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Roche Organ Transplantation Research Foundation

ANNUAL REPORT 2014



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The Roche Organ Transplantation Research Foundation

The mission of the Roche Organ Transplantation Research Foundation (ROTRF) is to advance the science of organ transplantation in order to improve the care of the thousands of patients undergoing transplantation every year. The results of the funded research projects will contribute to an understanding of many aspects of the clinical and scientific transplantation, such as the mechanisms of long-term organ deterioration and the consequences of tissue injury, and will provide the opportunities to intervene in these processes.

The Foundation is an independent medical research charity that provides operating funds to established academic staff at universities, transplant centres and research institutes. The Foundation supports research in organ transplantation, particularly where there is an unmet medical need.

The funding of the Foundation consists of donations from F. Hoffmann-La Roche Ltd, with an initial sum of 25 million Swiss francs (CHF) over the first five years and renewal donations of 15, 10 and 17.5 million CHF for the following eight and a half years (a total of 67.5 million CHF over 13.5 years). The funds are distributed as grants of up to 300,000 CHF distributed over two years. The foundation is legally independent from F. Hoffmann-La Roche Ltd and is guided solely by the Board of Trustees according to its charter.

The ROTRF is a charity registered in Switzerland, no. CH-270.7.022.678-7.



1. Preface

The ROTRF was established in 1998 as a legally independent, medical research charity with the mandate to award operating grants for research projects in organ transplantation to established members of academic staff at universities, transplant centres and research institutes.

The trustees are proud to announce that in 2014 the ROTRF completed its mission and that no further funding cycles are planned. In May 2014, seven projects were selected to receive funding totaling 1.7 million CHF. Since its establishment in 1998, the Foundation has therefore funded 289 innovative clinical research projects with over 50 million CHF (US\$ 48.5 million), contributing to a better understanding of many clinical and scientific aspects of organ transplantation.

As always, the grant applications were reviewed by a team of experts on the Scientific Advisory Committee and by ad hoc reviewers and these reviews formed the basis of the decisions made by the Board of Trustees. The main criteria for the evaluation of the proposals remained unchanged: scientific merit, originality, the relevance to organ transplantation and the potential for clinical application of the results in the near term. The projects selected for funding in this review cycle included investigations of the relationship between metabolic, immune and microcirculation marker profiles of donor organs and transplantation outcomes, the influence of HCV immunity on alloreactive T cell responses, examination of novel protocols for organ preservation and the role of regulatory and inflammatory B cells in renal allograft rejection. The Trustees wish the newly awarded grantees of Cycle XXIX every success in their investigations.

In addition, the Trustees awarded one-year extensions of 200,000 CHF to one project and 100,000 CHF each to six projects that were coming to the end of their funding, based on a thorough review of the Final Reports and a brief grant extension application. Extension grants were awarded to projects on specificity and function of graft infiltrating B cells, examining the molecular basis of PTLD, the role of circulating fibrocyte and epithelial progenitors in bronchiolitis obliterans syndrome, the role of post-transplantation immunosuppression on viral infection, the role of graft-derived cell-free DNA in biopsy-proven antibody-mediated rejection, strategies to improve cardiovascular risk management and investigations into the dynamics of the adaptive response of microbial communities during allograft colonisation.

The Board of Trustees would like to express its gratitude to F. Hoffmann-La Roche Ltd for the generous donation to support the Foundation. We would like to thank the members of the Scientific Advisory Committee, the ad hoc reviewers and the former Trustees for their commitment and crucial role in assessing the applicants, as well as the Applicants for their innovative proposals and outstanding research efforts. Together, the ROTRF has contributed to advancing the science of organ transplantation and improving the care of the thousands of patients who receive a transplant each year.

On behalf of the Board of Trustees

Philip F. Halloran, MD, PhD, OC Chairman, ROTRF Board of Trustees



2. Facts and Figures

Funding Cycle XXIX – Letters of Intent Submission in October 2013

The ROTRF is pleased to announce that 1.7 million CHF were awarded to seven clinical research projects in 2014.

