In 2013 I joined BASTION project as a Postdoctoral Researcher in the Department of Immunology, Medical University of Warsaw.

#### B. Publications published during BASTION project

|          | <b>Authors, title, journal, year</b>   | <b>IF</b>    |
|----------|--|--------------|
| <b>1</b> | Szewczyk M, Drzewinska J, Dzmitruk V, Shcharbin D, Klajnert B, Appelhans D, Bryszewska M. Stability of Dendriplexes Formed by Anti-HIV Genetic Material and Poly(propylene imine) Dendrimers in the Presence of Glucosaminoglycans. <i>The Journal of Physical Chemistry B</i> . 2012; 116(50):14525-32. | <b>3,377</b> |
| <b>2</b> | Drzewinska J, Appelhans D, Voit B, Bryszewska M, Klajnert B. Poly(propylene imine) dendrimers modified with maltose or maltotriose protect phosphorothioate oligodeoxynucleotides against nuclease activity. <i>Biochemical and Biophysical Research Communications</i> . 2012 Oct 12;427(1):197-201     | <b>2,281</b> |

#### C. Grant applications submitted during BASTION project

1. Application for internal grant from Medical University of Warsaw. Project title: Investigation of influence of tyrosine kinases inhibitors on epigenetic changes in chronic myeloid leukemia – a potential association with drug resistance.
2. Grant application to National Science Center for funding “Opus” project entitled “Role of epigenetic mechanisms in chronic myeloid leukemia progression and resistance to targeted therapy” application registration number 2013/11/B/NZ2/02679
3. Grant application to Polish Ministry of Science and Higher Education for funding “luventus Plus” project entitled „Rola metylacji DNA w progresji przewlekłej białaczki szpikowej i mechanizmach lekooporności na terapię celowaną”, application registration number IP2014 007973

#### D. Poster presentation at the conferences

1. Drzewinska-Chanko J., Seferynska I., Machnicki M., Bajorek K., Wnuk M., Glodkowska-Mrowka E., Stoklosa T. “Tyrosine kinase inhibitors do not affect expression of DNA Methyltransferases and global methylation level in chronic myeloid leukemia cells”. 19th Congress of the European Hematology Association”, Milano 2014
2. Drzewinska-Chanko J., Bajorek K., Glodkowska-Mrowka E., Barankiewicz J., Machnicki M., Seferynska I., Stoklosa T. „Badanie wpływu inhibitorów kinaz tyrozynowych I, II, i III generacji na ekspresję metylotransferaz DNA w przewlekłej białaczce szpikowej”. Międzynarodowa Konferencja Szkoleniowa PTHiT „Hematologia Kliniczna i Doświadczalna”, Kazimierz Dolny 2014
3. Stoklosa T, Deregowska A., Drzewinska-Chanko J., Barankiewicz J., Machnicki M., Pruszczyk K., Wnuk M., “Effects of First and Next-Generation Tyrosine Kinase Inhibitors on Telomere-Mediated Chromosomal Instability in Chronic Myeloid Leukemia Cells”. *Blood*: 124 (21) December 6, 2014 ( 56th Annual Meeting and Exposition of The American Society of Hematology)

#### E. Current research interests

Research activities of Joanna concentrate on studying mechanisms of drug resistance in tumors with the main focus on haematological malignancies such as chronic lymphocytic leukaemia (CLL) and chronic myeloid leukaemia (CML). Introduction of tyrosine kinase inhibitors (TKI) occurred to be a milestone in targeted therapy of CML. However, drug resistance becomes an emerging problem with novel targeted therapies. Many reports demonstrated that epigenetic processes remarkably modulate CML expression profiles and phenotypic outcomes, but a lot of questions regarding epigenetic mechanisms of pathogenesis in CML remain unanswered.



Hence, unravelling the mechanisms of epigenetic changes in leukaemia cells may contribute to inhibition of development of resistance to TKIs and malignant progression of the disease. Although aberrant DNA methylation is considered to be associated with CML progression, there are almost no useful epigenetic biomarkers which would allow stratifying CML patients into groups with different risk and to personalize or change their treatment before clinical resistance will develop. Thus, she is studying the epigenetic landscape in CML with special focus on leukaemia stem cells (LSCs) which are intrinsically resistant to targeted therapy with tyrosine kinase inhibitors. Employing the next-generation sequencing technology in a follow-back study, she intends to define patterns of epigenetic changes in early phase of CML which predispose patients to the progression of the disease.

#### F. Envisioned career paths in BASTION project

Joanna Drzewińska-Chańko went for maternal leave in August 2013 (due to health problems during pregnancy she had to take sick leave already from April 2013) and she is on maternal leave till the end of the Project. Due to family reasons she is planning to come back to her hometown, Lodz and pursue her scientific career there.

Since Joanna Drzewińska-Chańko, postdoc hired to Dr Stoklosa team, due to health problems during pregnancy had to take sick leave already from April 2013, recruitment process was carried out to hire a postdoc for replacement. The procedure was performed according to the procedures of the Medical University of Warsaw. Advertisement was published on the online Nature Jobs website ([www.nature.com/naturejobs/science/](http://www.nature.com/naturejobs/science/)), EURAXESS website (<http://ec.europa.eu/euraxess/index.cfm/jobs/index>). Moreover, advertisement was announced on the Medical University of Warsaw website ([www.wum.edu.pl](http://www.wum.edu.pl)), distributed through email lists to international and domestic research centers (The International Institute of Molecular and Cell Biology IIMCB, Nencki Institute of Experimental Biology, University of Warsaw, Mossakowski Medical Research Centre, Polish Academy of Sciences, Institute of Haematology and Transfusion Medicine, Institute of Fundamental Technological Research Polish Academy of Sciences) and published in the second highest selling newspaper in Poland (Gazeta Wyborcza).

Announcement was published with deadline on 15th July 2014 (attachment 1).

The response to the advertisement for research positions was very good, however numerous candidates were formally ineligible for the positions. Tomasz Stoklosa as a leader assessed candidates' suitability for positions and assessment processes was focused upon the formal criteria and work-related qualities needed for positions (attachment 2).

The following six selection criteria were used for postdoc position:

| CRITERIA                  | WEIGHT (TOTAL OF 100) |
|---------------------------|-----------------------|
| Motivation letter         | 5                     |
| reference letters         | 5                     |
| Publications              | 50                    |
| Experience in the area    | 20                    |
| International experience  | 10                    |
| Additional qualifications | 10                    |

#### List of candidates for team of Tomasz Stoklosa:

(full list will be rejected from the public report)

1. Olena Bakhuryńska
2. Piotr Banski
3. Thomas Fricke
4. Paulina Gapska
5. Iwona Solarska
6. Katarzyna Solarska-Sciuk
7. Przemek Swiecki
8. Abhishek Narain Singh





9. Juan Alfonso Redondo
10. María Ramírez Arroyo
11. Tomasz Pelczar
12. Sheik Asraf

Short list was prepared by Tomasz Stoklosa by 23<sup>rd</sup> July 2014 and selected candidates were informed via e-mail by WP3 leader about the results and that they proceeded to the next stage of assessment. Interview took place on 6<sup>th</sup> August, 2014 in the Department of Immunology, MUW.

During the interview all applicants were informed about the objectives of BASTION project and were evaluated against the selection criteria for the position and how far they could contribute towards the achievements of the BASTION goals. The selection process was made by a selection committee. The rating scale (1-40) was used when assessing candidates against the selection criteria.

| Rating                  | Description  | Points       |
|-------------------------|--|--------------|
| <b>Highly qualified</b> | The candidate demonstrated experience/expertise above the advertised classification level.                             | <b>36-40</b> |
| <b>Very qualified</b>   | The candidate demonstrated experience/expertise to a high degree as described for the advertised classification level. | <b>30-35</b> |
| <b>Qualified</b>        | The candidate demonstrated experience/expertise as described for the advertised classification level.                  | <b>25-29</b> |
| <b>Not qualified</b>    | The candidate demonstrated some aspects of experience/expertise for the advertised classification level.               | <b>20-24</b> |
| <b>Not qualified</b>    | The candidate failed to provide experience/expertise demonstrative of the requirements of this position.               | <b>1- 19</b> |

#### Selection criteria

|   | Skill/Quality                         | Rating      |
|---|---------------------------------------|-------------|
| 1 | Depth and breadth of experience       | <b>1-5</b>  |
| 2 | Technical knowledge                   | <b>1-5</b>  |
| 3 | Interpersonal skills, teamwork        | <b>1-5</b>  |
| 4 | Organization and planning             | <b>1-5</b>  |
| 5 | Creativity                            | <b>1-5</b>  |
| 6 | Project planning, grant applications  | <b>1-5</b>  |
| 7 | Written and oral communication skills | <b>1-5</b>  |
| 8 | Coping stress management              | <b>1-5</b>  |
|   | <b>TOTAL</b>                          | <b>1-40</b> |

#### Members of the selection board:

1. Radoslaw Zagozdzon
2. Magdalena Winiarska
3. Tomasz Stoklosa

#### List of invited candidates:

1. Iwona Solarska
2. Thomas Fricke (name will be rejected from the public report)

Results were announced on 7<sup>th</sup> July 2014. Both invited candidates were informed about the results.



