

## **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; 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 nerve-insulating myelin is produced by oligodendrocyte 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 Society 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; becomes first drug to speed recovery from MS relapses
1974	Society convinces U.S. Congress to appoint commission on MS; resulting report increases federal funding 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, revolutionizing MS diagnosis and treatment
1981	Identification of oligodendrocytes in MS brain with capability for regeneration of myelin
1982	Society partners with MS Society of Canada to convene international conference leading to
	standards for clinical trials in MS; the first controlled trial of natural human beta interferon in MS was an outgrowth 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
1988	Health Care Delivery and Policy Research Program established by Society
1988	First demonstration with MRI that MS brain lesions are active even when the disease is clinically quiet

1992	Society supports first comprehensive search for genes that make people susceptible to MS, initiating targeted research program in MS genetics
1993	Betaseron approved by FDA for relapsing-remitting MS, becoming the first disease-modifying therapy; the Society had supported the first study of any type of interferon in MS
1996	Society grantees find that aerobic exercise improves physical and psychological well-being in persons with MS
1996	Society develops international consensus on 4 clinical descriptions of MS (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 approved for treatment of spasticity
1997	Sylvia Lawry Physician Fellowships established to train doctors in conducting clinical trials in MS
1998	Society launches targeted research initiative into gender differences in MS
1999	Society initiates first-ever longitudinal patient-focused database effort, the Sonya Slifka MS Longitudinal 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
1999-00	Society initiates new clinical trial in estrogen treatment for women with MS and T-cell vaccination, both of which had their origins in early Society-funded 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	Society-funded Fellow shows that Black Americans tend to have a more aggressive course than white Americans
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	"McDonald Criteria" for diagnosing MS updated by Society Task Force, speeding time to diagnosis for 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	Long-term Care Caucus convened to create national agenda to foster spectrum of long-term care options for people with MS

2007	Society and MS International Federation convene a Stem Cell Research Summit to explore the potential of all types of stem cell research for MS and to set research priorities
2007	First large-scale trial of sex hormone estriol gets underway in women with MS, a result of the Society's 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 <i>Fast Forward</i> to speed treatments to people with MS by partnering with industry to develop new therapies
2007	Society's Task Force on MS Epidemiology meets to outline future research directions for funding agencies that will further the search for the cause of MS; through its efforts, a disease tracking system is being piloted by the Centers for Disease Control's ATSDR
2007-8	Congressional MS Caucus launched to raise awareness and engage in discussion about access to health care, increase in research funding, disability rights and other MS issues, with members from 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	Society holds first-ever Tykeson Fellows Conference to stimulate collaboration and strengthen 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

Aubagio approved by FDA as second oral therapy for relapsing forms of MS
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
Launch of studies into whether bacteria in the gut influence MS disease activity or risk
Tecfidera approved by FDA as third oral therapy for relapsing forms of MS
Barancik Prize for Innovation in MS Research is launched to inspire novel research
Society commits \$2.5 million to support the Network of Pediatric MS Centers to provide essential
infrastructure to facilitate research
Researchers co-funded by the Society transplant stem cells derived from human skin into mice to
successfully grow nerve-insulating myelin
Launch of MS Prevalence Work Group to update the estimate of MS prevalence in the US
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
New, less frequent dose of Copaxone approved
Plegridy, an interferon taken by subcutaneous injection every 2 weeks, is approved
Lemtrada, given in two courses of IV infusion, is approved for relapsing forms of MS
Society convenes Wellness conference to map out gaps and opportunities in research and
programs to find wellness solutions for people with MS
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
International Progressive MS Alliance awards its first 22 research grants
Ocrelizumab becomes first experimental therapy to show positive effects in primary progressive
MS in large-scale trial
First generic form of Copaxone, given by daily injections, approved
International initiative is launched to focus on how having MS and other conditions (co-morbidities
influences disease course and treatment
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
Results of phase 2 trial of anti-LINGO suggests it has potential as myelin repair strategy
Society co-hosts international conference on cell-based therapies to forge next steps for cell
therapy in MS
Society funding helps launch MS Microbiome Consortium to promote research on the role of gut
bacteria in MS progression and treatment
Society's 5-year, \$250 million NOW (No Opportunity Wasted) campaign successfully concluded,
launching more MS research and driving more life-changing progress than what occurred at any
other time in the Society's history
Positive results from a phase 3 trial of siponimod for secondary progressive MS break through a
long-standing barrier
Positive results announced from two studies of bone marrow-derived stem cells (HSCT) in people
with aggressive, relapsing MS
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 MS
The antihistamine clemastine, identified with Society support, shows evidence of promoting myeling
repair in a small clinical trial
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	More results were announced from several studies of bone marrow-derived stem cells in people
	with aggressive, relapsing MS
2019	A study sponsored by the National MS Society estimates that nearly 1 million adults are living with
	MS in the U.S., more than twice the previously reported number
2019	FDA approves oral Mavenclad for adults with relapsing forms of MS, and also approves oral
	Mayzent for adults with relapsing MS
2019	FDA approved Vumerity for relapsing MS
2020	National MS Society releases recommendations for a HSCT-bone marrow transplant for MS
2020	FDA approves generic form of Tecfidera for relapsing MS
2020	FDA approves Kesimpta 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
2020	COVID-19 Clinical Database and contribute data to global COVID tracking initiative
2020	FDA approves oral Bafiertam for relapsing MS
2020	FDA approves oral Zeposia for relapsing MS
2021	Study shows that stem cells derived from skin cells of people with MS make normal myelin, a plus
2021	for future repair strategies
2021	Studies suggest that changes in the nervous system related to MS begin well before there are
2021	perceptible symptoms Studies add to growing evidence that Black poople with MS may experience werse disease.
2021	Studies add to growing evidence that Black people with MS may experience worse disease FDA approves oral Ponvory for relapsing MS, and the PoNS nerve stimulator for use during physical
2021	therapy to treat walking problems in MS
2021	Society-supported researchers discover a subset of brain cells that fight inflammation with
2021	instructions from gut bacteria, opening new therapy approaches
2022	Pathways to MS Cures research roadmap is published and gains global endorsement
2022	International MS Microbiome study links gut bacteria to MS susceptibility, severity and treatment
2022	in novel study
2022	Two studies show that high doses of vitamin D did not reduce MS disease activity
2022	Study commissioned by the Society showed that the total estimated cost to the U.S. economy,
	including government, industry and individuals, is \$85.4 billion per year
2022	Two studies co-funded by the Society show links between socioeconomic status to increases in
	retinal (eye) nerve fiber damage and vision loss, and to depression and anxiety
2022	Society-funded researchers at Harvard published the strongest evidence yet that the common
	Epstein-Barr virus (EBV) can trigger MS when other risk factors are present
2022	FDA approves Briumvi for relansing MS