CONTENTS

Chapter 1  Introduction  9

Chapter 2  Finding ways to determine clinical disease progression: responsiveness and cut-off points for significant change in primary progressive MS  23

2.1  The search for responsive clinical endpoints in primary progressive multiple sclerosis  25

2.2  Progression on the Multiple Sclerosis Functional Composite in multiple sclerosis: what is the optimal cut-off for the three components?  37

Chapter 3  How do different commonly applied scales compare in predicting future clinically important outcome in progressive MS?  47

3.1  Clinical scales in progressive MS: predicting long-term disability  49

3.2  Walking speed, rather than Expanded Disability Status Scale, relates to long-term patient-reported impact in progressive MS  61

Chapter 4  Detecting clinically relevant changes in progressive MS  77

Chapter 5  Summary and general discussion  93

Samenvatting  105

Appendices  111

List of abbreviations  113

Clinical measures  115

About the author  143

Dankwoord  147