## VI. Iwona Solarska (TEAM of Tomasz Stokłosa) – hired for replacement



| Years       | Degree / experience                         | Place   | Supervisor                             |
|-------------|---|---|--|
| 06/1995     | Medical laboratory Technician               | Medical College in Warsaw, Poland   | -                                      |
| 1995 - 1999 | Technician                                  | Department of Microbiology, Institute of Hematology and Transfusion Medicine, Warsaw, Poland                        | M.Sc. Irena Bednarska                  |
| 06/1999     | M.Sc.                                       | Faculty of Biology, University of Lodz, Lodz, Poland  | Prof. Barbara Różalska                 |
| 1999 - 2004 | Biologist                                   | Department of Microbiology, Institute of Hematology and Blood Transfusion, Warsaw, Poland                           | M.Sc. Maria Zaleska                    |
| 2004 – 2014 | Assistant-specialist in laboratory medicine | Genetic Laboratory, Diagnostic Hematology Department, Institute of Hematology and Blood Transfusion, Warsaw, Poland | Prof. Monika Prochorec-Sobieszek, M.D. |
| 06/2009     | Ph.D.                                       | Institute of Hematology and Blood Transfusion, Warsaw, Poland   | Prof. Krzysztof Warzocha, M.D.         |
| 2014-2015   | Postdoc                                     | Department of Immunology, Medical University of Warsaw, Poland  | Tomasz Stokłosa M.D. Ph.D              |



### Biosketch (provided by Iwona Solarska)

I graduated from the Faculty of Biology at University of Lodz in 1999. My Master's thesis, entitled 'Phenotype characteristic of the *Staphylococcus* isolated from the blood of patients with hematological disorders' was performed in the Department of Infectious Biology, Institute of the Microbiology and Immunology, University of Lodz, Poland and in the Department of Microbiology in the Institute of Hematology and Blood Transfusion in Warsaw, where I was employed first as a technician, and then as a biologist (from 1999).

In October, 2006 I commenced PhD studies at the Molecular Genetic Department in the Institute of Hematology and Blood Transfusion in Warsaw, under supervision of Professor Krzysztof Warzocha. From the beginning of my work I was strongly interested in the area of chronic myeloid leukemia (CML), so the research I conducted within the topic of my PhD thesis consisted of the analysis of the minimal residual disease (*BCR-ABL* gene expression) in a group of CML patients treated with allogeneic stem cell transplantation. I was trying to determine the prognostic levels of minimal residual disease that correlate with risk of leukemia relapse. Finally I received my doctoral degree in medical sciences in June, 2009 from the Institute of Hematology and Blood Transfusion in Warsaw.

In the 2004 – 2014 in the Institute of Hematology and Blood Transfusion I was responsible for managing the registry of newly diagnosed patients with CML within the cooperation with Polish Society of Hematology and Transfusiology and European Leukemia Net (ELN). I participated also in Polish first control for the standardization of *BCR-ABL* quantitative PCR method within the collaboration with ELN, in a lab rounds control for minor-*BCR-ABL* within the collaboration between European Study Group on MRD detection in ALL (ESG-MRD-ALL) and European Working Group for Adult Acute Lymphoblastic Leukemia (EWALL), and in the Polish project MapTest – detection of the *BCR-ABL* mutations in a Polish population CML and ALL Ph+ patients. I was committed to some researches, including investigation of molecular mechanisms of progression and primary resistance to imatinib in CML (2006 – 2009), investigation of clinical implications of the somatic hypermutation VDJ genes, T-cell receptor and protooncogenes in non-Hodgkin lymphomas (2005 – 2007) or analysis the multi-drug resistance *MDR1*, *MRP*, *LRP*, *BCRP* genes in adult patients with non-lymphoblastic acute leukemia (2009 – 2012). In 2011 – 2012 I supervised the project 'Analysis of coexpression of endogenous transcripts of bidirectional genes *BAALC* and *C8orf56* in AML patients and its clinical implications'.

In 2014 I joined BASTION project as a Postdoctoral Researcher in the Department of Immunology, Medical University of Warsaw.

#### A. Publications published during BASTION project

|   | Authors, title, journal, year  | IF    |
|---|--|-------|
| 1 | Nasilowska-Adamska B, Czyz A, Markiewicz M, Rzepecki P, Piatkowska-Jakubas B, Paluszewska M, <b>Solarska I</b> , Dzierzak-Mietla M, Borg K, Prochorec-Sobieszek M, Szydlo R, Lewandowski K, Skotnicki A, Jedrzejczak WW, Kyrzch-Krzemien S, Komarnicki M, Warzocha K. Mild chronic graft versus host disease may alleviate poor prognosis associated with FLT3-internal tandem duplication for adult acute myeloid leukemia following allogeneic stem cell transplantation with myeloablative conditioning in first complete remission; a retrospective study. <b>Eur J Haematol.</b> 2015 Apr 27. [Epub ahead of print] | 2,066 |
| 2 | Nasilowska-Adamska B, <b>Solarska I</b> , Paluszewska M, Malinowska I, Jedrzejczak WW, Warzocha K. FLT3-ITD and MLL-PTD influence the expression of MDR-1, MRP-1, and BCRP mRNA but not LRP mRNA assessed with RQ-PCR method in adult acute myeloid leukemia. <b>Ann. Hematol.</b> 2014, 93(4): 577-593.   | 2,634 |
| 3 | Glodkowska-Mrowka E, <b>Solarska I</b> , Mrowka P, Bajorek K, Niesiobedzka-Krezel J, Seferynska I, Borg K, Stoklosa T. Differential expression of BIRC family genes in chronic myeloid leukemia BIRC3 and BIRC8 as potential new candidates to identify disease progression. <b>Br J Haematol</b> 2014, 164(5): 740-742.   | 4,711 |



## **B. Participation in the conferences during BASTION project**

1. Conference: 'CML Forum Experts Meeting'. Gdansk, 19-21 February 2015.
2. Translational Research in Oncology in New Member State Economies International Conference, Warsaw, 21-22 May 2015
3. European Hematology Association Congress, Vienna, Austria, 11-14 June 2015

## **C. Poster presentation at the conferences**

1. Marcin M. Machnicki, Joanna Niesiobedzka-Krezel, Iwona Solarska, Piotr Stawinski, Rafal Ploski, Tomasz Stokłosa. *Unraveling the mechanism of chronic myeloid leukemia progression by next-generation sequencing of leukemic progenitor and stem cells.*
2. FEBS-EMBO Conference, 1-4 September 2014, Paris, France. Abstract in: FEBS Journal 281(Suppl.1):782
3. Marcin M. Machnicki, Joanna Niesiobędzka-Kręzel, Iwona Solarska, Piotr Stawiński, Rafał Płoski i Tomasz Stokłosa. *Poszukiwanie nowych zmian genetycznych odpowiedzialnych za oporność na terapię celowaną i progresję przewlekłej białaczki szpikowej przy zastosowaniu sekwencjonowania wysokoprzepustowego.*
4. Międzynarodowa Konferencja Szkoleniowa PTHiT „Hematologia Kliniczna i Doświadczalna”. 16-18 May, 2014, Kazimierz Dolny, Poland.
5. Marcin M. Machnicki, Joanna Niesiobedzka-Krezel, Iwona Solarska, Piotr Stawinski, Rafal Ploski and Tomasz Stokłosa. *Exome and custom-gene sequencing as tools for analysis of chronic myeloid leukemia progression. Conference: "NGS Milan 2015: From the clinic To single cell analysis", 9-10 March, 2015.*
6. Iwona Solarska, Marcin M. Machnicki, Joanna Niesiobedzka-Krezel, Piotr Stawinski, Rafal Ploski, Tomasz Stokłosa. *Comparison of whole-exom and custom-gene sequencing as tools for analysis of chronic myeloid leukemia progression.*
7. Translational Research in Oncology in New Member State Economies Conference, Warsaw, 21-22 May, 2015
8. Iwona Solarska, Marcin M. Machnicki, Barbara Nasiłowska-Adamska, Barbara Pieńkowska-Grela, Ilona Seferyńska, Piotr Stawiński, Rafał Płoski and Tomasz Stokłosa
9. *Selection of RUNX1-mutated clone associated with relapse and blast crisis in chronic myeloid leukemia patient after allo-HSCT as revealed by targeted enrichment and deep sequencing.*
10. European Hematology Association, Vienna, Austria 11-14 June, 2015
11. Marta Libura, Sebastian Giebel, Beata Piątkowska-Jakubas, Marta Przestrzelska Pawełczyk, Izabella Florek, Karolina Matiakowska, Bożena Jaźwiec, Katarzyna Borg, Iwona Solarska, Magdalena Zawada, Sylwia Czekalska, Jolanta Libura, Małgorzata Jakóbczyk, Karolina Karabin, Małgorzata Calbecka, Justyna Gajkowska-Kulig, Grażyna Gadomska, Marek Kiełbiński, Anna Ejduk, Dariusz Kata, Sebastian Grosicki, Agnieszka Wierzbowska, Sławomira Kyrz-Krzemień, Krzysztof Warzocha, Kazimierz Kuliczkowski, Aleksander Skotnicki, Jerzy Holowiecki, Wiesław Jedrzejczak, Olga Haus.
12. *Favorable outcome of patients with normal karyotype acute myeloid leukemia harboring FLT3-ITD and treated with cladribine added induction.* European Hematology Association, Vienna, Austria 11-14 June, 2015



#### **D. Participation in courses/trainings/workshops**

1. Workshop “Genome-wide methods in cancer genetics”; Medical University of Warsaw, Poland, 28 October 2014;
2. Workshop “Application of flow cytometry in Molecular Oncology”; Medical University of Warsaw, Poland, 15-16 October, 2014;
3. Workshop: “Molecular diagnostic in cancer” organized by BASTION, Medical University of Warsaw, 8 June 2015.

