

## Strengthening collaborations in systems biology

French and German experts hear of a possible new computer-based system to identify new biomarkers for colon cancer. And perhaps there is a new way to design vaccinations based on new understanding that memory cells are generated early in the immune response to infection, Bettina Döbereiner reports.

About 90 participants gathered in the French Embassy in Berlin this November to hear 16 lectures in just one day. In a French-German joint workshop entitled 'Perspectives of systems biology - from modelling to therapy of complex diseases', the talks covered general funding strategies, with presentations of selected institutions and departments specialising in systems biology and introductions of core areas of systems biology research in both countries.

The workshop was initiated by the Scientific Department of the French Embassy and BioTOP Berlin-Brandenburg, which is a central contact and coordination office for



1. David Henderson, 2. Jean-Michael Heard, 3. Prof. Thomas Höfer Pictures: CH.Wegen





all issues concerning biotechnology in the region, and is funded by the Federal States of Berlin and Brandenburg. Welcoming participants, BioTOP's CEO, Dr Kai Uwe Bindseil, explained that the workshop is part of a series of annual conferences organised with the French Embassy since 2007, to cover current scientific issues such as regenerative medicine, obesity or functional food. This time scientific partners were the École Normale Supérieure (ENS) in Paris, and the Max Delbrück Centre for Molecular Medicine (MDC) in Berlin-Buch, Germany.

There are plenty of definitions of systems biology in the literature and even at the workshop definitions varied. However, it is beyond dispute that, as a relatively new field of research in life sciences, systems biology can be seen as a shift of perspective from studying single and specific components of the organism to the focus on relations and interactions between the single components in terms of the whole system in time. Thereby scientists inter alia hope to understand the functioning of complex diseases better and develop enhanced diagnostic tools and more effective therapies. In this approach, mathematical analysis and computational modelling play a key

role to comprehend the complex biological interrelations.

Therefore, multidiscipline is a pre-condition for this new research branch, as was seen in the gathering, which included biologists, chemists, physicists, mathematicians, computer scientists as well as physicians and other healthcare professionals.

That systems biology advances to become a key-issue in life sciences also may be seen in the funding strategies of France and Germany. In her lecture, Karin Efferts PhD from the German Ministry of Education and Research (BMBF) introduced e:Med - Measures for establishing systems biology - the new fund-







4. Bodo Lange, 5. Karin Effertz, 6. Dr Kai Uwe Bindseil

ing concept for systems biology in Germany, launched in 2012. The programme predicts €200 million in funding over the next eight years. Recently, systems biology projects in France have also been funded on a large scale. As Jean-Michel Heard PhD from the French National Research Agency (ANR) reported, in 2011 and 2012, as a part of its so-called future programme, France began funding systems biology for the next ten years, which will amount to around €300 million dedicated to infrastructure, equipment and projects related to systems biology.

All lectures held at the workshops primarily addressed the analysis of complex diseases such as neurodegenerative, cardiovascular or inflammatory, as well as cancer on a very basic, pre-clinical level. One interesting typical systems biology example of the cooperation between biology and the computational sciences is a project headed by Professor Thomas Höfer from German Cancer Research Centre and the BioQuant Centre for Quantitative Analysis of Molecular and Cellular Biosystems, at Heidelberg University. He presented his recent results regarding the fate decisions of lymphocytes during infectious diseases. By means of computer modelling he and his

team recently developed a model for the pathway of the differentiation of naive antigen-specific T cells into long-living memory cells. In the first phase of project, Professor Dirk Busch and his team at Munich Technical University observed the fate of individual naive T cells in infected mice.

Prof. Höfer and colleagues then generated a set of different potential pedigrees of naive T-cells through mathematical analysis. Later on, these models of pedigrees were compared to the observations made in adoptive transfer models. The results were clear - only one of these models was suitable to explain the data. It indicates that the precursors of memory cells are generated early during the immune response to infection, rather than - as had been previously thought - at its very end. If further validated, this new finding for example will have implications vaccination design.

The ppp-project OncoTrack - a more practical oriented research -was presented by two industry representatives, David Henderson PhD, Principal Scientist at Bayer Healthcare and project coordinator, and Bodo Lange PhD, CEO of Alacris Theranostics.

OncoTrack is one of the largest ppp (public/private partnership) projects in systems biology. With funding of around €27 million from the European Innovative Medicines Initiative (IMI) and partners from the pharmaceutical industry, the project involves 21 partners in Europe and is scheduled for completion in five years. The ambitious aim of OncoTrack is to develop a computerbased system to identify new biomarkers for colon cancer. Called ModCell, the computer model was developed by Prof. Hans Lehrach's department at the Max Planck Institute of Molecular Genetics, Berlin, and exclusively licensed to the spin-off company Alacris in 2010.

The system integrates data from sources such as genome, transcriptome, drug action, protein function and their mutant variants obtained from the literature and from public databases. With this unique systems biology approach the project partners analyse the obtained genomic and patient data on colon cancer.

On cell and mouse xenografts the ModCell system is validated to predict optimised drug actions for patients based on genome and transcriptome information from the individual patient's tumour. This approach also aims to identify complex biomarkers that will improve diagnostics and therapy.

## MRSA, C Di staphyloco

New, promising methods, tools and tests for prevention and treatments

Jacquie Michels

A recent webinar, hosted by the American Association of Clinical Chemistry, featured doctors Duane Newton of the University of Michigan and Susan Novak-Weekly of Kaiser Permanente. The presentation, entitled Molecular Testing for Methicillin-Resistant Staphylococcus Aureus (MRSA) and C. Difficile (C. diff): 'Using New Tools to Identify and Reduce Healthcare-Associated Infections' (HAIs), outlined the problem of rising rates of these two virulent and possibly fatal infections, possible new testing methods, and suggestions for controlling outbreaks where risk is greatest.

'C. diff and MRSA are healthcareassociated infections,' Dr Newton pointed out. 'They are among the top ten causes of death in any such setting - and all are preventable.' According to Newton, there are up to two million infections caused in the

healthcare setting, and as many as 1.7 million related deaths. These include urinary tract, bloodstream, surgical site and pulmonary infections.

Dr Newton is not surprised by these risks. 'Healthcare institutions use surgical procedures, injections, and in-dwelling devices, such as catheters, and there are countless sources of contamination,' he said. 'Infection can be passed between provider and patient. And over-use of antibiotics has made this problem infinitely more complicated.'

The USA's Department of Health and Human Services has devised some initiatives to eliminate health-care-based infections.'A steering committee was formed at the federal level, and an action plan was published in 2010 that contains clinical, administrative, and governmental directives and strategies to manage this problem,' Dr Newton said. 'Some of their findings seem self-evident, but calling in federal involvement and organising

