

## **Timeline of Progress in MS Research**

This timeline highlights a selection of major landmarks in research into multiple sclerosis and the launch of programs to propel this research forward. These are only a few of thousands of advances in basic and clinical research that have laid the foundation for the rapid progress we're seeing today.

1421	First documented case of MS; St. Lidwina of Schiedam
1860-70	First studies of myelin and glial cells in brain tissue
1868	First correlation of MS clinical symptoms with central nervous system pathology; disease named "Sclerose en plaques" by Jean Martin Charcot
1869	First attempts to treat MS with gold chloride, zinc, sulfate, silver nitrate, strychnine and electrical stimulation (by Charcot)
1928	Discovery that myelin is produced by oligodendrocyte glial cells
1933	Acute experimental allergic encephalomyelitis (EAE) developed as model for MS
1936	Discovery that lymphocytes are involved in immune function
1943	First detailed description of the composition of myelin
1946	National MS Society USA Launched
1947	First research grant to Elvin A. Kabat, MD
1950	National Institute of Neurological Diseases founded at National Institutes of Health through the work of the Society
1950	Society funds first major survey of MS in the U.S. and Canada
1954	First Fellowship programs to train MS Scientists offered
1965	Society-convened panel of experts develops precise criteria for diagnosing MS
1969	Society co-funds research of steroid ACTH, which becomes the first drug shown to speed recovery from MS relapses
1974 federal fundin	Society convinces U.S. Congress to appoint commission on MS; resulting report increases g for MS research
1980	Society funds first large trial of any form of interferon (interferon alpha), stimulating interest in interferons for treating MS
1981	First MRI pictures of a brain affected by MS are produced, revolutionizing MS diagnosis
1981	Identification of oligodendrocytes in MS brain with capability for regeneration of myelin
	Society partners with MS Society of Canada to convene international conference leading to clinical trials in MS. The first controlled trial of natural human beta interferon in MS was of this meeting.

1983	Society supports studies of the anti-cancer agent mitoxantrone in an animal model of MS. In 2000, this drug (Novantrone) is approved by FDA to treat worsening MS		
1984	First modern documentation of cognitive problems in MS		
1987	Pilot Research Program established by Society to quickly test novel, high-risk ideas		
1987 rehabilitation	Society establishes program to attract research proposals in patient management, care and		
1988	Health Care Delivery and Policy Research Program established by Society		
1988	First demonstration, using MRI, that there is significant lesion activity in MS brain, even when the disease is clinically quiescent		
1992	Society supports first comprehensive search for genes that make people susceptible to MS, initiating targeted research program in MS genetics		
1993 effects under	Betaseron approved by FDA for relapsing-remitting MS, becoming first drug approved that lying disease. The Society had supported the first study of any type of interferon in MS		
1996 in persons wit	Society grantees find that aerobic exercise improves physical and psychological well-being th MS		
1996	Society develops international consensus on 4 clinical descriptions of M (relapsing-remitting, secondary-progressive, primary-progressive, progressive-relapsing)		
1996	Avonex and Copaxone approved for relapsing-remitting MS. The Society funded early basic research underlying both therapies		
1997	Zanaflex (tizanidine) approved for treatment of spasticity		
1997	Sylvia Lawry Physician Fellowship Program established to train doctors in conducting clinical trials in MS		
1998	Society launches targeted research initiative into gender differences in MS		
1999 Longitudinal S	Society initiates first-ever longitudinal patient-focused database effort, the Sonya Slifka MS Study, providing quality of life and socioeconomic data		
1999	Society grantees first to isolate immature cells in the adult brain capable of developing into replacements for myelin-making cells destroyed by MS		
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1999/2000	Society initiates new clinical trial in estrogen treatment for women with MS and T-cell vaccination, both of which had their origins in early fundamental NMSS research		
2000	Society initiates international collaborative research effort to better correlate the MS lesion with disease state and MRI: The MS Lesion Project		
2000	Novantrone approved in U.S. for worsening forms of MS		
2001	Society task force develops new diagnostic criteria for MS, which may shorten the time it takes a person to receive a firm diagnosis.		
2001	Society collaborates with NIH on a \$20-million research initiative on gender-based differences in immune responses to increase understanding and treatments		
2002	Rebif approved in U.S. to treat relapsing-remitting MS		