#### **E. Current research interests**

Iwona Solarska is a molecular biologist working in the field of molecular hematology. Her current research interest is focused on the molecular mechanisms that determined drug resistance and progression of hematological malignancies. The projects she is currently involved in aim at understanding molecular pathogenesis of chronic myeloid leukemia (CML) with special regard to mechanisms of resistance to tyrosine kinase inhibitors (TKI) and potential ways to overcome such resistance. TKIs resistance becomes an emerging problem with novel targeted therapies. For most of CML patients a therapy with TKIs is very effective, but some of them are resistant to therapy or even progressed to more advanced phases of the disease. That's why there are still a lot of questions regarding drug resistance mechanisms or progression pathways of pathogenesis in CML to answered. The second line of her scientific interest is focused on the acute myeloid leukemias (AML) including investigation of the multi-drug resistance genes and its correlation with other prognostic factors. This is a very heterogeneous group of malignancies, so every finding appeared to be a milestone to understanding the biology and improving prognosis for AML patients.

#### **F. Envisioned career paths in BASTION project**

Iwona Solarska hired for replacement in September 2014 after the end of BASTION Project will come back to her previous position in the Institute of Hematology and Blood Transfusion in Warsaw and will continue to collaborate in the genetic studies on leukemia with Dr Stoklosa team.



## VII. Magdalena Banach-Orlowska (TEAM of Pawel Wlodarski)



| DATE (YEARS)         | DEGREE/EXPERIENCE             | PLACE   | SUPERVISOR              |
|----------------------|-------------------------------|---|-------------------------|
| 1999                 | M.Sc. in molecular biology    | Department of Genetics, University of Warsaw, Poland  | Prof. Piotr Weglenski   |
| 2005                 | PhD in biochemistry           | Institute of Biochemistry and Biophysics, PAS in Warsaw, Poland                                     | Prof. Piotr Jonczyk     |
| 2006 - 2013          | Postdoc                       | Laboratory of Cell Biology at International Institute of Molecular and Cell Biology, Warsaw, Poland | Prof. Marta Miaczynska  |
| 2013 – until present | Postdoc (research specialist) | Department of Histology and Embryology, Medical University of Warsaw                                | Dr hab. Pawel Wlodarski |

### A. Biosketch (provided by Magdalena Banach-Orlowska)

I graduated from the Department of Genetics, University of Warsaw in 1999. During my graduate studies in the laboratory of Prof. Weglenski I studied the regulation of *Aspergillus nidulans* *agaA* gene.

After completing my M.Sc., I successfully applied for the PhD-tract in the Laboratory of Mutagenesis and DNA Repair at the Institute of Biochemistry and Biophysics, PAS. The aim of my PhD project, being a part of systematic studies on a replication fidelity in Prokaryota, was to determine the role of DNA polymerase II and DNA polymerase IV in replication. After publishing the results of my studies I received my doctoral degree (PhD) in biochemistry in 2005. In the years 2005–2006, I continued to work with the group of Prof. Fijalkowska and Prof. Jonczyk at the Institute of Biochemistry and Biophysics as a research fellow.

In the years 2006–2013, I was employed as a Postdoctoral Fellow at the International Institute of Molecular and Cell Biology. At that time I participated in studies concerning the role of endocytic proteins in signal transduction in mammalian cells. I have characterized the relationship between APPL1 endocytic protein and the nuclear repressor complex NuRD and its consequence for gene expression. We also demonstrated the role of APPL2 protein in survival of glioma cells. Since APPL adapter proteins interact with many partners involved in signal transduction we investigated their role in several signalling pathways. In 2010, I received grant from Foundation for Polish Science for investigation of the role of APPL1 protein in Wnt signaling. During my postdoctoral fellowship at Prof. Miaczynka Lab I was also involved in the project concerning the role of endocytic proteins in the NFκB pathway.

In 2013, I moved to dr hab. Pawel Wlodarski Lab at the Department of Histology and Embryology, Medical University of Warsaw. Since then I have been involved in three lines of work. The first one focuses on genetic basis of endometriosis. The second line of studies concerns the epigenetic regulation of gene expression. Employing NGS technology we plan to perform systematic analysis of methylation profile and in consequence find the changes in gene expression in response to female sex hormones (estradiol and progesterone).

The third line of work conducted in cooperation with Prof. Tomasz Ciach Lab from Warsaw University of Technology is devoted to investigation of the novel nanoparticles containing anticancer drug. Within this project we perform in vitro and in vivo studies of the new drug. I am particularly involved in investigating the intracellular trafficking of modified drug in breast and ovarian cancer cell lines.



## B. Publications published during BASTION project

|          | <b>Authors, title, journal, year</b>  | <b>IF</b>    |
|----------|---|--------------|
| <b>1</b> | Pyrzynska B, Banach-Orlowska M, Teperek-Tkacz M, Miekus K, Drabik G, Majka M, Miaczynska M. Multifunctional protein APPL2 contributes to survival of human glioma cells. <i>Mol Oncol.</i> 2013 Feb;7(1):67-84. | <b>5,935</b> |
| <b>2</b> | Banach-Orlowska M, Szymanska E, Miaczynska M. APPL1 endocytic adaptor as a fine tuner of Dvl2-induced transcription. <i>FEBS Lett.</i> 2015 Feb 13;589(4):532-9.  | <b>3,169</b> |

## C. Grant applications submitted during BASTION project

1. Searching for the novel miRNAs and isomiRNAs in endometriosis (OPUS 8 – NSC) – as a principal investigator – failed

## D. Participation in grants during BASTION project

1. Exome-wide search for somatic mutations in pathogenesis of endometriosis – OPUS 5 (NSC) - Main contractor
2. Epigenetic regulation of expression of genes involved in extracellular matrix remodeling and angiogenesis during development of endometriosis – contractor

## E. Participation in the conferences during BASTION project

1. Application of flow cytometry in molecular oncology (BASTION workshop, 15-17 October, 2014)
2. Cancer genetics for medical community, July, 2013 Warsaw

## F. Participation in courses/trainings/workshops

1. Microdissection - MicroBeam IV (training performed by Advanced Imaging Microscopy Specialist - Zeiss ) October, 2013
2. Technology Day- miRNA solutions from profiling to validation" - Life Technologies 14.11.2013, Warsaw

## G. Current research interests

For several years scientific interest of Magdalena concentrated on signal transduction and gene regulation. Since she joined BASTION program her scientific activity focuses on understanding the mechanism of endometriosis development – disease which in many cases leads to endometrioid or clear-cell ovarian cancer. The aim of this project is to identify mutation predisposing to endometriosis development and establish the origin of ectopic lesions. Since in some of the affected individuals, endometriosis develops into endometrioid or clear-cell ovarian cancer the identification of novel mutations could be helpful in understanding the cancer development. We have been sequencing DNA isolated from the eutopic and ectopic endometrium of affected woman as well as eutopic endometrium of healthy individuals using powerful Next Generation Sequencing (NGS) technology. The proposal for this project (accepted for founding by National Science Center) has been prepared by Dr Wlodarski group with Magdalena contribution, and she was to be the main executor of the NGS experiments.

## H. Envisioned career paths in BASTION project

Dr Magdalena Banach- Orłowska has unexpectedly left the team on April 30, 2015. Her plans have not been disclosed neither to any of the team members (who are currently continuing her tasks) not to the group leader.



Since Magdalena Banach-Orlowska, postdoc hired to Dr Wlodarski team, unexpectedly left the team, recruitment process was carried out to hire a postdoc for replacement. The procedure was performed according to the procedures of the Medical University of Warsaw.

Advertisement was published on the online Nature Jobs website ([www.nature.com/naturejobs/science/](http://www.nature.com/naturejobs/science/)), EURAXESS website (<http://ec.europa.eu/euraxess/index.cfm/jobs/index>). Moreover, advertisement was announced on the Medical University of Warsaw website ([www.wum.edu.pl](http://www.wum.edu.pl)) and published in the second highest selling newspaper in Poland (Gazeta Wyborcza).

Announcement was published with deadline on 29th April 2015 (attachment 3).

The response to the advertisement for research position was rather poor – only one candidate declared an interest. Pawel Wlodarski as a leader assessed candidate's suitability for position and assessment process was focused upon the formal criteria and work-related qualities needed for position (attachment 2).

The following six selection criteria were used for postdoc position:

| CRITERIA                  | WEIGHT (TOTAL OF 100) |
|---------------------------|-----------------------|
| Motivation letter         | 5                     |
| reference letters         | 5                     |
| Publications              | 50                    |
| Experience in the area    | 20                    |
| International experience  | 10                    |
| Additional qualifications | 10                    |

#### List of candidates for team of Pawel Wlodarski:

1. Agnieszka Pollak

The candidate was informed via e-mail by WP3 leader about the interview process. Interview took place on 30<sup>th</sup> April, 2015 in the Department of Medical Genetics, MUW.

