

## Prof. dr hab. Antoni Więdołcha – informacja biograficzna

**Name: Antoni Wiedlocha born:** 12 June 1959, Opole, Poland

**Home address:** Oppsaltoppen 16C, N-0687 Oslo, Norway

**Work address:** Centre for Cancer Biomedicine, Department of Biochemistry, Institute for Cancer Research, The Norwegian Radiumhospital, Oslo University Hospital, Montebello, N-0379 Oslo, Norway.

### EDUCATION:

**M. Sc., 1983** Biology, University of Wrocław, Wrocław, Poland.

**November 1986** UICC – Yamagiva-Yoshida Fellowship, Institute for Cancer Research, Department of Biochemistry, Oslo, Norway.

**January–April 1988** Norwegian and Polish Government Fellowship, Institute for Cancer Research, Department of Biochemistry, Oslo, Norway.

**Ph.D., 1989** Tumor immunotherapy, Ludwik Hirszfild Institute of Immunology and Experimental Therapy, Polish Academy of Sciences, Wrocław, Poland.

**Thesis entitle:** *The use of anti-L1210V leukemic monoclonal antibody and its ricin A-chain immunotoxin in experimental immunotherapy.*

**Habilitation, 1997** Mechanisms of fibroblast growth factor signaling, Ludwik Hirszfild Institute of Immunology and Experimental Therapy, Polish Academy of Sciences, Wrocław, Poland.

**Thesis entitle:** *Diphtheria toxin as a molecular tool in the study of acidic growth factor intracrine signaling.*

### Courses:

**Spring 1995** Growth factors and Oncogenes, EMBL/EMBO, Heidelberg, Germany

**2006–2007** Harvard Business School Publishing, research leadership course (*Forskerledelse*) at Institute for Cancer Research, The Norwegian Radium Hospital, Oslo, Norway.

### POSITIONS HELD:

**1984–1990** **Ph.D. student/assistant** at Department of Tumor Immunology, Ludwik Hirszfild Institute of Immunology and Experimental Therapy, Polish Academy of Sciences, Wrocław, Poland.

**1991–1994** **Postdoctoral fellow** of Norwegian Cancer Society, Department of Biochemistry, Institute of Cancer Research, Oslo, Norway.

**1995–1998** **Research fellow** at Department of Biochemistry, Institute for Cancer Research, The Norwegian Radium Hospital, Oslo, Norway.

**Since 1998** **Senior scientist** (*professor competence*) at Department of Biochemistry, Institute for Cancer Research, The Norwegian Radium Hospital, Oslo, Norway.

**2004–2009** **Project leader** at Department of Biochemistry, Institute for Cancer Research, The Norwegian Radium Hospital, Centre for Cancer Biomedicine, Oslo, Norway.

**Current position:** **Group leader** at Department of Biochemistry, Institute for Cancer Research, The Norwegian Radium Hospital, Centre for Cancer Biomedicine, University of Oslo, Oslo, Norway.

#### SCIENTIFIC ACTIVITY:

##### Publication and meetings:

- Published 68 original articles in peer-review international journals
- Published 14 reviews
- Invited speaker at 14 international conferences and 4 courses
- Invited speaker at 2 Gordon Research Conference (2000 and 2008)

##### Scientific experience:

- Tumor immunobiology
- Experimental anti-cancer therapy
- Molecular and cell biology
- Biochemistry

##### Current scientific interest:

- Molecular determinants of malignant phenotype
- Network of cell signaling
- Endocytosis and intracellular sorting
- Nuclear transport – nuclear signaling
- Targeted anti-cancer therapy

#### TEACHING AND SUPERVISION:

**Supervision:** supervisor of M.Sc. (4), Ph.D. degrees (2), co-supervisor Ph.D. degrees (4) and postdocs (4) .

**Teaching experience:** Since 1996 teaching at course MBV4240 – Biochemical mechanism in intracellular transport, at University of Oslo.

