

# Masitinib in patients with primary progressive or secondary progressive multiple sclerosis

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		<input type="checkbox"/> Protocol
<b>Registration date</b> 20/05/2022	<b>Overall study status</b> Ongoing	<input type="checkbox"/> Statistical analysis plan
		<input type="checkbox"/> Results
<b>Last Edited</b> 07/06/2022	<b>Condition category</b> Nervous System Diseases	<input type="checkbox"/> Individual participant data
		<input type="checkbox"/> Record updated in last year

## Plain English summary of protocol

### Background and study aims

Multiple sclerosis (MS) is an inflammatory neurodegenerative disease of the central nervous system (CNS). The vast majority of MS drugs primarily benefit active/relapsing forms of MS with limited efficacy in the progressive forms. It is estimated that 10-15,000 people have primary progressive MS in the UK.

Masitinib is a new chemical entity under development in progressive forms of multiple sclerosis. Masitinib targets newly discovered disease mechanisms via its dual action against activated macrophage/microglia and mast cells.

### Who can participate?

Patients with primary and secondary progressive MS without relapse will be eligible to be treated during the study.

### What does the study involve?

Patients will be treated for 96 weeks with masitinib or placebo. They will be then offered the option to participate in open-label extension. As part of the study, patients will have a physical exam, vital signs, and neurological exam. They will have efficacy assessments and questionnaires to evaluate their medical condition and its progress. Patients will have ECGs to monitor cardiac safety and also routine blood samples and PK blood samples. There will be urinalysis and urine cytology. For some patients there will be regular permanency testing.

### What are the possible benefits and risks of participating?

#### Benefits:

Participants will be reimbursed for their travel expenses upon presentation of costs' proofs.

#### Risks:

Masitinib is an experimental drug and there could be adverse events that are not known yet. This is why participants will be monitored closely throughout the study. Participants will need to attend regular clinic visits which could be a burden but this is the only option to monitor their safety and well-being.

### Where is the study run from?

Dokumed (Latvia)

When is the study starting and how long is it expected to run for?  
March 2022 to June 2026

Who is funding the study?  
AB Science (France)

Who is the main contact?  
Dr Seema Kalra, Seema.Kalra@uhnm.nhs.uk  
Dr Maria Carvalho, maria.carvalho@ab-science.com

## Contact information

### Type(s)

Principal investigator

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Scientific

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## Additional identifiers

### Clinical Trials Information System (CTIS)

2021-000639-30

### Integrated Research Application System (IRAS)

1005160

### Central Portfolio Management System (CPMS)

52039

**Protocol serial number**

AB20009

## **Study information**

**Scientific Title**

A 96-week, prospective, multicenter, randomised, double-blind, placebo controlled, phase 3 study to compare efficacy and safety of masitinib dose titration to 4.5 mg/kg/day versus placebo in the treatment of patients with primary progressive or secondary progressive multiple sclerosis without relapse.

**Study objectives**

To evaluate the efficacy and safety of oral masitinib versus placebo in the treatment of patients with primary progressive or secondary progressive multiple sclerosis without relapse.

The secondary objectives of the study are to evaluate the efficacy of masitinib compared with placebo on a range of clinical parameters of multiple sclerosis. The secondary objectives also include the assessment of safety and tolerability of masitinib as compared to placebo in terms of adverse events, vital signs, physical examination, ECG, and clinical laboratory tests.

**Ethics approval required**

Old ethics approval format

**Ethics approval(s)**

Approval pending, ref: 22/WM/0092

**Study design**

Interventional double blind randomized parallel group placebo controlled trial

**Primary study design**

Interventional

**Study type(s)**

Treatment

**Health condition(s) or problem(s) studied**

Multiple sclerosis

**Interventions**

There are 2 parallel arms: masitinib and placebo. Patients will be treated for 96 weeks and will be offered an additional treatment extension to week 108 if they have benefit. The drug is administered orally twice a day. Drug formulation is AB1010/Masitinib (100 mg and 200 mg tablets) or placebo tablets (100 mg and 200 mg tablets). Dose: 3.0 mg/kg/day from Week 0 to Week 4, and then to 4.5 mg/kg/day from Week 5 to Week 96/Early Termination or to Week 108. The study uses IWRS and investigators will receive notification of the arm the patient is randomized on.

**Intervention Type**

Drug

## Phase

Phase III

## Drug/device/biological/vaccine name(s)

mansitinib

## Primary outcome(s)

Time to confirmed (12-weeks CDP [Confirmed Disability Progression]) Expanded Disability Status Scale (EDSS) progression. The EDSS progression is defined as 1-point worsening when EDSS baseline score  $\leq 5.5$  or 0.5 if baseline score  $> 5.5$  from randomization to Week 96.