During the interview applicant was informed about the objectives of BASTION project and was evaluated against the selection criteria for the position and how far they could contribute towards the achievements of the BASTION goals. The selection process was made by a selection committee. The rating scale (1-40) was used when assessing candidates against the selection criteria.

| Rating                  | Description  | Points       |
|-------------------------|--|--------------|
| <b>Highly qualified</b> | The candidate demonstrated experience/expertise above the advertised classification level.                             | <b>36-40</b> |
| <b>Very qualified</b>   | The candidate demonstrated experience/expertise to a high degree as described for the advertised classification level. | <b>30-35</b> |
| <b>Qualified</b>        | The candidate demonstrated experience/expertise as described for the advertised classification level.                  | <b>25-29</b> |
| <b>Not qualified</b>    | The candidate demonstrated some aspects of experience/expertise for the advertised classification level.               | <b>20-24</b> |
| <b>Not qualified</b>    | The candidate failed to provide experience/expertise demonstrative of the requirements of this position.               | <b>1- 19</b> |

#### Selection criteria

|   | Skill/Quality                   | Rating     |
|---|---------------------------------|------------|
| 1 | Depth and breadth of experience | <b>1-5</b> |
| 2 | Technical knowledge             | <b>1-5</b> |
| 3 | Interpersonal skills, teamwork  | <b>1-5</b> |





|   |                                       |             |
|---|---------------------------------------|-------------|
| 4 | Organization and planning             | 1-5         |
| 5 | Creativity                            | 1-5         |
| 6 | Project planning, grant applications  | 1-5         |
| 7 | Written and oral communication skills | 1-5         |
| 8 | Coping stress management              | 1-5         |
|   | <b>TOTAL</b>                          | <b>1-40</b> |

**Members of the selection board:**

1. Pawel Wlodarski
2. Rafal Ploski
3. Tomasz Stoklosa

**List of invited candidates:**

1. Agnieszka Pollak



### VIII. Agnieszka Pollak (TEAM of Pawel Wlodarski) – hired for replacement



| DATE (YEARS) | DEGREE/EXPERIENCE | PLACE  |
|--------------|-------------------|--|
| 1995–2000    | M.Sc.             | M.Sc. in Biology, with a specialization in Molecular Biology at University of Warsaw. Dissertation title: “Analysis of <i>suDpro</i> gene, proline mutations suppressor in <i>Aspergillus nidulans</i> ”         |
| 2003-10-15   |                   | title "laboratory diagnostician"   |
| 2010         | PhD               | Ph.D. in Medicine with a specialization in Medical Biology at Warsaw Medical University. Dissertation title: “Connexins associated deafness: spectrum of mutations and clinical phenotype among polish patients” |
| 2012         |                   | start of specialization in medical genetics laboratory   |

#### A. Publications (last 5 years)

1. Iwanicka-Pronicka K, **Pollak A**, Skórka A, Lechowicz U, Korniszewski L, Westfal P, Skarżyński H, Płoski R. Audio profiles in mitochondrial deafness m.1555A>G and m.3243A>G show distinct differences. *Med Sci Monit.* 2015 Mar 6;21:694-700. doi: 10.12659/MSM.890965. **IF=1,21**
2. Barg E, Skarżyńska M, **Pollak A**, Ślęzak R, Głąb E, Petriczko E, Józwa A, Sąsiadek MM. Uncommon constellation of multiglandular deficiency with 2 mutations in AIRE gene in an 18-year-old girl - 12 years of observation. *Endokrynol Pol.* 2014;65(6):514-8. doi: 10.5603/EP.2014.0070. **IF=1,21**
3. \*Ołdak M, Ścieżyńska A, Młynarski W, Borowiec M, Ruszkowska E, Szulborski K, **Pollak A**, Kosińska J, Mueller-Malesińska M, Stawiński P, Szaflik JP, Płoski R. Evidence against RAB40AL being the locus for Martin-Probst X-linked deafness-intellectual disability syndrome. *Hum Mutat.* 2014 Oct;35(10):1171-4. doi: 10.1002/humu.22620. Epub 2014 Aug 7. PubMed PMID: 25044830. **IF= 5.05**
4. \*Ołdak M, Ruszkowska E, **Pollak A**, Sobczyk-Kopciół A, Kowalewski C, Piwońska A, Drygas W, Płoski R. A note of caution on the diagnosis of Martin-Probst syndrome by the detection of the p.D59G mutation in the RAB40AL gene. *Eur J Pediatr.* 2014 Nov 5. [Epub ahead of print] PubMed PMID: 25370018. **IF=1.98**
5. \*Kostera-Pruszczyk A, Kosinska J, **Pollak A**, Stawinski P, Walczak A, Wasilewska K, Potulska-Chromik A, Szczudlik P, Kaminska A, Ploski R. Exome sequencing reveals mutations in MFN2 and GDAP1 in severe



Charcot-Marie-Tooth disease. *J Peripher Nerv Syst*. 2014 Nov 18. doi: 10.1111/jns.12088. [Epub ahead of print] PubMed PMID: 25403865. **IF= 2.50**

6. Pawlak A, Pronicki M, Iwanicka-Pronicka K, Kuśnierz J, Płoski R, **Pollak A**, Gil RJ. [Cardiological manifestation of MELAS syndrome associated with mutation at position 3234]. *Kardiol Pol*. 2014;72(1):83. doi: 10.5603/KP.2014.0009. Polish. PubMed PMID: 24469753. **IF=0,53**
7. \*Ploski R, **Pollak A**, Müller S, Franaszczyk M, Michalak E, Kosinska J, Stawinski P, Spiewak M, Seggewiss H, Bilinska ZT. Does p.Q247X in TRIM63 cause human hypertrophic cardiomyopathy? *Circ Res*. 2014 Jan 17;114(2):e2-5. doi: 10.1161/CIRCRESAHA.114.302662. PubMed PMID: 24436435. **IF=11,86**
8. Iwanicka-Pronicka K, **Pollak A**, Skórka A, Lechowicz U, Pajdowska M, Furmanek M, Rzeski M, Korniszewski L, Skarżyński H, Płoski R. Postlingual hearing loss as a mitochondrial 3243A>G mutation phenotype. *PLoS One*. 2012;7(10):e44054. doi: 10.1371/journal.pone.0044054. Epub 2012 Oct 25. PubMed PMID: 23133508; PubMed Central PMCID: PMC3485002. **IF=3,73**
9. Moraitou M, Dimitriou E, Mavridou I, Michelakakis H, Georgouli H, Ploski R, **Pollak A**. Transferrin isoelectric focusing and plasma lysosomal enzyme activities in the diagnosis and follow-up of hereditary fructose intolerance. *Clin Chim Acta*. 2012 Oct 9;413(19-20):1714-5. doi: 10.1016/j.cca.2012.06.010. Epub 2012 Jun 8. PubMed PMID: 22713622. **IF=2,53**
10. **Pollak A**, Mueller-Malesinska M, Lechowicz U, Skorka A, Korniszewski L, Sobczyk-Kopciol A, Waskiewicz A, Broda G, Iwanicka-Pronicka K, Oldak M, Skarzynski H, Płoski R. MTHFR 677T is a strong determinant of the degree of hearing loss among Polish males with postlingual sensorineural hearing impairment. *DNA Cell Biol*. 2012 Jul;31(7):1267-73. doi: 10.1089/dna.2012.1607. Epub 2012 Mar 16. PubMed PMID: 22424391; PubMed Central PMCID: PMC3391488. **IF=2,34**
11. Rydzanicz M, Cywińska K, Wróbel M, **Pollak A**, Gawęcki W, Wojsyk-Banaszak I, Lechowicz U, Mueller-Malesińska M, Ołdak M, Płoski R, Skarżyński H, Szyfter K, Szyfter W. The contribution of the mitochondrial COI/tRNA(Ser(UCN)) gene mutations to non-syndromic and aminoglycoside-induced hearing loss in Polish patients. *Mol Genet Metab*. 2011 Sep-Oct;104(1-2):153-9. doi: 10.1016/j.ymgme.2011.05.004. Epub 2011 May 13. PubMed PMID: 21621438. **IF=3,19**
12. Rydzanicz M, Wróbel M, **Pollak A**, Gawecki W, Brauze D, Kostrzewska-Poczekaj M, Wojsyk-Banaszak I, Lechowicz U, Mueller-Malesińska M, Ołdak M, Płoski R, Skarżyński H, Szyfter K. Mutation analysis of mitochondrial 12S rRNA gene in Polish patients with non-syndromic and aminoglycoside-induced hearing loss. *Biochem Biophys Res Commun*. 2010 Apr 23;395(1):116-21. doi: 10.1016/j.bbrc.2010.03.149. Epub 2010 Mar 28. PubMed PMID: 20353758. **IF=2,59**

## **B. Envisioned career path**

Dr Agnieszka Pollak, who has been recruited in May to complete the tasks assigned initially to dr Banach, is planning to work with the team and finish the project on endometriosis, that is planned beyond the date of cessation of BASTION.



## IX. Oksana Kovtonyuk (TEAM of Piotr Religa)



| DATE (YEARS) | DEGREE/ EXPERIENCE | PLACE  | SUPERVISOR                                      |
|--------------|--------------------|--|---|
| 1995–2000    | M.Sc.              | Zhytomyr State University, Faculty of Biology and Chemistry.   | Dr. Sergej V.Verevka<br>Dr. Vladimir N. Listvan |
| 2008         | PhD                | R.E. Kavetsky Institute of Experimental Pathology, Oncology and Radiobiology of the National Academy of Sciences of Ukraine, Department of Anticancer Therapy Mechanisms | Prof., MD. Vasyl F. Chekhun                     |
| 2009-2010    | Postdoc            | Jagiellonian University Faculty of Biochemistry, Biophysics and Biotechnology, Laboratory of Molecular Genetics and Virology   | Prof. Hanna Rokita                              |
| 2011-2012    | Postdoc            | Taras Shevchenko National University of Kyiv, Ukraine. The Institute of Higher Technology, Molecular Biology, Biotechnology and Biophysics Dept.                         | Prof. Lidia S. Kholodna                         |
| 2012         | Postdoc            | Laboratory of Mutagenesis and DNA Repair, Institute of Biochemistry and Biophysics, Polish Academy of Sciences   | Prof. Iwona J. Fijalkowska                      |

### A. Biosketch (provided by Oksana Kovtonyuk)

In 2000, after graduating from the Faculty of Natural Sciences, Zhytomyr I. Franko State University (Ukraine), I joined the Department of Anticancer Therapy Mechanisms at R.E. Kavetsky Institute of Experimental Pathology, Oncology and Radiobiology of the National Academy of Sciences of Ukraine and started my PhD thesis. Starting from 2004, I was employed as a young scientist in the same department. I worked on the study of the proteinase-antiproteinase balance in the dynamics of the growth of Guerin carcinoma and Lewis lung carcinoma with induced cisplatin resistance. In these studies we were able to show that tumor resistance to cisplatin is accompanied by significant changes of the kinetics of tumour growth. The change of the growth kinetics has been found to associate with the elevation of total proteolytic activity as well as the level of  $\alpha_1$ -proteinase inhibitor and decreased  $\alpha_2$ -macroglobuline level in blood plasma. It has been shown that cisplatin resistance is accompanied by the imbalance between proteolytic and antiproteolytic activities shifted to the total proteolytic activity increase in blood plasma and tumour tissue. Furthermore, we were able to demonstrate that Lewis lung carcinoma cisplatin drug resistance development is accompanied by the increase of its metastasis together with the elevation of total proteolytic activity in blood plasma.