2002	New Career Transition Fellowship Program launched to foster promising young MS investigators
2003	Society convenes international Task Force on Nervous System Repair to identify ways to speed development of repair strategies to restore nerve function in MS
2003	Society launches Collaborative MS Research Centers, 5-year awards to team up scientists and clinicians from a variety of fields to work on promising avenues
2003	Italian researchers transplant cells to enhance nerve tissue repair in mice with MS
2004	A pivotal study by a Society Fellow compares the clinical characteristics of MS among hundreds of African- and Caucasian Americans, showing that African-Americans tend to have a more aggressive course
2005	Society launches Promise:2010 campaign to raise at least \$30 million for cutting-edge research and clinical care initiatives including 6 Pediatric MS Centers of Excellence
2005	Society launches initiative to speed nervous system repair and protection clinical trials in MS with 4 international teams funded for \$15.6 million
2005 diagnosis for	"McDonald Criteria" for diagnosing MS updated by Society Task Force, speeding time to many
2005	Society collaborates with NIH, MS Society of Canada and University of Washington on international workshop to move MS rehabilitation research forward.
2006	Tysabri approved in U.S. for treating relapsing MS
2006 term care op	Long-term Care Caucus convened to create national agenda to foster spectrum of long-tions for people with MS
	Society and MS International Federation convene a Stem Cell Research Summit ding stem cell and MS experts to explore the potential of all types of stem cell rch for MS and to set research priorities
2007 the Society's	First large-scale trial of sex hormone estriol gets underway in women with MS, a result of targeting of gender differences
2007	With support from Society to International MS Genetics Consortium, two genes are confirmed to be linked to susceptibility to MS; more likely to be uncovered
2007	Society launches Fast Forward to speed treatments to people with MS by partnering with industry to develop new therapies
	Society's Task Force on MS Epidemiology meets to outline future research directions for cies that will further the search for the cause of MS; through its efforts, a disease tracking piloted by the Centers for Disease Control's ATSDR
	Congressional MS Caucus launched to raise awareness and engage in discussion about s to health care, increase in research funding, disability rights and other MS issues, with m the House and Senate
2008	MS activists secure a place for MS research in the \$50 million Congressionally Directed Medical Research Program, administered through the Department of Defense, plus \$5 million restricted to MS research within the CDMRP
2008	Society funds genome scan by International MS Genetics Consortium of 10,000 patients to validate a large-scale study and to study the influence of copy number variants and gene-to-gene interactions in MS susceptibility
2009	Fast Forward makes investments in 7 companies working on early stage MS therapies

2009	International task force convened by Society publishes landmark guidelines on the complex process of telling MS from look-alike disorders ("differential diagnosis")
2009	Extavia is approved by the FDA as a new brand of interferon beta-1b
2009	Society convenes international workshop on strategies to find the cause of MS, and factors that drive progression and ways to estimate MS frequency
2009 and strengthe	Society holds first-ever Don Tykeson Fellows Conference to stimulate new research ideas in the commitment of bright young people to MS research
2009	Fast Forward joins forces with EMD Serono to fast-track new therapies for MS
2010	Gilenya, first oral disease-modifying therapy for MS, approved for relapsing forms
2010	Ampyra approved to improve walking ability in people with all types of MS
2010	Fast Forward, Juvenile Diabetes Research Foundation and Axxam SpA Join Forces to accelerate development of treatments
2010	Society releases a request for proposals resulting in funding 2 feasibility studies on risk factors that drive MS progression and prognosis
2010	Nuedexta and Botox approved to treat specific symptoms that interfere with quality of life in people with MS
2010	Rapid collaboration and support for research on CCSVI
2010	International team co-supported by Society revises MS diagnostic criteria to reduce the wait for accurate diagnosis
2011	Society supports controlled study to determine whether vitamin D supplements can reduce MS disease activity
2011	Promise: 2010 Nervous System Repair initiative is completed, having jump-started the field, trained scores of promising young investigators, produced over 180 research papers, and leveraged millions of dollars in new funding
2011	Society-supported global consortium doubles the number of MS risk genes identified
2012	Launch of International Progressive MS Alliance to speed the development of therapies
2012	Aubagio approved by FDA as second oral therapy for relapsing forms of MS
2012	Society task force launches effort to create a better tool to track benefits of therapies on disability in MS, leading to formation of the MS Outcome Assessments Consortium
2012	Launch of studies into whether bacteria in the gut influence MS disease activity or risk
2013	Tecfidera approved by FDA as third oral therapy for relapsing forms of MS
2013	Society partners with two foundations to form a research network of 10 leading universities aimed at screening drugs that show potential for treating brain diseases
2013	Barancik Prize for Innovation in MS Research is launched to inspire novel research
2013	Society commits \$2.5 million to support the Network of Pediatric MS Centers to provide essential infrastructure to facilitate research
2013	Researchers co-funded by the Society transplant stem cells derived from human skin into mice to successfully grow nerve-insulating myelin
2013	Launch of MS Prevalence Work Group to update the estimate of MS prevalence in the US
2013	Society-funded researchers report that dietary salt can speed the development of an MS-like disease in mice and may influence immune activity in MS