In Cycle XXIX, 98 eligible Letters of Intent (LOI) for research projects were received up until the deadline of 1 October 2013. The majority was received from North America (47%; USA: 46%; Canada: 1%) and from Europe (45%), with the major European countries being the UK, Germany (7% each), Italy, the Netherlands (6% each), Spain and France (5% each). The remaining 8% of the LOIs were received from Australasia. Based on the reviews by the Scientific Advisory Committee, the Board of Trustees invited 26 applicants to submit a Full Paper Application (FPA), and grants were awarded to seven of the applicants.

The projects funded by the ROTRF focus on important issues in transplantation and have potential for near-term clinical application. Abstracts of all funded projects are available on www.rotrf.org.

Grant Extensions - Submission of extension request in 2014

As of 2014, the ROTRF accepts the submission of grant award applications for extensions to projects that are about to close. During 2014, fourteen applications were received, seven from the USA, and one application each from Australia, Canada, France, Spain, Germany, Switzerland and the UK. Funding was awarded to seven of the applicants, six of which received 100,000 CHF funding each for one-year extensions to their grants and one received 200,000 CHF funding for a one-year extension to his grant.

The recipients were Prof. Joren Madsen, Prof. Olivia Martinez, Dr Bernhard Schmidt, Prof. Christian van Delden (who had each been awarded an ROTRF grant in Cycle XXV), Dr He Xu (Cycle XXVI) and Dr Howard Slater (Cycle XXVII).



Statistics on Applications to the ROTRF

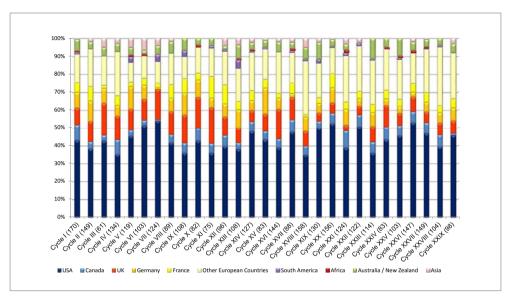


Figure 1. Geographical distribution of the applicants who submitted Letters of Intent (LOI) during the first 29 ROTRF funding cycles. The total number of LOIs submitted per cycle is shown in brackets.

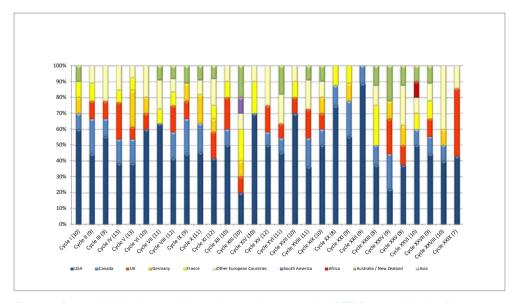
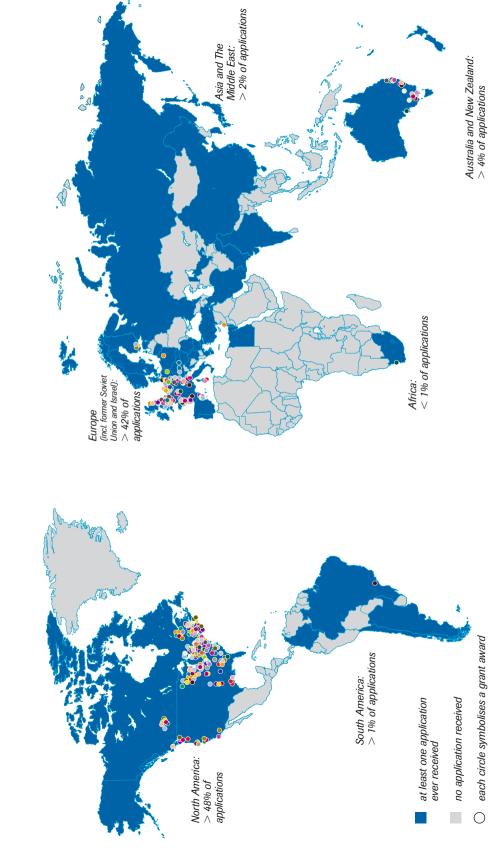


Figure 2. Geographical distribution of the applicants who were awarded ROTRF grants during the first 29 ROTRF funding cycles. The total number of grants awarded per cycle is shown in brackets.

