

advancements and a unique feature of the CDTM series is the integration of 'Technical Application' sessions. For CDTM2010 Olav Scheimann (St Andrews University, UK) presented a session on EPR spectroscopy and its value in resolving conformations and dynamics of membrane associated proteins. Werner Witke (Leica Microsystems, DE) presented a session on Total Internal Reflection Microscopy (TIRF) and its capacity to study crucial events that occur on and very close to the plasma membrane. Dries Vercauteren (Ghent University, BE) presented a session comparing pharmacological and molecular approaches to inhibit endocytic

pathways and the requirements to integrate both to gain a more accurate picture of uptake pathways.

CDTM2010 welcomed over 180 registered delegates from 27 different countries and from five continents. Over 90 delegate posters were presented, with six of these also selected to highlight their work via short talks. The publication of the CDTM2010 abstracts in the journal *Drug Discovery Today* is a major landmark for the symposia series reflecting its international standing in the scientific community, its maturation into an important event in the Drug Delivery calendar and its ability to

consistently deliver high quality science both on the podium and through the delegate contributions. It is hoped that all who attended CDTM2010 departed with renewed energy for the scientific challenges they face. We look forward to CDTM2012.

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## DELEGATE ABSTRACTS

### A1 Design and development of polymeric nanoparticles for targeted delivery of nucleic acid-based therapeutics to tumor sites

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Nucleic acids are widely used as potent therapeutics in cancer research. They can either promote gene expression by bringing a gene either not expressed or under-expressed into tumor cells (cDNA), or alternatively silence expression of genes such as oncogenes (RNAi mediators). However, before they can be efficiently translated to the clinic, this technology requires some optimization: nucleic acids and their vehicles need for instance to be protected from rapid elimination from the bloodstream (opsonization, clearance, and nuclease-mediated degradation) and the specificity of tumor addressing has to be validated. Hence a polymeric nanoparticulate carrier encapsulating nucleic acids, either plasmid DNA or siRNA, was developed. Nanoparticles are composed of (1) PLGA, a well tolerated and biodegradable polymer, (2) PEG groups to

avoid opsonization, (3) PEI moieties to complex nucleic acids and to enhance cytosolic delivery and (4) RGD sequence for active tumor targeting. Nanoparticles were formulated by double emulsion or water-in-oil-in-water method. Physical properties of such nanoparticles were assessed by dynamic light scattering (size and polydispersity index) and laser doppler electrophoresis (zeta potential). The efficiency of nucleic acid encapsulation into the carrier was determined by the Picogreen assay. Cytotoxicity and transfection capacity were assessed in an *in vitro* model of B16F10 melanoma cells. To date, various designs of nanoparticles were successfully formulated with appropriate size, surface charge and encapsulation efficiency. The PLGA nanoparticles did not show cytotoxic effects on cells and, although less efficient than PEI alone, allowed DNA delivery into tumor cells.

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### A3 Pulmonary delivery of mRNA: *in vitro* and *in vivo* evaluation

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Gene therapy is a very promising field of research in medicine. The success of gene based therapeutics will depend on a well

thought-out and well-designed delivery system, which should guide the nucleic acids into the desired compartment of the selected cells. However, humans and other organisms have developed natural barriers that protect their body against different kinds of pathogens or intruders. During the evolution of the human being, these barriers have become almost perfect and difficult to overcome. The nuclear membrane, one of the final barriers that protect our genes, appears to be the most important and the crucial one to overcome in non-viral gene delivery. In this work we try to avoid the need to overcome this barrier by intracellular delivery of mRNA instead of pDNA. mRNA delivery has many advantages. First, mRNA does not have to overcome the nuclear barrier and therefore mRNA can transfect also non-dividing cells or dividing cells independent of their cell cycle. Second, mRNA cannot integrate in the genome. Consequently, mRNA mediated gene expression is transient and the risk of insertional mutagenesis can be excluded. Third, there is no need to select a promoter [1]. In this work we evaluate whether mRNA complexed with cationic liposomes (composed of e.g. the cationic lipid GL67) are able to transfect the respiratory tissue of mice. The efficacy of the mRNA:liposome complexes and the gene expression kinetics will be studied and compared with pDNA:liposome complexes. In this study we focus in particular on GL67-based liposomes. GL67 is an amphiphile consisting of a cholesterol anchor lined to a spermine headgroup in a 'T-shape' configuration. It was proven that GL67 based liposomes are the most effective non-viral pulmonary gene delivery systems [2]. Evaluation of the