#### AWARDS AND HONORS:

1988 – Award of Polish Biochemical Society

1989 – Award of President of Polish Academy of Sciences

2002 – Dr. Ragnar Mørks Legat Price

## LIST OF PUBLICATION 2004-2015

### Original papers

1. Zakrzewska, M., Krowarsch, D., **Wiedlocha, A.**, and Otlewski, J. (2004). Design of fully active FGF-1 variants with increased stability. *Protein Eng. Des. Sel* 17, 603–611.
2. Malecki, J., Wesche, J., Skjerpen, C.S., **Wiedlocha, A.**, and Olsnes, S. (2004). Translocation of FGF-1 and FGF-2 across vesicular membranes occurs during G1-phase by a common mechanism. *Mol. Biol. Cell* 15, 801–814.
3. Zakrzewska, M., Krowarsch, D., **Wiedlocha, A.**, Olsnes, S., and Otlewski, J. (2005). Highly stable mutants of human fibroblast growth factor-1 exhibit prolonged biological action. *J. Mol. Biol.* 352, 860–875.
4. **Wiedlocha, A.**, Nilsen, T., Wesche, J., Sorensen, V., Malecki, J., Marcinkowska, E., and Olsnes, S. (2005). Phosphorylation-regulated nucleocytoplasmic trafficking of internalized fibroblast growth factor-1. *Mol. Biol. Cell* 16, 794–810.
5. Wesche, J., Malecki, J., **Wiedlocha, A.**, Ehsani, M., Marcinkowska, E., Nilsen, T., and Olsnes, S. (2005). Two nuclear localization signals required for transport from the cytosol to the nucleus of externally added FGF-1 translocated into cells. *Biochemistry*, 44, 6071–6080.
6. Sorensen, V., **Wiedlocha, A.**, Haugsten, E.M., Khnykin, D., Wesche, J., and Olsnes, S. (2006). Different abilities of the four FGFRs to mediate FGF-1 translocation are linked to differences in the receptor C-terminal tail. *J. Cell Sci.* 119, 4332–4341.
7. Wesche, J., Malecki, J., **Wiedlocha, A.**, Skjerpen, C.S., Claus, P., and Olsnes, S. (2006). FGF-1 and FGF-2 require the cytosolic chaperone Hsp90 for translocation into the cytosol and the cell nucleus. *J. Biol. Chem.* 281, 11405–11412.
8. Marcinkowska, E., Superat, K., and **Wiedlocha, A.** (2006). FGF-1 as a possible carrier for targeted drug delivery. *Oncol. Res.* 16, 27–34.
9. Zakrzewska, M., Krowarsch, D., **Wiedlocha, A.**, Olsnes, S., and Otlewski, J. (2006). Structural requirements of FGF-1 for receptor binding and translocation into cells. *Biochemistry* 45, 15338–15348.
10. Nilsen, T., Rosendal, K.R., Sorensen, V., Wesche, J., Olsnes, S., and **Wiedlocha, A.** (2007). A nuclear export sequence located on a beta-strand in fibroblast growth factor-1. *J. Biol. Chem.* 282, 26245–26256.
11. Zhen, Y., Sorensen, V., Jin, Y., Suo, Z., and **Wiedlocha, A.** (2007). Indirubin-3'-monoxime inhibits autophosphorylation of FGFR1 and stimulates ERK1/2 activity via p38 MAPK. *Oncogene* 26, 6372–6385.
12. Sorensen, V., Zhen, Y., Zakrzewska, M., Haugsten, E.M., Walchli, S., Nilsen, T., Olsnes, S., and **Wiedlocha, A.** (2008). Phosphorylation of fibroblast growth factor (FGF) receptor 1 at Ser777 by p38 mitogen-activated protein kinase regulates translocation of exogenous FGF1 to the cytosol and nucleus. *Mol. Cell. Biol.* 28, 4129–4141.
13. Zakrzewska, M., **Wiedlocha, A.**, Szlachcic, A., Krowarsch, D., Otlewski, J. and Olsnes, S. (2009). Increased protein stability of FGF1 mutants can compensate for lower heparin-binding. *J. Biol. Chem.*, 284, 25388–25403.
14. Zakrzewska, M., Zhen, Y., **Wiedlocha, A.**, Olsnes, S. and Wesche, J. (2009). FGF1 can carry heterologous polypeptides into cytosol and the nucleus of mammalian cells. *Biochemistry*, 48, 7209–7218.