The Global View: Distribution of the ROTRF Applications and Grant Awards Worldwide





3. ROTRF Grant Awards in Cycle XXIX

The abstracts of these projects are available on the ROTRF homepage.

Research Grant Awards - Cycle XXIX

Dr Agustin Lahoz, IIS-Hospital La Fe, Valencia, Spain "Fast Metabolomic Assessment of Donor Liver Quality prior to Transplant"

Dr Marc Martinez-Llordella, King's College, London, UK "Influence of HCV heterologous immunity on alloreactive T cell responses in liver transplantation"

Prof. Vassilios Papalois, Imperial College London, London, UK "Targeting Post Transplant Kidney Injury through Machine Organ Preconditioning"

Dr James Reynolds, Case Western Reserve University, Cleveland, USA "S-nitrosylation, Oxygenation, and Donor Support"

Prof. David Rothstein, University of Pittsburgh, Pittsburgh, USA "The Role of Regulatory and Inflammatory B Cells in Renal Allograft Rejection"

Dr Candice Roufosse, Imperial College, London, UK "Microcirculation injury in the renal allograft - transcripts, ultrastructure and immunophenotype"

Prof. Megan Sykes, Columbia University, New York, USA "A blood biomarker of kidney transplant outcomes: tracking and donor-reactive T cells"



4. Board of Trustees

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Chairman: Professor Philip Halloran	Director, Alberta Transplant Applied Genomics Centre Division of Nephrology & Immunology University of Alberta Edmonton, Canada
Dr Dominique Borie	Senior Medical Director Product Development Immunology Genentech San Francisco, USA
Professor Andrew Bradley	Clinical Director of Transplantation Services Professor of Surgery Department of Surgery University of Cambridge Cambridge, UK
Professor Robert Colvin	Director, Immunopathology Research Laboratory Department of Pathology Harvard Medical School Massachusetts General Hospital Boston, USA
Professor Allan Kirk	Chairman of Surgery, Department of Surgery, Duke University Durham, USA
Professor Gerhard Opelz	Director, Department of Transplantation Immunology University of Heidelberg Heidelberg, Germany
Professor Giuseppe Remuzzi	Director, Department of Immunology and Clinical Transplantation Ospedali Riuniti di Bergamo and Istituto di Ricerche Farmacologiche "Mario Negri" Bergamo, Italy
Professor Alberto Sanchez-Fueyo	Professor, Institute of Liver Studies King's College London King's College Hospital London London, UK



5. Scientific Advisory Committee

Prof. Maria-Luisa Alegre

Dept of Medicine The University of Chicago Chicago, USA

Prof. Anthony d'Apice

Dept of Clinical Immunology St. Vincent's Hospital Melbourne, Australia

Prof. Jeremy Chapman

University of Sydney Westmead Hospital Westmead, Australia

Prof. Sandy Feng

UCSF Medical Center Transplant Surgery University of California San Francisco, USA

Prof. Jay Fishman

Infectious Disease & Compromised Host Program Massachusetts General Hospital and Harvard Medical School Transplant Boston, USA

Prof. Xavier Forns

Liver Unit Hospital Clinic Barcelona, Spain

Prof. Gregg Hadley

Division of Transplant Surgery The Ohio State University Columbus, USA

Prof. Bruce Kaplan

Dept of Medicine Arizona Health Science Center Tucson, USA

Prof. Philip O'Connell

Transplantation
Westmead Hospital
Westmead. Australia

Prof. Jeffrey Platt

Surgery and Microbiology and Immunology University of Michigan Ann Arbor, USA

Prof. Heinz Regele

Dept of Pathology Medical University of Innsbruck Innsbruck, Austria

Prof. Angus Thomson

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