2014	New, less frequent dose of Copaxone approved
2014	Plegridy, an interferon taken by subcutaneous injection every 2 weeks, is approved
2014	Lemtrada, given in two courses of IV infusion, is approved for relapsing forms of MS
2014	Society convenes Wellness conference to map out gaps and opportunities in research and programs to find wellness solutions for people with MS
2014	Large trial of ibudilast, funded by NIH and Society, begins recruiting participants with progressive MS to test its ability to protect the nervous system from damage
2014	International Progressive MS Alliance awards its first 22 research grants
2015	Ocrelizumab becomes first experimental therapy to show positive effects in primary progressive MS in large-scale trial
2015	First generic form of Copaxone, given by daily injections, approved
2015	International initiative is launched to focus on how having MS and other conditions (comorbidities) influences disease course and treatment
2015	A phase 2 clinical trial co-funded by the Society suggests a pill used to treat epilepsy (phenytoin) has the potential to slow the accumulation of disability in people with MS
2015	Results of phase 2 trial of anti-LINGO suggests it has potential as myelin repair strategy
2015	Society co-hosts international conference on cell-based therapies to forge next steps for cell therapy in MS
2015	Society funding helps launch MS Microbiome Consortium to promote research on the role of gut bacteria in MS progression and treatment
2015	Society's 5-year, \$250 million NOW (No Opportunity Wasted) campaign was successfully concluded, launching more MS research and driving more life-changing progress than what occurred at any other time in the Society's history
2016	Positive results from a phase 3 trial of siponimod for secondary progressive MS break through a long-standing barrier
2016	Positive results announces from two studies of bone marrow-derived stem cells (HSCT) in people with aggressive, relapsing MS; more research focuses on who might benefit and how to reduce risks
2016	Society-funded International Consortium of MS Genetics identifies 200 genetic variations linked to MS, offering new leads to how genes and other factors that make people susceptible to developing MS
2016	The antihistamine clemastine, identified with Society support, shows evidence of promoting myelin repair in a small clinical trial
2016	Society launches two new studies testing the ability of dietary approaches to treat MS symptoms and improve quality of life
2016-17	International Progressive MS Alliance awards three large-scale Collaborative Network Awards to promote solutions for people with progressive MS
2017	FDA approves Ocrevus (ocrelizumab) as first disease-modifying therapy for primary progressive MS, and also as a therapy for relapsing MS
2018	International team co-supported by Society revises MS diagnostic criteria to speed the diagnostic process and reduce the chance of misdiagnosis

2018	FDA approves expansion of the use of Gilenya to include children and adolescents 10 years of age or older with relapsing MS, the first therapy specifically approved to treat pediatric MS
2018	A Phase 2 trial showed that ibudilast could reduce brain atrophy in progressive MS. The trial was funded collaboratively by the National MS Society, the NINDS, its NeuroNext trials network, with support by MediciNova, the supplier of ibudilast
2018-19	Results were announced from several studies of bone marrow-derived stem cells in people with aggressive, relapsing MS. More research is focused on who might benefit and how to reduce risks
2019	A study sponsored by the National MS Society conducted by leading experts estimates that nearly 1 million adults are living with MS in the United States. This is more than twice the previously reported number
2019	FDA approves oral Mavenclad (cladribine) for adults with relapsing forms of MS, and also approves oral Mayzent (siponimod) for adults with clinically isolated syndrome (an initial neurological episode) and relapsing forms of MS
2019	FDA approved Vumerity™ (diroximel fumarate, Biogen and Alkermes plc) for relapsing MS
2020	National MS Society releases recommendations for aHSCT-bone marrow transplant for MS
2020	FDA approves generic form of Tecfidera for relapsing MS
2020	FDA approves Kesimpta® (ofatumumab) for relapsing MS
2020	Society-supported Wellness Research Group publishes exercise and physical activity recommendations for ALL people with MS
2020	National MS Society, Consortium of MS Centers, and others establish the North American MS COVID-19 Clinical Database
2020	FDA approves Oral Bafiertam™ (Monomethyl Fumarate) for relapsing MS
2020	FDA approves Oral Zeposia® (Ozanimod) for relapsing MS