In 2008, I have received my doctoral degree in oncology. To increase my expertise in the field of cancer biology, I moved to the Laboratory of Molecular Genetics and Virology of Prof. H. Rokita (Faculty of Biochemistry, Biophysics and Biotechnology, Jagiellonian University, Krakow, Poland) as a Postdoctoral Fellow within Marie-Curie fellowship funded under the EU's Seventh Framework Programme. My research project focused on MCPIP



(Monocyte chemoattractant protein-induced protein) function in human neuroblastoma cell lines. During my 13-month stay at the lab, I worked on stable and transient transfection of neuroblastoma cells with mutant MCPIP forms and characterization of the clones at the level of MCPIP content, their proliferation and viability. In the years 2011-2012, I worked as a Staff Scientist at the Taras Shevchenko National University of Kyiv, Ukraine. In 2012, I joined as a postdoctoral training fellow the laboratory of Prof. Iwona Fijalkowska at the Institute of Biochemistry and Biophysics, Polish Academy of Sciences. During my 6 month stay at the lab I was working on a research project entitled "New players involved in the maintenance of genomic stability". I studied the role of PSF1 (a subunit of the GINS complex, which plays a key role at DNA replication forks) mutants in ensuring genomic microsatellite stability using *Saccharomyces Cerevisiae* cell as a model organism.

#### B. Publications published during BASTION project

|   | Authors, title, journal, year   | IF    |
|---|---|-------|
| 1 | PatenYaiw KC, Mohammad AA, Taher C, Wilhelmi V, Davoudi B, Straat K, Assinger A, Ovchinnikova O, Shlyakhto E, Rahbar A, Kovtonyuk O, Religa P, Butler L, Khan Z, Streblow D, Pernow J, Söderberg-Nauclér C. Human cytomegalovirus induces upregulation of arginase II: possible implications for vasculopathies. <i>Basic Res Cardiol.</i> 2014 Mar;109(2):401. doi: 10.1007/s00395-014-0401-5. Epub 2014 Jan 19. | 5,414 |

#### C. Grant applications submitted during BASTION project

1. CTCs-based novel diagnostic and screening method in cancer diseases – 3<sup>rd</sup> competition of PBS (NCBiR) – not funded
2. OPUS 8 (National Science Center) grant submitted: "Role of fibronectin in regulation of endothelial-mesenchymal transition in *in vitro* and *in vivo* models of colorectal cancer". Function: Principal Investigator – not funded

#### D. Participation in grants during BASTION project

1. grant of National Center of Science (Grant number 2011/01/B/NZ4/06635) „Effect of tumor biology of circulating tumor cells” . Co-investigator. 2013-2014.

#### E. Participation in the conferences during BASTION project

1. COST working group meeting (19.09.2013-20.09.2013) Warsaw, Poland.
2. International Conference Translational Research in Oncology in New Member State Economies, Warsaw, Poland, 2015.

#### F. Poster presentation at the conferences

1. International Conference Translational Research in Oncology in New Member State Economies ("TRON") Kovtonyuk O, Ananthaseshan S., Soin J., Religa P. Evaluation of the role of CMV infection in colorectal cancer progression, Warsaw, Poland, 21-22 May 2015.
2. COST working group meeting. Soin J., Kovtonyuk O., Bojakowski K., Religa P. The role of CMV infection in tumor progression. Warsaw, Poland, 2013.

#### G. Participation in courses/trainings/workshops

1. Employing genome-wide technologies for functional regulation in development and disease course (2742), RIKEN Division of Genomic Technologies and Karolinska Institutet, 24 February – 28 April 2014.
2. Tumor virology course (1592) at Karolinska Institutet, 31 March -04 April 2014.
3. Apoptosis: Theory and Methods course (1201) at Karolinska Institutet, 07-11 April 2014.



4. "Application of flow cytometry in molecular oncology" workshop organized by the Medical University of Warsaw as part of the BASTION project, Warsaw, 15-16 October 2014.
5. "Genome wide methods in cancer genetics" workshop organized by the Medical University of Warsaw as part of the BASTION project, Warsaw, 28 October 2014.
6. Workshop "Idea 2 Business"; Medical University of Warsaw, Poland, 26 November – 17 December 2014.
7. Workshop on flow cytometry, BASTION project, 6 May 2015.
8. "Exome analysis using the next generation sequencing platform Ion Proton", BASTION project, 30-31 July 2015.
9. "Transcriptome analysis (RNA-seq)", BASTION project, 5-7 August 2015.

#### **H. Students supervision**

1. PhD student Varsha Prakash
2. PhD student Sharan Ananthaseshan

#### **I. Collaboration with other research teams started during BASTION project**

1. Prof. Cecilia Söderberg-Nauclér research group, Department of Medicine, Karolinska Institutet, Cell and Molecular Immunology, Karolinska University Hospital, Stockholm, Sweden.
2. Alice Assinger, PhD, Institute of Physiology, Centre of Physiology and Pharmacology, Vienna, Austria
3. Klas Strååt, PhD, Department of Cell and Molecular Biology, Karolinska University Hospital, Stockholm, Sweden.
4. Mariusz Sacharczuk, PhD, DSc, Department of Molecular Cytogenetics, Institute of Genetics and Animal Breeding of the Polish Academy of Sciences, Jastrzębiec, Poland.
5. Krzysztof Bojakowski, MD, PhD, DSc, Department of General, Vascular and Oncological Surgery, CSK MSW, Warsaw, Poland; Department of Immunology, Biochemistry and Nutrition, Medical University of Warsaw, Warsaw, Poland.

#### **J. International research visits during BASTION project**

Twinning at Karolinska Institute 03 November 2013 –22 December 2013, 16 February 2014 –12 April 2014, 30 May 2014 –29 June 2014

#### **K. Current research interests**

Recent studies indicate that circulating tumor cells (CTC), released by primary tumours into blood, represent an independent prognostic factor for patient survival. These are biomarkers which are increasingly being used in clinical trials.

A critical concept that has emerged to be relevant to CTCs is the epithelial to mesenchymal transition (EMT), which enables epithelial cells to lose their apical–basal polarity, detach from neighboring cells, acquire a fibroblast-like morphology and invade through the surrounding stroma. During this process, tumor cells lose expression of specific epithelial markers including E-cadherin and cytokeratin, gain expression of mesenchymal cytoskeletal and adhesion proteins such as vimentin, CD44 and N-cadherin, and upregulate kinases and growth factors including c-MET, TGF- $\beta$ , Wnt. Our preliminary results show that the number and the structure of the vessels in a tumor mass is a better predictor of tumor dissemination and CTC number than tumor size. The project aims to identify the role of vascular factors in interrelationships between CTC with metastasis. Moreover, she works on modulation of angiogenesis and inhibition of tumor growth in colon cancer model through  $\alpha 5\beta 1$ -integrin/c-Met/FAK/Src-dependent signaling pathway to identify possible molecular players that are involved in this process. Moreover, the important part of research work of Oksana is the study of anti-viral treatment of cytomegalovirus (CMV) -infected tumors (colon cancer, brain tumors) aimed to better understand the CMV role in cancer.



#### **L. Envisioned career paths in BASTION project**

Collaboration of Dr. Piotr Religa with Dr. Oksana Kovtonyuk ends with the finalization of the BASTION project. Oksana is going to look for next postdoctoral position.



## X. Marzena Lazarczyk (TEAM of Zbigniew Gaciong)



| DATE (YEARS) | DEGREE/EXPERIENCE | PLACE   | SUPERVISOR             |
|--------------|-------------------|---|------------------------|
| 2005         | MSc.              | University of Warsaw, Faculty of Biology, Warsaw, Poland                              | Prof. Mieczysław Kuras |
| 2009         | PhD               | Mossakowski Medical Research Centre; Polish Academy of Sciences (PAS), Warsaw, Poland | Prof. Ewa Matyja       |

### A. Biosketch (provided by Marzena Lazarczyk)

My research was previously focused on plant compounds and their potential role in anticancer therapy. Since 2009 I still continued my cancer research within PhD. I studied potential anti-proliferative properties of tachykinin and opioid peptides analogues (substance P/NK1 receptor antagonists, opioids agonists and hybrid peptides consisted of tachykinin-like and opioid sequences), as well as platinum (II) peptide complexes on human and rat glioma cell lines. The achievements of mentioned research activities were partially included in the Final Activity Report for Normolife project Specific targeted research or innovation project (Development of new therapeutic substances and strategies for treatment of pain in patients with advanced stages of cancer within Sixth Framework Programme Life Sciences, Genomics And Biotechnology For Health Liefescihealth-6). In the meantime (2010-2013) I gained comprehensive knowledge and practical experience in clinical trials field cooperating with clinical research organizations. As a project coordinator and medical writer I had an excellent opportunity to further develop my scientific background in pre-clinical research getting familiar with clinical studies. I dealt with clinical data management, prepared numerous and completed clinical trial documentation for registration, including clinical study protocols submitted to European Medicines Agency and clinical study reports submitted to U.S. Food and Drug Administration.

