



Project BASTION "From Basic to Translational Research in Oncology"

Creation and coordinating the Bioinformatics/System Modelling Group

Radoslaw Zagozdzon MD PhD



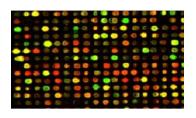
Professional background

- MD and PhD at Medical University of Warsaw – cancer immunology
- 8 years in BIDMC, Boston, USA molecular biology of cancer
- 4 years UCD, Dublin, Ireland cancer biomarkers, in vivo cancer models

Bioinfomatic support for BASTION activities

- Organizing and coordinating the research group (2 postdocs, 2 IT specialists)
- Generating computer cluster and data storage
- Support for:
 - Next-generation sequencing
 - Databases (transcriptomic, proteomic and clinical data)

Biomarker Validation



1. Biomarker discovery using various microarray datasets



2. Biomarker specificity assessment & optimisation



expression

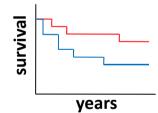
gene

Cell lines with modulated



3. Biomarker assessed on TMA/RPPA

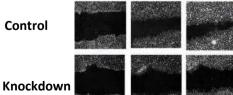




4. Biomarker validation



Control

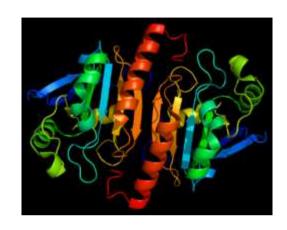


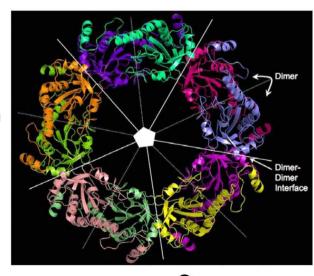
5. Utilise in vitro assays



Peroxiredoxin-1 (PRDX1)

- Hs578T cell lines
 - Parental
 - Invasive clone (-i8) decreased expression of PRDX1
- Peroxiredoxin family of antioxidant enzymes
- Multifunctional
 - Scavenger of hydrogen peroxide (dimer and decamer)
 - Chaperone of signal-transducing molecules (decamer)
 - Regulator of transcription
 - Ligand for toll-like receptor 4 (TLR4)
- Differentially expressed in prostate, lung & ovarian carcinoma.
- Contradicting results in breast cancer



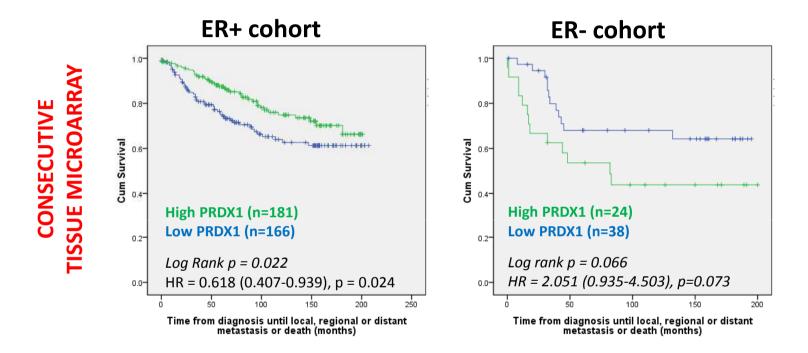


Lee W et al. J. Biol. Chem. 2007;282:22011-22022

Summary of Patient Cohorts

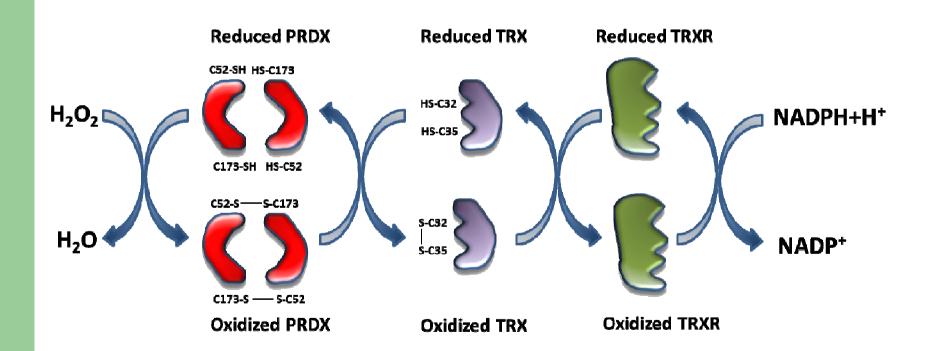
Cohort	Platform	Patient number	Protein	Clinico- pathological data	Survival	Gene Expression
Consecutive array	TMA	512	Yes (PRDX1)	Yes	Yes	No
Houston RPPA	RPPA	712	Yes (PRDX1)	Yes	Yes	No
The Cancer Genome Atlas – Breast cancer cohort	RPPA	410	~150 antibodies	Yes	Short follow-up	Yes

PRDX1 & it's prognostic impact in ER-positive patients

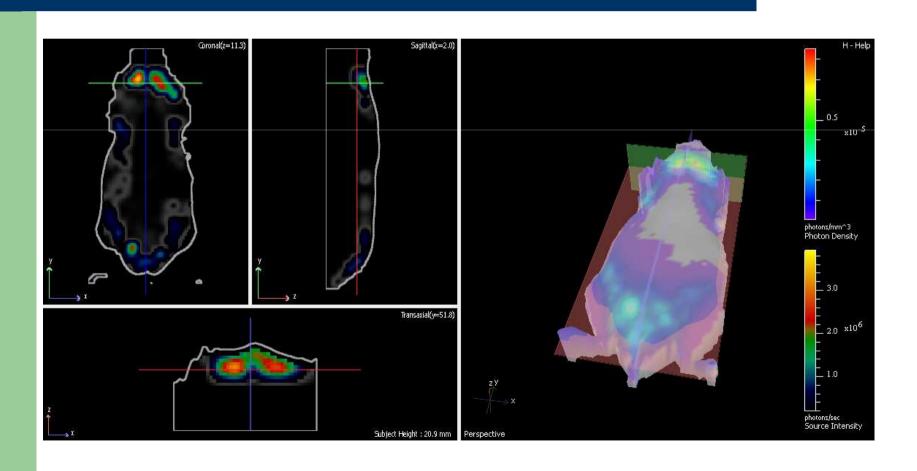


- Median split
- Prognostic value of PRDX1 in estrogen receptor positive (ER+) patients
- In vitro studies have shown that PRDX1 acts as a protector of ER receptor under oxidative stress

Molecular Biology Studies



In vivo models



Thank you