

# Gene therapy for Wiskott-Aldrich Syndrome (WAS)

<b>Submission date</b> 03/05/2011	<b>Recruitment status</b> No longer recruiting	<input type="checkbox"/> Prospectively registered
		<input type="checkbox"/> Protocol
<b>Registration date</b> 20/05/2011	<b>Overall study status</b> Completed	<input type="checkbox"/> Statistical analysis plan
		<input checked="" type="checkbox"/> Results
<b>Last Edited</b> 11/04/2019	<b>Condition category</b> Haematological Disorders	<input type="checkbox"/> Individual participant data

**Plain English summary of protocol**  
Not provided at time of registration

## Contact information

**Type(s)**  
Scientific

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75015

## Additional identifiers

**EudraCT/CTIS number**

**IRAS number**

**ClinicalTrials.gov number**  
NCT01347346

**Secondary identifying numbers**  
GTG003.08

# Study information

## Scientific Title

Phase I/II clinical trial of haematopoietic stem cell gene therapy for the Wiskott-Aldrich Syndrome

## Study objectives

Studying the safety and efficacy of an ex vivo gene therapy using a lentiviral vector containing the human Wiskott-Aldrich Syndrome protein gene in patients with WAS.

## Ethics approval required

Old ethics approval format

## Ethics approval(s)

Committee to Protect People (Comité de Protection des Personnes) - Ile de France 2 approved on 11th August 2009, (ref : 2009-04-01)

## Study design

Open labelled non-randomised single centre phase I/II cohort study

## Primary study design

Interventional

## Secondary study design

Non randomised controlled trial

## Study setting(s)

Hospital

## Study type(s)

Treatment

## Participant information sheet

Not available in web format, please use the contact details below to request a patient information sheet

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

Wiskott-Aldrich Syndrome

## Interventions

Ex vivo gene therapy using patient's autologous CD34+ cells transduced with a lentiviral vector containing the human WASP gene.

Patients undergo either a bone marrow harvest or a leukapheresis. They then receive a conditioning myeloablative regimen while CD34+ cells are selected in their bone marrow and transduced with the lentiviral vector (3 days). Patients then receive their transduced CD34+ cells (as in autologous bone marrow transplantation).

There are no real doses, simply quantity of CD34+ cells transduced will depend on the amount of bone marrow harvest and quality of transduction. This is part of the parameters that are being assessed in the trial.

Duration of the study follow-up is 2 years.

## **Intervention Type**

Drug

## **Phase**

Phase I/II

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

Gene therapy

## **Primary outcome measure**

1. Safety of conditioning regimen (haematopoietic recovery within 6 weeks assessed by absolute neutrophil count (ANC) above  $0.5 \times 10^9 /l$ )
2. Safety of the transduction procedure [as assessed by availability of greater than  $1 \times 10^6$  CD34+ cells per kg; retrospective undetectable (replication-competent lentiviruses) RCL; and cell viability equal to or greater than 70%, in accordance with the GMO release criteria].
3. Engraftment of genetically corrected haematopoietic progenitors and/or differentiated cells in peripheral blood and/or in bone marrow (as assessed by evidence of vector sequences or transgene expression in the cells)
4. Reconstitution of cell mediated and humoral immunity (as assessed by evidence of changes in T cell function and circulating immunoglobulin levels).
5. Correction of microthrombocytopenia (as assessed by increased blood platelet counts, expected to rise above 50,000/mm<sup>3</sup> and platelets size restoration)

## **Secondary outcome measures**

1. Reduction in frequency of infections (evaluated from 2nd year after treatment by clinical history, complete physical examinations, haematological and microbiological tests)
2. Resolution/reduction of autoimmunity (a decrease from baseline observations assessed by clinical examination)
3. Improvement in eczema (a decrease from baseline observations assessed by clinical examination)
4. Reduction in bruising and bleeding episodes (as assessed by clinical monitoring)

## **Overall study start date**

16/05/2011

## **Completion date**

31/12/2013

# **Eligibility**

## **Key inclusion criteria**

1. Males of all ages
2. Severe WAS (clinical score 3-5) or absence of WAS protein in peripheral blood mononuclear cells determined by Western blotting and flow cytometry

3. Molecular confirmation by WAS gene DNA sequencing
4. Unless disease severity indicates that one cannot wait for 3 months (score 5; refractory thrombocytopenia with platelets < 5000 with bleeding or severe autoimmunity)
5. Lack of HLA-genotypically identical bone marrow after 3 month search
6. Lack of a 10/10 or 9/10 antigen HLA-matched unrelated donor after 3 month search
7. Lack of a HLA-matched cord blood after 3 month search
8. Parental, guardian, patient signed informed consent/assessment
9. Willing to return for follow-up during the 2 year study and lifelong for off study review
10. Only for patients who have received previous allogenic haematopoietic stem cell transplant
- 10.1. Failed allogenic haematopoietic stem cell transplant
- 10.2. Contraindication to repeat allogenic transplantation

**Participant type(s)**

Patient

**Age group**

Adult

**Sex**

Male

**Target number of participants**

5

**Key exclusion criteria**

1. Patient with HLA-genotypically identical bone marrow
2. Patient with 10/10 or 9/10 antigen HLA-matched unrelated donor or with HLA-matched cord blood
3. Contraindication to leukapheresis
  - 3.1. Anaemia (Hb < 8g/dl)
  - 3.2. Severe vascularitis
  - 3.3. Refractory thrombopenia
    - 3.3.1. Contraindication to bone marrow harvest
    - 3.3.2. Contraindication to administration of conditioning medication
4. Human immunodeficiency virus (HIV) seropositive patient

**Date of first enrolment**

16/05/2011

**Date of final enrolment**

31/12/2013

**Locations****Countries of recruitment**

France

**Study participating centre**

Unité d'Immunologie et d'Hématologie Pédiatriques  
Paris  
France  
75015

## Sponsor information

### Organisation

Genethon (France)

### Sponsor details

1 bis, rue de l'Internationale  
Evry  
France  
91000

### Sponsor type

Industry

### Website

<http://www.genethon.fr>

### ROR

<https://ror.org/03fj96t64>

## Funder(s)

### Funder type

Industry

### Funder Name

Genethon (France)

## Results and Publications

### Publication and dissemination plan

Not provided at time of registration

### Intention to publish date

### Individual participant data (IPD) sharing plan

## IPD sharing plan summary

Not provided at time of registration

## Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
<a href="#">Results article</a>	results	21/04/2015	11/04/2019	Yes	No