### B. Grant applications filled during BASTION project

1. Preparation and coordination of project submitted for competitions: PBS III NCBiR and: Symfonia II, Symfonia III, Tango I, Sonata, Opus NCN
2. Sonata 9, NCN "Role of CCL9 chemokine in colon cancer progression" – principal investigator, failed

### C. Participation in courses/trainings/workshops

1. Attendance at workshop "Application of flow cytometry in molecular oncology" 15 – 16 October 2014

### D. Collaboration with other research teams started during BASTION project

Collaboration with CePT IMDiK and Institute of Genetics and Animal Breeding





#### **E. International research visits during BASTION project**

Twinning in Centre for Molecular Medicine in Stockholm (23 October – 21 December 2014, 30 June – 30 July 2015)

#### **F. Current research interests**

Since 2013 Marzena continues her cancer research trying to combine her experience in pharmaceutical industry /clinical trial companies and business area with scientific activity to commercialize the research results. We are investigating the role of distinct factors affecting tumour progression i.e. cytomegalovirus (CMV) and chemokines- mediated mechanisms of cancer metastasis. It is known that CMV contributes to increased motility of tumour cells and facilitate their migration. CMV virus presence has been detected in numerous cases of distinct human cancer types, including breast cancer, colon cancer, sarcomas, glioblastoma, medulloblastoma and neuroblastoma. We are intending to test selected compounds against CMV virus in animal model of human malignances to develop therapeutic strategies towards metastatic diseases.

Second line of research she is involved in relies on attempts to find explanation of the role of chemokines and their receptors in cancer invasiveness and migration. It has been demonstrated that chemokines can control organ predilection of metastasis. We assume that detailed insight into chemokines signaling may provide additional information on mechanisms of cancer metastasis. Upcoming studies of Marzena will focus on CXCL9-related paracrine and autocrine mechanisms by which tumors retain their own ability to spread.

#### **G. Envisioned career paths in BASTION project**

Dr Marzena Łazarczyk will be employed in a new Laboratory for Experimental Angiology, and her salary will be funded from NCN grants. She will try to build her position of an independent researcher applying for funding to Polish National Science Centre.



## XI. TEAM of Rafal Ploski

### Lech Trzeciak



| DATE (YEARS) | DEGREE/EXPERIENCE | PLACE   | SUPERVISOR         |
|--------------|-------------------|---|--------------------|
| 1984-1990    | M.Sc.             | Medical University of Warsaw, First Faculty of Medicine               | - (none)           |
| 1991-1994    | Assistantship     | Medical Centre for Postgraduate Education, Warsaw                     | prof. J. Ostrowski |
| 1994-1995    | Fellowship        | University of Washington, Seattle                                     | Prof. K. Bomsztyk  |
| 1995-1999    | PhD               | Medical Centre for Postgraduate Education, Warsaw                     | prof. J. Ostrowski |
| 2000-2001    | Postdoc           | M. Nencki Institute of Experimental Biology, Warsaw                   | prof. M. Żylicz    |
| 2001-2004    | Postdoc           | International Institute of Molecular and Cell Biology, Warsaw, Poland | prof. M. Żylicz    |

#### A. Biosketch (provided by Lech Trzeciak)

I began my scientific career in 1987 as a medical student, joining the research team of prof. Ostrowski at the Department of Gastroenterology, Medical Centre for Postgraduate Education in Warsaw. I started from biochemical research (such as HPLC analysis of various components or measuring activity of enzymes, looking for biomarkers of certain diseases, including neoplasms). Soon I made a transition to molecular biology, meanwhile graduating from Medical Faculty in 1990. My primary interest was cancer biology (molecular causes of transformation and metastasis) with particular emphasis on protein phosphorylation (culminated in 1.5 year fellowship in the molecular lab of prof. Bomsztyk in University of Washington, Seattle, USA, for studying phosphorylation of RNA-binding proteins, 1994-1995).

Initially I was involved in studies on the action of growth factors (via kinase receptors) on cancer cells, but it was then already clear that the reason for increased activity of certain receptors in neoplasms was DNA



mutation. In 1994 I have successfully applied for a grant for investigating p53 mutations and expression in colorectal cancer, and in 1999 completed my PhD thesis based on the results from this project that involved sequencing on first generation semi-automated DNA sequencers from ABI. In course of these studies we also looked for the contribution of DNA methylation to cancer development. I was hoping to extend these studies to cover more genes, reasoning that cancer development relies on interplay of at least several altered pathways, but we soon realized that semi-automated sequencing, although being a great improvement over radiolabeled manual method, wasn't really powered enough for a large scale cancer DNA study, considering the size and exon-intron composition of several just-cloned cancer-related genes.

In 2000 I got another grant, for cloning and studying a then-novel human protein kinase (discovered by myself through a PCR-based screen of a cancer cell transcriptome) and moved with this project to the International Institute for Molecular and Cellular Biology in Warsaw under supervision of prof. Maciej Żylicz. The study got an unexpected aid from HUGO project: an accelerated publication of a draft of human genome essentially produced a complete sequence of the gene we were attempting to clone. We followed with the studies on protein function, finding a plausible activation mechanism for the kinase and looking for its interacting partners. However, the kinase appeared to be unlikely involved in carcinogenesis. Meanwhile, we were again reminded that to study molecular biology of cancer one needs sufficient resources to cover multiple interaction networks at once.

Since then I had spent several years working in science/education, first for Polish edition of Scientific American and then for two medical book publishers, closely following the advances in the field. The development of next generation sequencing turned former impossibility into nearly a routine. Therefore I took this opportunity, quit my recent job and successfully applied for a position within BASTION project.

#### **B. Grant applications submitted during BASTION project**

1. Searching for genetic factors linked to early mortality in Polish population by means of whole-exome sequencing – co-author (primary investigator: prof. R. Płoski) – accepted for financing in May 2014
2. Pathogenesis of ovarian cancer and predicting its drug sensitivity from its molecular profile – co-author/consultant – application failed

#### **C. Participation in grants during BASTION project**

1. Searching for genetic factors linked to early mortality in Polish population by means of whole-exome sequencing – role: analyzing NGS data

#### **D. Participation in the conferences during BASTION project**

1. "Science and Business" - main topics: 1. management of intellectual property rights 2. applying for grants to governmental agencies, Warsaw, Poland, 22 November 2013
2. Workshop "Genome-wide methods in cancer genetics" Warsaw, Poland, 28 November 2014
3. "Translational Research in Oncology in New Member State Economies (TRON)", Warsaw, Poland, 21-22 May 2015.

#### **E. Poster presentation at the conferences**

1. "c.449-1G>T mutation of TMC8 gene - an unreported frequent cause of epidermodysplasia verruciformis in Polish population – a case study and molecular basis" TRON, Warsaw 2015.05.21-22.
2. "Functional analysis of SMAD4 mutants in an in vitro system reveals upregulation of SMAD2, SMAD3 and SMAD4 by Myhre syndrome-associated variants" TRON, Warsaw 21-22 May 2015.

#### **F. Participation in courses/trainings/workshops**



1. NGS workshop on SureSelect vs Haloplex; SureDesign, SureCall (dr Andreas Polten, Agilent Technologies; 29 November 2013)
2. training/tests of NimbleGen SeqCap EZ (03-07 February 2014)
3. training on the usage of Fluidigm system (13-14 March 2014)
4. Leica Microscopy Workshop in Cologne on DLS, GSD and STED (16-17 June 2015)
5. completion of postgraduate studies on statistics in biomedical research (1 year, 2014/2015) at Medical University Łódź (organized in cooperation with StatSoft)

#### **G. Students supervision**

Temporary supervising Joanna Sałkowska-Wanat, PhD student from Department of Dermatology

#### **H. Collaboration with other research teams started during BASTION project**

- 1) Twinning with the Institute of Virology (University of Cologne) / German National Reference Center for Papillomaviruses (head: prof. Herbert Pfister)
- 2) Collaboration with Department of Dermatology, WUM (also in conjunction with 1)
- 3) Collaboration with R. Zagożdżon team (BASTION member) on a preliminary study for possible grant application on SMAD4 function

#### **I. International research visits during BASTION project**

Twinning to Institute of Virology, University of Cologne, 23 April 2014 – 11 May 2014, 03 June 2015 – 17 July 2015

#### **J. Current research interests**

Research interests of Lech Trzeciak revolve around the role of genes in the development and outcome of cancer (incl. cancer therapy). The advent of Next Generation Sequencing allows now for a comprehensive (and relatively inexpensive) study of individual cancer exomes/genomes, methylomes and transcriptomes and compare these to the corresponding normal tissue of the same individual or its relatives, if appropriate. This approach may be used in several ways.

First, genetic predispositions towards cancer may be elucidated. Another possible way of using NGS data is to correlate an individual tumor mutation profile to the clinical parameters, most importantly to the susceptibility to treatment with conventional chemotherapeutics as well as modern targeted therapies (such as low-molecular weight kinase inhibitors etc). Lech is also interested in another use of NGS information, namely correlating mutational profile to immunologic parameters of a neoplasm.

