

Damien Lhenry , Franck Denat , Christine Goze , Victor Goncalves , Manuel Larrouy
ICMUB Institute Chemistry, 9 Avenue Alain Savary, Dijon 21078, France

- 28 **Site-specific labeling of cysteine-tagged Nanobodies for use in molecular imaging**
Sam Massa^{1,2,3}, Catarina Xavier¹, Jens de Vos^{1,2,3}, Vicky Cavelliers^{1,4}, Tony Lahoutte^{1,4}, Serge Muyldermans^{2,3}, Nick Devoogdt^{1,2}
¹Vrije Universiteit Brussel In vivo Cellular and Molecular Imaging laboratory, Laarbeeklaan 101, 1090 Brussels, Belgium
²Vrije Universiteit Brussel Laboratory of Cellular and Molecular Immunology, Pleinlaan 2, 1050 Brussels, Belgium
³VIB Department of Structural Biology, Pleinlaan 2, 1050 Brussels, Belgium
⁴Universitair Ziekenhuis Brussel Nuclear medicine department, Laarbeeklaan 103, 1090 Brussels, Belgium
- 29 **Development of a hybrid probe for specific imaging of bacterial infections**
Bunschoten Anton, Welling Mick M., Rood Marcus T. M., Buckle Tessa, van Leeuwen Fijs
Leiden University Medical Center, Radiology, Interventional Molecular Imaging Lab - Leiden, Netherlands
- 30 **Development of radiotracers for PET imaging of CB2 receptor**
Slavik Roger¹, Grether Uwe², Müller Adrienne¹, Gobbi Luca², Krämer Stefanie D.¹, Weber Markus³, Schibli Roger¹, Ametamey Simon M.¹, Linijing Mu¹
¹ETH Zürich, Institute of Pharmaceutical Sciences - Zürich, Switzerland
²F. Hoffmann-La Roche Ltd., Discovery Chemistry - Basel, Switzerland
³Neuromuscular Diseases Unit/ALS Clinic - St. Gallen, Switzerland
- 31 **Clinically-relevant, customizable agents for multimodal imaging**
Srinivas Mangala¹, Lajoinie G.2, de Korte Chris L.³, **Heerschap Arend**³, Versluis Michel², Figdor Carl G.¹, de Vries I. J. M.¹
¹Radboud Institute for Molecular Life Sciences, Tumor Immunology - 6500HB, Netherlands
²University of Twente, Physics of Fluids - Enschede, Netherlands
³Radboud UMC, Radiology - Nijmegen, Netherlands
- 32 **The Development of Nuclear Receptor Imaging Agents**
Louis Allott¹, John Greenman¹, Christopher Cawthorne¹, Graham Smith²
¹The University of Hull The School of Biological, Biomedical and Environmental Sciences, Cottingham Road, Hull, Great Britain
²Institute of Cancer Research Division of Radiotherapy and Imaging, Sutton, Great Britain
- 33 **Synthesis and preclinical evaluation of ¹⁸F-labelled F3 derivatives for the PET imaging of Tumour Angiogenesis**
Phoebe Y. H. Lam , Katherine A. Vallis
University of Oxford Gray Institute for Radiation Oncology and Biology, Radiobiology Research Institute, Churchill Hospital, Oxford, OX3 7LJ, Great Britain
- 34 **Nanoplatfoms for Bimodal Imaging of Sentinel Lymph Node**
Maurício Morais¹, Maria P. C. Campello¹, Catarina Xavier², Sophie Hernot², João D. G. Correia¹, Vicky Cavelliers^{2,3}, Tony Lahoutte^{2,3}, **Isabel Santos**¹
¹University of Lisbon C2TN/IST, CTN, Pólo de Loures do IST, Estrada Nacional 10 (km 139,7), 2695-066 Bobadela LRS, Portugal
²Vrije Universiteit Brussel In Vivo Cellular and Molecular Imaging Laboratory, ---- Brussels, Belgium
³UZ Brussel Nuclear Medicine Department, Brussels, Belgium

Introduction

Sentinel Lymph Node Detection (SLND) is very important for cancer management [1]. The preoperative detection of SLN requires an intradermal injection of a radiolabeled colloid, whereas its intraoperative localization depends on the acoustic signal coming from the hand-held gamma probe as well as on visual confirmation of the radioactive node stained with a blue dye. Although widely used in the clinical setting for SLND, the properties of ^{99m}Tc-based colloids and blue dyes are far from ideal, hampering the surgeon's ability to identify and excise the SLN in a minimal invasive way [2, 3]. The presence of two imaging reporters (radioisotope and fluorophore) in the same probe would enable fast and accurate excision of SLN by radio- or near-infrared (NIR) guided surgery. Therefore, herein we report on new fluorescent radiolabeled mannosylated dextran-based probes for preoperative visualization of SLN by SPECT and PET, respectively, as well as on their use for real-time guidance during surgical excision by NIR optical imaging.

Methods

We have prepared bimodal imaging probes containing a radiometal, ^{99m}Tc or ⁶⁸Ga, a NIR fluorophore and mannose units for Mannose Receptor (MR) targeting. The pharmacokinetics and SLN mapping of the probes were evaluated in a Wistar rat model using nuclear and optical imaging.

Results

The probes, obtained with high radiochemical yield (> 95%) and specific activity, are stable *in vitro*. The biodistribution studies have shown that both probes presented an increased accumulation in the popliteal node with minimal spread to other organs after subcutaneous injection. Moreover, both probes enabled real time optical-guided excision of the SLN.

Conclusions

The new bimodal probes exhibit suitable biological properties for pre- and intraoperative mapping and excision of the SLN.

Acknowledgement / References

COST Action TD1004 is acknowledged for a STSM (STSM-TD1004-040213-027456). The authors also wish to thank the IAEA. M. M. thanks the Fundação para a Ciência e Tecnologia (FCT) for a PhD fellowship (SFRH/BD/48066/2008). FCT is also acknowledged for funding (EXCL/QEQ-MED/0233/2012).

[1] Alitalo A, Detmar M. Interaction of tumor cells and lymphatic vessels in cancer progression. *Oncogene*. 2012;31:4499-508. doi:10.1038/onc.2011.602 onc2011602.

[2] Sharma R, Wendt JA, Rasmussen JC, Adams KE, Marshall MV, Sevic-Muraca EM. New horizons for imaging lymphatic function. *Lymphatic Continuum Revisited*. 2008;1131:13-36. doi:DOI 10.1196/annals.1413.002.

[3] Morais M, Subramanian S, Pandey U, Samuel G, Venkatesh M, Martins M, Pereira S, Correia JDG, Santos I. Mannosylated dextran derivatives labeled with fac-[M(CO)₃]⁺ (M = ^{99m}Tc, Re) for specific targeting of sentinel lymph node. *Mol Pharm*. 2011;8:609-20. doi:10.1021/mp100425p.

Keywords: Bi-Modal Imaging, Nanoparticles, Sentinel lymph node

- 35 **Gold nanoparticles for targeted delivery of radiometals**
Campello Maria P. C.¹, **Francisco Silva**¹, Paulo António¹, Gano Lurdes¹, Morais Maurício¹, Santos Isabel¹, Kannan Raghuraman²