15. Zakrzewska, M., Sørensen, V., Jin, Y., **Wiedlocha, A.** and Olsnes, S. (2011). Translocation of exogenous FGF1 into cytosol and nucleus is a periodic event independent of receptor kinase activity. *Exp. Cell Res.*, 317, 1005–1015.
16. Jin, Y., Zhen, Y., Haugsten, E.M. and **Wiedlocha, A.** (2011). The driver of malignancy KG-1a cells, *FGFR1OP2-FGFR1*, encodes an HSP90 addicted oncoprotein. *Cell. Sig.*, 23, 1758–1766.
17. Zhen Y., Sørensen V., Skjerpen C.S., Haugsten E.M., Jin Y., Walchli S., Olsnes S. and **Wiedlocha A.** (2012). Nuclear import of exogenous FGF1 requires the ER-protein LRRC59 and the importins Kpn1 and Kpn2. *Traffic*, 13, 650–664.
18. Szlechcic, A., Pala, K., Zakrzewska, M., Jakimowicz P., **Wiedlocha A.**, Otlewski J. (2012). FGF1-gold nanoparticle conjugates targeting FGFR efficiently decrease cell viability upon NIR irradiation. *Inter. J. Nanomed.*, 7, 5915–5927.
19. Zakrzewska, M., Haugsten, E.M., Nadratowska-Wesolowska, B., Oppelt, A., Hausott, B., Jin, Y., Otlewski, J., Wesche, J., **Wiedlocha, A.** (2013). ERK-mediated phosphorylation of fibroblast growth factor receptor on Ser777 inhibits signaling. *Science Sig.*, 6, 1–15.
20. Nadratowska-Wesolowska, B., Haugsten, E.M., Zakrzewska, M., Jakimowicz, P., Zhen, Y., Pajdzik, D., Wesche, J., **Wiedlocha, A.** (2014). RSK2 regulates endocytosis of FGF receptor 1 by phosphorylation on serine 789. *Oncogene*, 40, 4823–4836.
21. Sletten T., Kostas M., Bober J., Sorensen V., Olsnes J., Otlewski J., Zakrzewska M., **Wiedlocha, A.** (2014). Nucleolin regulates phosphorylation and nuclear export of fibroblast growth factor 1 (FGF1). *PLOS ONE*, 9, e90687
22. Buchtova, M., Chaloupkova, R., Zakrzewska, M., Vesela, I., Cela, P., Barathova, J., Gudernova, I., Zajickova, R., Trantirek, L., Martin, J., Kostas, M., Otlewski, J., Damborsky, J., Kozubik, A., **Wiedlocha, A.**, Krejci, P. (2015). Instability restricts signaling of multiple fibroblast growth factors. *Cell Mol Life Sci.*, Feb 18.

#### **Review papers:**

1. **Wiedlocha, A.** and Sørensen, V. (2004). Signaling, internalization, and intracellular activity of fibroblast growth factor. *Curr. Top. Microbiol. Immunol.*, 286, 45–79.
2. Sorensen, V., Nilsen, T., and **Wiedlocha, A.**, (2006). Functional diversity of FGF-2 isoforms by intracellular sorting. *BioEssays*, 28, 504–514.
3. Zakrzewska, M., Marcinkowska, E., and **Wiedlocha, A.** (2008). FGF-1: from biology through engineering to potential medical applications. *Crit Rev. Clin. Lab Sci.*, 45, 91–135.
4. Haugsten, E., **Wiedlocha, A.**, Olsnes, S., and Wesche, J. (2010). The role of fibroblast growth factor receptors (FGFR) in carcinogenesis. *Mol. Canc. Res.*, 8, 1439–1452.