#### **K. Envisioned career paths in BASTION project**

Regrettably, it is now impossible to extend current employment of Lech Trzeciak at Department of Medical Genetics. He is looking for a new job in the field of cancer studies, applying for a position in the newly opening laboratory studying hedgehog signaling and its role in medulloblastoma. This pathway is also of paramount importance for basal cell skin carcinomas, a field closely related to HPV-induced squamous cell skin carcinoma, so common in cases of erythrodermia verruciformis that he studied in twinning cooperation with Institute of Virology Cologne. There are also strong links between Hedgehog and TGF $\beta$  signaling pathways, what would let Lech Trzeciak capitalize on his work with SMAD4 mutants during BASTION. He also plans to remain involved in NGS data analysis in the Department of Medical Genetics, hoping to continue developing skills and to introduce data analysis techniques he has learned last year in course of biostatistics studies.



### 3. Summary of research activity of recruited postdocs

BASTION project has fully used its opportunity to recruit nine top-level qualified researchers with high ability to increase research potential in basic and translational oncology at Medical University of Warsaw. Since two postdocs had to quit, we managed to hire for replacement two another highly qualified researchers. The technological expertise and scientific background of all eleven recruits fit BASTION effort to strengthen the existing areas of excellence in oncology research. Moreover, each individual used the opportunity to bring in know-how and experience in translational oncology work and helped to bridge the gaps and create links among research groups working at MUW. All leaders have succeeded in recruiting extremely diligent and hardworking postdocs showing a great enthusiasm for their work in the field of experimental oncology. All newly employed researchers contributed to the great success of BASTION project.

In summary, postdocs recruited to nine research groups in BASTION project are authors and co-authors of 32 publications, they managed to secure funding for their research and get 9 grants (7 grants as Principal Investigators, 2 as supervisors), they were awarded with 7 different awards and are authors of 4 patent applications).

#### **Working space:**

All recruited researchers were provided with research and office space by leaders of research groups already existing at MUW. Since four research groups in BASTION project were located at the Department of Immunology the option to increase the research and office space was necessary and inevitable. We managed to redesign and renovate one room to provide new researchers with sufficient working space. Team of Pawel Wlodarski during BASTION project moved to a new laboratory located in a newly built CePT (Centre for Preclinical Research and Technology) building.

#### **Status of recruited researchers:**

Number of faculty positions at Medical University of Warsaw is regulated by a quota of teaching hours (pensum). Thus, according to the recruitment policy of Medical University of Warsaw recruited researchers were employed at the university as the experienced research specialists. They were entitled to all benefits of governmental employees.

#### **Research funding:**

BASTION project did not directly provide research support for newly employed post docs. All recruited researchers were eligible for applying for national funding from National Science Centre (NCN), The National Centre for Research and Development (NCBiR), The Foundation for Polish Science (FNP) and Ministry of Science and Higher Education. All researchers made attempts to get funds for their research and prolong their employment.

For three researchers (Joanna Drzewinska-Chanko, Magdalena Banach-Orlowska and Oksana Kovtonyuk) it was not possible to extend employment at MUW after completion of the BASTION project.

Two researchers (Iwona Solarska and Lech Trzeciak), due to shortage of money, with the finalization of BASTION project end their cooperation with team leaders. However, they declare their interest in BASTION projects and willingness to cooperate with BASTION leaders.

Three postdocs (Anna Wojcicka, Beata Pyrzynska and Malgorzata Czystowska-Kuzmicz) have managed to secure funding for research and their salaries as principal investigators:

1. Anna Wojcicka until 2017 (National Centre for Research and Development Lider Grant: The use of next-generation sequencing for elucidation of a sensitive and specific molecular panel for diagnostics of thyroid cancers and Ministry of Science and Higher Education Iuventus Plus Grant: Evaluation of the possibility of using microRNA inhibitors as adjuvant therapy for thyroid cancer )



2. Beata Pyrzyńska until 2017 (grant OPUS, NCN; 2013/11/B/NZ5/03240) Influence of AKT signaling pathway on CD20 expression and antitumor activity of therapeutic monoclonal antibodies)
3. Malgorzata Czystowska-Kuzmicz until 2017 (grant OPUS, NCN Elucidation of the role of tumor-derived and exosomal arginases in avoiding immune responses by ovarian cancer)

Moreover, team leaders will support the employment of three other researchers (Malgorzata Firczuk, Agnieszka Pollak and Marzena Lazarczyk) with their grant funding:

1. Malgorzata Firczuk as a postdoc until 2016 in team of Radoslaw Zagodzón (OPUS, NCN, Evaluation of peroxiredoxins 1 and 2 along with the thioredoxin-thioredoxin reductase system as new therapeutic targets in B cell lymphomas)
2. Agnieszka Pollak as a postdoc until 2017 in team of Pawel Wlodarski (OPUS, NCN, Exome-wide search for somatic mutations in pathogenesis of endometriosis)
3. Marzena Lazarczyk as a postdoc until 2018 in team of Zbigniew Gaciong (OPUS, NCN, Red cell heterogeneity as a risk factor for thrombotic complications)

### Corresponding estimated/\* budget

| PERSONNEL, TRAVEL, OTHER MAJOR DIRECT COST ITEMS FOR BENEFICIARY "1" FOR M19-M36 |                              |              |   |  |
|--|------------------------------|--------------|---|--|
| WP No  | Item description             | Amount [EUR] | Explanations  |  |
| WP3<br>Task 3 &<br>3.1   | Personnel costs              | 571,383.12   | Fees of the WP3 leader, Co-leader, recruitment committee members (4,25 PM); salaries of 9+2 newly hired (eleven) Postdocs - experienced researchers (160,55 PM) |  |
|  | Travel                       | 0,00         |   |  |
|  | Remaining direct costs       |              | 1,053.16  | Recruitment press announcements            |
|  |                              |              | 807.80  | Projector for postdocs meetings            |
|  |                              |              | 151.93  | Refreshments for BASTION postdocs meetings |
|  | <b>TOTAL DIRECT WP3 COST</b> |              | <b>573,396.01</b>   |  |

*/\* - exact costs for M19-M36 will be presented in the 11<sup>th</sup> Period Report and Form C (October 2015)*

Prof. Jakub Golab  
BASTION Project Coordinator  
Dr Magdalena Winiarska  
WP3 Leader

*Warsaw, August 2015*



## Attachment 1

### Announcement for postdoc position in the team of Tomasz Stoklosa



**Medical University of Warsaw, Poland**  
**REGPOT-2012-2013-1 Program EU FP7**

**Project BASTION (From Basic to Translational Research in Oncology)**  
is looking for a

**POSTDOCTORAL FELLOW (temporary position-substitution)**

**in the project aimed at looking for therapeutic targets and mechanisms of drug resistance in hematological and solid tumors - beginning in the third quarter of 2014:**

**Ref. no: APK2/1210-19/2014**

#### Requirements:

- PhD degree (or equivalent) in medical sciences, genetics, biotechnology or molecular biology,
- Extensive experience in the field of molecular biology, genetics, experimental oncology (minimum two-year post-doctoral employment, preferably previous post doctoral positions different from the PhD awarding institutions),
- Outstanding publication record,
- Strong background in tumor biology and gene-expression analysis, including qPCR;
- Experience in cell culture and cytotoxicity assays;
- Experience in working with biological or medical databases and bioinformatics tools;
- Ability to travel for short term assignments and work with international partners;
- Experience in dealing with clinical data will be an additional asset
- Proficiency in English

#### Required documents and declarations:

- CV
- Letter-of-intent
- Reference letter(s)
- Copy of degree diploma(s),
- Copy of certificate(s) of employment,
- Contact information, including e-mail address and phone number
- Declaration about authorization for personal data processing: „I hereby authorize you to process my personal data included in my job application for the needs of the recruitment process (in accordance with the Personal Data Protection Act, Journal of Laws of 2002, no 101, item 926 as amended)
- The candidates may include additional information or copies of documents/certificates in support of the application.

#### Selection criteria

A detailed analysis of the received applications will be based on the following evaluation criteria:

- List of publications: 0-50 points
- Professional experience: 0-25 points
- Previous international experience: 0-10 points
- Adequacy of the prepared letter-of-intent with the target-project: 0-5 points
- Reference(s): 0-5 points
- Certificates of extra qualifications that may be of some value for the execution of the project: 0-5 points

**Position is offered for 6 months with the possibility of extension to 12 months.**

Applications should be submitted by 3p.m. (Warsaw time) on 15<sup>th</sup> July 2014 to

[magdalena.winiarska@wum.edu.pl](mailto:magdalena.winiarska@wum.edu.pl) with a note in the e-mail subject:

**“Competition for the position of Postdoctoral fellow ref. no APK2/1210-19/2014 in “BASTION” project**

The admission procedure will be carried out in two steps. First, the applicants are requested to submit application documents. Short listing will be carried out within 3 days after the closing date. Applications will be assessed against person specification criteria and 3-4 applicants will be invited for interview. During the interview candidates will be scored with regard to communication skills, teamwork and project competency. Successful candidates will be offered a position within 2 days after the interview date.

For more information on the project visit our website at <http://bastion.wum.edu.pl/>

Please be advised that only selected candidates will be contacted, and sent documents will not be returned.



