

central european institute of technology

CEITEC

Genomics and proteomics at MU Jiří Fajkus



Genomics

- ✓ interested in GENOME complete genetic info (DNA/RNA sequence) of an organism (3Gb-human)
- ✓ ca. 25 000 hum<u>an genes</u>
- $\checkmark \approx 2\%$ genome







 Potential of genomics : fast analysis of gene mutations and changes in gene expression (at the mRNA level)

 \Rightarrow detection of inherited disorders, pathogens, genetic manipulations, molecular oncology...

✓ Novel DNA sequencing platforms✓ cDNA microarrays



✓ Why PROTEOMICS?

- Only limited repertoire of events directly depends on DNA sequence
 - (25 000 genes \times several millions proteins)
 - 1 organism \Rightarrow 1 genome \times many proteomes













Core facility "Genomics and Proteomics"

✓ proteome analysis

bacteria, plants, animals, human, stem cells, etc.









Core facility "Genomics and Proteomics": integration of sophisticated technologies

- $\checkmark\,$ isolation, separation of proteins
- ✓ identification of proteins, posttranslational modifications
- ✓ analysis of protein complexes

✓ mass spectrometry – major proteomic tool



Core facility "Genomics and Proteomics"





Examples of present scientific projects

- ✓ Bacterial genetics and genomics
- ✓ Plant cytogenomics
- Molecular analysis of plant development
- ✓ Biology of telomeres
- ✓ Molecular diagnostics and therapy (collab. UH Brno, UH S.A.)



Bacterial genetics and genomics

- ✓ Human, emerging and re-emerging pathogens
- ✓ Whole genome sequencing strategies
 - Whole genome fingerprinting (physical mapping)
 - Comparative genome sequencing (microarray)
 - Pyrosequencing, Solexa sequencing, SOLiD sequencing, Sanger sequencing
- ✓ Analyses of bacterial transcriptome
 - ✓ Gene annotations
 - ✓ Analyses of gene regulatory networks





Goals of whole-genome sequencing

- ✓ Identification of virulence factors (better treatment of infectious diseases)
- ✓ Identification of prominent bacterial antigens (development of new vaccines)
- ✓ Identification of unique genomic sequences (better diagnostics)
- Metagenomic sequencing (definition of human microbiome)
- Epidemiological studies (better preventive measures)







CGS sequencing





Physical mapping of the genome

DNA microarray



Plant Sciences at MU - Why plants?

- ✓ Scientific reasons: unique developmental model
- Political and economical reasons: Plants as the only <u>sustainable</u> and <u>self-renewable</u> tool of energy conservation on Earth; Plants as a <u>multipurpose</u> and <u>low-cost production system</u>













Laboratory of Plant Cytogenomics Methodology

- ✓ construction of large insert DNA libraries
- high-throughput DNA sequencing
- ✓ genome comparisons on a sequence and chromosomal level
 - bioinformatics
 - molecular cytogenetics
 - comparative phylogenomics









Laboratory of Plant Cytogenomics

- ✓ analysis of inter-species
 chromosome and genome collinearity
- ✓ karyotype and genome evolution within phylogenetic frameworks (phylogenomics)
 - ✓ evolution of repetitive DNA elements and structure of plant genomes
 - ✓ karyotype evolution and speciation





Molecular analysis of plant growth and development

- ✓ Functional genomics and proteomics of model plants (*Arabidopsis*) and crop plants (oil-seed rape) as a lowcost production systems for e.g. pharmaceutically active proteins (antibodies) and biomass production
- Analysis of the protein structure and protein engineering



 Identification of Proteins Regulating Plant Biomass Production - cytokinins and their receptors regulate biomass production in plants (in frame of collaboration with Korean colleagues) - application for patent cooperation treaty (PCT) submitted by MU and Korean partner.







✓ Biology of telomeres

- ✓ Structure, evolution and maintenance of plant telomeres (DNA, proteins, chromatin structure, epigenetics
- ✓ Telomeres and telomerase in oncology diagnostics and therapy





Regulation of telomere length in normal and cancer cells by telomerase



 Key processes in telomere biology (and in living cell, in general): interaction networks of proteins, DNA and RNA





Laboratory of separation and analysis of proteins and their complexes

- ✓ description of important protein complexes and their compositions
- kinetics and thermodynamics of protein-protein and protein-DNA interactions
- ✓ study of mechanisms of function and action of biomolecular complexes





SUMMARY – Genomics and proteomics

- Concentration of "universal" and sophisticated genomic and proteomic equipment and expertise in core facility – optimum cost-efficiency
- Potential for synergistic interactions among research groups, capability to solve complex projects
- ✓ Higher competitive strength and attractiveness to international R&D community ⇒ raising of funds for science
- Emphasis on current internationally competitive research groups, yet flexibility in routing of future development
- Connection to major medical institutions (university hospitals, res. institutes), biotechnology incubators and institutes of Czech Academy of Sciences