## Key secondary outcome(s)

1. Expanded Disability Status Scale (EDSS):

1.1. Time to confirmed (24-weeks CDP) EDSS progression. Progression is defined as 1-point worsening when EDSS score  $\leq 5.5$ , or 0.5 if baseline score  $> 5.5$ )

1.2. Expanded Disability Status Scale (EDSS): Absolute and ordinal change from baseline considering all measurements up to Week 96

2. Time to EDSS score of 7.0 Clinical Global Assessment Tools:

2.1. Timed 25-foot walk (T25-FW) from baseline up to Week 96 and 12 weeks confirmed worsening using 20% threshold

2.2. Nine-hole peg test (9-HPT), right and left hands sides (finger dexterity) from baseline up to Week 96 and 12 weeks confirmed worsening using 20% threshold

2.3. The Symbol Digit Modalities Test (SDMT) from baseline up to Week 96 and 12 weeks confirmed worsening using 4-point threshold

3. Brain MRI Assessments:

3.1. Brain Volume and Lesions will be measured and assessed at Baseline, Week 48 and Week 96, or early termination (only if patient discontinues after Week 48 and more than 24 weeks have elapsed since last MRI) for the following endpoints: Brain atrophy - Percent brain volume change (PBVC) from baseline at Week 96 or early termination

3.2. New/newly enlarged T2 lesion count (compared with baseline MRI scan) at Week 96 or early termination

4. Quality of Life assessment:

4.1. Multiple Sclerosis Quality of Life (MSQOL)-54 instrument from baseline up to Week 96

4.2. Modified Fatigue Impact Scale (MFIS) from baseline up to Week 96

4.3. Hamilton Depression Rating Scale (HAM-D) from baseline up to Week 96

4.4. Disability Impact Profile (DIP) from baseline up to Week 96

5. Relapses measured using patient records:

5.1. Occurrence of new or worsening neurological symptoms attributable to MS

5.2. Symptoms persisting for  $> 24$  hours

5.3. Symptoms not attributable to confounding clinical factors (e.g., fever, infection, injury, adverse reactions to medications)

5.4. Symptoms immediately preceded by a stable or improving neurological state for at least 30 days

5.5. Symptoms accompanied by objective neurological worsening consistent with an increase of at least half a step on the EDSS scale

6. Biomarker(s):

6.1. Comparison of serum Neurofilament Light Chain (NFL) and Glial Fibrillary Acidic Protein

(GFAP) levels at Baseline and Week 96 or early termination (only if patient discontinues after Week 48 and more than 24 weeks have elapsed since last test) These biomarkers are to be tested in a subgroup of 200 patients of pre-selected sites

**Completion date**

01/06/2026

## Eligibility

**Key inclusion criteria**

1. Patients with either primary progressive or secondary progressive multiple sclerosis with onset of symptoms at least five years before inclusion and with no relapse diagnosed according to the 2017 revised McDonald's criteria at least two years before screening
2. Patients with Expanded Disability Status Scale (EDSS) score between 3.0 to 6.0 (both inclusive) at screening and baseline
3. Patients with an EDSS score progression  $\geq 1$  point with no improvement during 2 years before screening
4. Absence of T1 Gadolinium-enhancing brain lesions at baseline as measured by MRI at screening

**Participant type(s)**

Patient

**Healthy volunteers allowed**

No

**Age group**

Adult

**Sex**

All

**Key exclusion criteria**

1. Patients suffering from a disease other than MS that would better explain the patient's neurological clinical signs and symptoms and/or MRI lesions observed at screening
2. Inability to complete screening MRI (contraindications for MRI) and/or any known allergy or hypersensitivity or any contra-indication to gadolinium macrocyclic
3. Patients treated with other disease modifying treatments in the time frames and conditions mentioned under previous treatment wash out period, assessed at baseline
4. Patients with lymphocytes  $< 1.0 \times 10^9/L$  at screening and at baseline

**Date of first enrolment**

31/03/2022

**Date of final enrolment**

01/06/2026

## Locations

**Countries of recruitment**

United Kingdom

England

Argentina

Belgium

France

Germany

Greece

Hungary

Italy

Netherlands

Norway

Poland

Portugal

Romania

Russian Federation

South Africa

Spain

Sweden

Ukraine

**Study participating centre**  
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## Sponsor information

**Organisation**  
Dokumeds

## Funder(s)

**Funder type**  
Industry

**Funder Name**  
AB Science

## Results and Publications

### Individual participant data (IPD) sharing plan

The current data sharing plans for this study are unknown and will be available at a later date

### IPD sharing plan summary

Data sharing statement to be made available at a later date

### Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
<a href="#">HRA research summary</a>			28/06/2023	No	No