## Attachment 2

### Candidates for Tomasz Stoklosa team:

| Lp           | Name                     | Formal requirements       |               | Points |
|--------------|--------------------------|---------------------------|---------------|--------|
| 1            | Olena Bakhuryńska        | Ph.D. diploma             | +             | -      |
|              |                          | CV                        | +             | -      |
|              |                          | Motivation letter         | +             | -      |
|              |                          | reference letters         | -             | -      |
|              |                          | Publications              | +             | -      |
|              |                          | Experience in the area    | +             | -      |
|              |                          | International experience  | +             | -      |
|              |                          | Additional qualifications | +             | -      |
| <b>TOTAL</b> |                          |                           | <b>failed</b> |        |
| 2            | Piotr Banski             | Ph.D. diploma             | -             | -      |
|              |                          | CV                        | +             | -      |
|              |                          | Motivation letter         | +             | -      |
|              |                          | reference letters         | -             | -      |
|              |                          | Publications              | +             | -      |
|              |                          | Experience in the area    | +             | -      |
|              |                          | International experience  | +             | -      |
|              |                          | Additional qualifications | +             | -      |
| <b>TOTAL</b> |                          |                           | <b>failed</b> |        |
| 3            | Thomas Fricke            | Ph.D. diploma             | +             | +      |
|              |                          | CV                        | +             | +      |
|              |                          | Motivation letter         | +             | 5      |
|              |                          | reference letters         | +             | 5      |
|              |                          | Publications              | +             | 50     |
|              |                          | Experience in the area    | +             | 5      |
|              |                          | International experience  | +             | 10     |
|              |                          | Additional qualifications | +             | 10     |
| <b>TOTAL</b> |                          |                           | <b>75</b>     |        |
| 4            | Paulina Gapska           | Ph.D. diploma             | +             | +      |
|              |                          | CV                        | +             | +      |
|              |                          | Motivation letter         | +             | 5      |
|              |                          | reference letters         | +             | 5      |
|              |                          | Publications              | +             | 10     |
|              |                          | Experience in the area    | +             | 5      |
|              |                          | International experience  | -             | 5      |
|              |                          | Additional qualifications | +             | 5      |
| <b>TOTAL</b> |                          |                           | <b>35</b>     |        |
| 5            | Iwona Solarska           | Ph.D. diploma             | +             | +      |
|              |                          | CV                        | +             | +      |
|              |                          | Motivation letter         | +             | 5      |
|              |                          | reference letters         | +             | 5      |
|              |                          | Publications              | +             | 40     |
|              |                          | Experience in the area    | +             | 20     |
|              |                          | International experience  | +             | 5      |
|              |                          | Additional qualifications | +             | 10     |
| <b>TOTAL</b> |                          |                           | <b>85</b>     |        |
| 6            | Katarzyna Solarska-Sciuk | Ph.D. diploma             | +             | +      |





|    |                       |                           |    |               |
|----|-----------------------|---------------------------|----|---------------|
|    |                       | CV                        | +  | +             |
|    |                       | Motivation letter         | +  | 5             |
|    |                       | reference letters         | +  | 5             |
|    |                       | Publications              | +- | 25            |
|    |                       | Experience in the area    | +  | 5             |
|    |                       | International experience  | +  | 5             |
|    |                       | Additional qualifications | +  | 5             |
|    |                       | <b>TOTAL</b>              |    | <b>50</b>     |
| 7  | Przemek Swiecki       | Ph.D. diploma             | -  | -             |
|    |                       | CV                        | +  | -             |
|    |                       | Motivation letter         | -  | -             |
|    |                       | reference letters         | -  | -             |
|    |                       | Publications              | -  | -             |
|    |                       | Experience in the area    | -  | -             |
|    |                       | International experience  | +  | -             |
|    |                       | Additional qualifications | +  | -             |
|    |                       | <b>TOTAL</b>              |    | <b>failed</b> |
| 8  | Abhishek Narain Singh | Ph.D. diploma             | -  | -             |
|    |                       | CV                        | +  | -             |
|    |                       | Motivation letter         | -  | -             |
|    |                       | reference letters         | -  | -             |
|    |                       | Publications              | -  | -             |
|    |                       | Experience in the area    | +  | -             |
|    |                       | International experience  | +  | -             |
|    |                       | Additional qualifications | +  | -             |
|    |                       | <b>TOTAL</b>              |    | <b>failed</b> |
| 9  | Juan Alfonso Redondo  | Ph.D. diploma             | +  | +             |
|    |                       | CV                        | +  | +             |
|    |                       | Motivation letter         | +  | 5             |
|    |                       | reference letters         | -  | -             |
|    |                       | Publications              | +  | 10            |
|    |                       | Experience in the area    | +  | 5             |
|    |                       | International experience  | +  | 10            |
|    |                       | Additional qualifications | +  | 5             |
|    |                       | <b>TOTAL</b>              |    | <b>35</b>     |
| 10 | María Ramírez Arroyo  | Ph.D. diploma             | -  | -             |
|    |                       | CV                        | +  | -             |
|    |                       | Motivation letter         | -  | -             |
|    |                       | reference letters         | -  | -             |
|    |                       | Publications              | -  | -             |
|    |                       | Experience in the area    | +  | -             |
|    |                       | International experience  | +  | -             |
|    |                       | Additional qualifications | +  | -             |
|    |                       | <b>TOTAL</b>              |    | <b>failed</b> |
| 11 | Tomasz Pelczar        | Ph.D. diploma             | -  | -             |
|    |                       | CV                        | +  | -             |
|    |                       | Motivation letter         | -  | -             |
|    |                       | reference letters         | -  | -             |
|    |                       | Publications              | -  | -             |
|    |                       | Experience in the area    | -  | -             |
|    |                       | International experience  | -  | -             |
|    |                       | Additional qualifications | -  | -             |



|    |             |                           |   |               |
|----|-------------|---------------------------|---|---------------|
|    |             | <b>TOTAL</b>              |   | <b>failed</b> |
| 12 | Sheik Asraf | Ph.D. diploma             | + | -             |
|    |             | CV                        | + | -             |
|    |             | Motivation letter         | - | -             |
|    |             | reference letters         | - | -             |
|    |             | Publications              | + | -             |
|    |             | Experience in the area    | + | -             |
|    |             | International experience  | + | -             |
|    |             | Additional qualifications | + | -             |
|    |             | <b>TOTAL</b>              |   | <b>failed</b> |



### Attachment 3

### Announcement for postdoc position in the team of Pawel Wlodarski



Medical University of Warsaw, Poland  
REGPOT-2012-2013-1 Program EU FP7

Project BASTION (From Basic to Translational Research in Oncology)  
is looking for a  
**POSTDOCTORAL FELLOW**

in the project on the role of somatic genetic changes in tissue homeostasis, using  
NGS techniques on analyses of DNA (including DNA methylation) and RNA (including  
microRNA)

The project begins in May 2015.  
Ref. no: APK2/1210-13/2015

#### Requirements:

- PhD degree (or equivalent) in molecular biology, biochemistry, biology, chemistry, physics or medical sciences, and extensive experience in the field
- outstanding publication record,
- be experienced in the analysis of NGS data
- be experienced in preparation of libraries for NGS (demonstrated via publications, references of the candidate's thesis tutor, previous post-doctoral positions different from the PhD awarding institutions),
- be proficient in English.

#### Required documents and declarations:

- CV
- Letter-of-intent
- Reference letter(s)
- Copy of degree diploma(s),
- Copy of certificate(s) of employment,
- Contact information, including e-mail address and phone number
- Declaration about authorization for personal data processing: „I hereby authorize you to process my personal data included in my job application for the needs of the recruitment process (in accordance with the Personal Data Protection Act, Journal of Laws of 2002, no 101, item 926 as amended)
- The candidates may include additional information or copies of documents/certificates in support of the application.

#### Selection criteria

A detailed analysis of the received applications will be based on the following evaluation criteria:

- List of publications: 0-50 points
- Professional experience: 0-25 points
- Previous international experience: 0-10 points
- Adequacy of the prepared letter-of-intent with the target-project: 0-5 points
- Reference(s): 0-5 points
- Certificates of extra qualifications that may be of some value for the execution of the project: 0-5 points

#### Position is offered for 4 months.

Applications should be submitted by 3p.m. (Warsaw time) on 29<sup>th</sup> April 2015 to  
[pawel.wlodarski@wum.edu.pl](mailto:pawel.wlodarski@wum.edu.pl) with a note in the e-mail subject:

**"Competition for the position of Postdoctoral fellow ref. no APK2/1210-.../2015 in "BASTION" project**

The admission procedure will be carried out in two steps. First, the applicants are requested to submit application documents. Short listing will be carried out within 3 days after the closing date. Applications will be assessed against person specification criteria and 3-4 applicants will be invited for interview. During the interview candidates will be scored with regard to communication skills, teamwork and project competency. Successful candidates will be offered a position within 2 days after the interview date.

For more information on the project visit our website at <http://bastion.wum.edu.pl>  
Please be advised that only selected candidates will be contacted, and sent documents will not be returned.



#### Attachment 4

Candidates for Pawel Wlodarski team:

| Lp | Name             | Formal requirements       |   | Points |
|----|------------------|---------------------------|---|--------|
| 1  | Agnieszka Pollak | Ph.D. diploma             | + | -      |
|    |                  | CV                        | + | -      |
|    |                  | Motivation letter         | + | 5      |
|    |                  | 2 reference letters       | + | 5      |
|    |                  | Publications              | + | 35     |
|    |                  | Experience in the area    | + | 20     |
|    |                  | International experience  | + | 0      |
|    |                  | Additional qualifications | + | 10     |
|    |                  | <b>TOTAL</b>              |   |        |