

University of Groningen

## Translational multiple sclerosis research in primates

Dunham, Jordon Tyler-Nathan

**IMPORTANT NOTE:** You are advised to consult the publisher's version (publisher's PDF) if you wish to cite from it. Please check the document version below.

*Document Version*

Publisher's PDF, also known as Version of record

*Publication date:*

2017

[Link to publication in University of Groningen/UMCG research database](#)

*Citation for published version (APA):*

Dunham, J. T-N. (2017). Translational multiple sclerosis research in primates: Mind the gap [Groningen]: Rijksuniversiteit Groningen

**Copyright**

Other than for strictly personal use, it is not permitted to download or to forward/distribute the text or part of it without the consent of the author(s) and/or copyright holder(s), unless the work is under an open content license (like Creative Commons).

**Take-down policy**

If you believe that this document breaches copyright please contact us providing details, and we will remove access to the work immediately and investigate your claim.

Downloaded from the University of Groningen/UMCG research database (Pure): <http://www.rug.nl/research/portal>. For technical reasons the number of authors shown on this cover page is limited to 10 maximum.

# **Translational multiple sclerosis research in primates**

Mind the gap

# **Translationeel multiple sclerose onderzoek in primaten**

Denk aan de kloof

**Jordon Tyler-Nathan Dunham**





# Translational multiple sclerosis research in primates

Mind the gap

## PhD thesis

to obtain the degree of PhD at the  
 University of Groningen  
 on the authority of the  
 Rector Magnificus Prof. E. Sterken  
 and in accordance with  
 the decision by the College of Deans.

This thesis will be defended in public on

Wednesday 18 October 2017 at 12.45

by

**Jordon Tyler-Nathan Dunham**

born on 21 August 1986  
 in Marion, Indiana, USA



**Supervisors**

Prof. J.D. Laman

Prof. B.A. 't Hart

**Co-supervisor**

Dr. Y.S. Kap

**Assessment Committee**

Prof. J.J.G. Geurts

Prof. O.C.M. Sibon

Prof. T. van Laar



*"It's a funny thing about coming home. Looks the same, smells the same, and feels the same.  
You realized what has changed is you."*

F. Scott Fitzgerald



The research described in this thesis was performed at the Dept. of Immunobiology at the Biomedical Primate Research Centre, Rijswijk, the Netherlands and University Medical Center, Dept. of Neuroscience, at the University of Groningen, The Netherlands.

Financial support was provided by the European Union's Seventh Framework Programme for research, technological development and demonstration under grant agreement no. 31672 (Neurokine).

Financial support for publication of this thesis was provided by the Biomedical Primate Research Centre, U-CyTech and the University Medical Center, Groningen.

Illustrations: F. van Hassel, H. van Westbroek and J. Dunham  
Lay-out: F. van Hassel  
Printed by: Ridderprint BV  
ISBN: 978-94-034-0037-2

© Jordon Dunham, 2017. No parts of this thesis may be reproduced or transmitted, in any form, without permission in writing from the author.



# Table of Contents

<b>1. Introduction</b> .....	<b>9</b>
1.1 General introduction .....	11
1.2 The common marmoset as an indispensable animal model for immunotherapy development in multiple sclerosis .....	39
1.3 Aims and outline of thesis .....	53
<b>2. Peripheral mechanisms</b> .....	<b>57</b>
2.1 Blockade of CD127 exerts a dichotomous clinical effect in marmoset experimental autoimmune encephalomyelitis .....	59
2.2 Analysis of the cross-talk between Epstein-Barr virus-infected B cells and T cells in the marmoset .....	79
<b>3. Into the CNS</b> .....	<b>103</b>
3.1 Oxidative injury and iron redistribution are pathological hallmarks of marmoset experimental autoimmune encephalomyelitis .....	105
3.2 A qualitative and quantitative analysis of grey matter demyelination in different experimental autoimmune encephalomyelitis induction protocols in marmosets .....	125
3.3 Severe oxidative stress in an acute inflammatory demyelinating model in the rhesus monkey .....	145
<b>4. General Discussion</b> .....	<b>161</b>
<b>Appendices</b> .....	<b>177</b>
Abbreviations .....	178
Summary .....	181
Samenvatting .....	183
Acknowledgements .....	185
Curriculum vitae .....	189
List of publications .....	190





# Introduction

Multiple Sclerosis  
Inflammation  
Neurodegeneration  
Progressive MS  
EAE  
Grey Matter  
Axon  
Mitochondria  
TH17  
Sclerosis  
Damage  
Erosion



