# Development of a haemophilia physiotherapy intervention for optimum musculoskeletal health in children

Submission date	Recruitment status  No longer recruiting	[X] Prospectively registered		
07/09/2020		☐ Protocol		
Registration date	Overall study status Completed Condition category	Statistical analysis plan		
04/11/2020		☐ Results		
Last Edited		Individual participant data		
04/11/2020	Haematological Disorders	Record updated in last year		

#### Plain English summary of protocol

Background and study aims

Children are born with haemophilia. Females carry the disorder and usually males are affected. It is a disorder affecting 1 in 10,000 people where the blood does not clot normally, leading to bleeding into muscles and joints. As a result, muscles become weak and joints become painful and difficult to move. "Being able to participate in games and activities with their friends" is one of the things that matters most to boys with haemophilia. At present, there is a lack of robust evidence to determine whether muscle strengthening exercise can improve or negatively affect outcomes for young children with haemophilia. With the help of boys with haemophilia, their parents and physiotherapists, the researchers have developed an exercise programme designed to increase muscle strength. This study aims to find out if the exercise programme might have an effect on joint pain and movement participation in games and activities and improve health in the long term.

Who can participate?

Boys aged between 6 and 12 with severe or moderate haemophilia

#### What does the study involve?

66 boys will be allocated to a group that is asked to complete a 12-week exercise routine to strengthen their leg muscles and another 66 boys to a group that does not do the exercises. The boys will be allocated at random, so that each boy has an equal chance of being in either group. The researchers will monitor the boys throughout the study by measuring their muscle strength, how far they can walk in 6 minutes and time taken to ascend and descend 12 steps. The researchers will also record how physically active the boys are using a wrist band as well as how satisfied they are with their health.

What are the possible benefits and risks of participating?

It is not known whether participants will benefit directly from participating in this study. However, the researchers will collect information which might improve future care for all patients with haemophilia including patients in this study. They do not expect any reactions or side effects of doing the exercises, but they cannot be certain. When muscles are worked harder

than they are used to, or in a different way, soreness in muscles may occur 1-2 days after exercise. This is common when starting an exercise programme and some people may notice some discomfort or aching in their muscles 1-2 days after doing exercises.

Where is the study run from?

This research is being organised by the Haemophilia Centre, East Kent Hospitals University NHS Foundation Trust. However, participants can participate at their local Haemophilia Centre.

When is the study starting and how long is it expected to run for? September 2019 to September 2023

Who is funding the study? National Institute for Health Research (NIHR) (UK)

Who is the main contact?
Dr David Stephensen
david.stephensen@nhs.net

# Contact information

#### Type(s)

Scientific

#### Contact name

Dr David Stephensen

#### **ORCID ID**

https://orcid.org/0000-0002-6175-3343

#### Contact details

Kent Haemophilia Centre
East Kent Hospitals University NHS Foundation Trust
Canterbury
United Kingdom
CT1 3NG
+44 (0)1227866481
david.stephensen@nhs.net

# Additional identifiers

Clinical Trials Information System (CTIS)

Nil known

Integrated Research Application System (IRAS)

282478

ClinicalTrials.gov (NCT)

Nil known

Protocol serial number

# Study information

#### Scientific Title

Single-blinded two-arm pragmatic randomised controlled trial of an exercise intervention versus usual care in children with haemophilia

#### **Acronym**

DOLPHIN-II

#### **Study objectives**

A 12-week exercise intervention increases muscle strength of the dominant knee extensors more than usual care at 24 weeks in children with haemophilia aged 6-12 years.

#### Ethics approval required

Old ethics approval format

#### Ethics approval(s)

Approved 03/11/2020, South West - Cornwall & Plymouth Research Ethics Committee (Level 3, Block B, Whitefriars, Lewins Mead, Bristol, BS1 2NT, UK; +44 (0)2071048071; cornwallandplymouth.rec@hra.nhs.uk), REC ref: 20/SW/0154

#### Study design

Single-blinded two-arm pragmatic randomized controlled trial

### Primary study design

Interventional

## Study type(s)

Treatment

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

Children with haemophilia

#### **Interventions**

Randomisation of eligible participants will be performed with the online randomisation service http://sealedenvelope.com/. Utilising the secure, centralised and independent service, participants will be randomly allocated into one of the two groups (treatment group 1: 12-week muscle strengthening exercise intervention; treatment group 2: usual physiotherapy care for 12 weeks) on a ratio of 1:1. Allocation will be stratified by centre to ensure the same numbers of participants at each site are allocated to the two groups. The unblinded principal investigator at each site will be issued with a password to access the service to create a new randomisation for study participants. The trial manager will oversee randomisations by site and over time.

#### Intervention Type

Behavioural

#### Primary outcome(s)

Muscle strength of the dominant knee extensors measured using maximum voluntary isometric contraction at baseline, 12 and 24 weeks

#### Key secondary outcome(s))

- 1. Muscle strength of the ankle plantarflexors and non-dominant knee extensors measured using maximum voluntary isometric contraction at baseline, 12 and 24 weeks
- 2. Physical function measured using the 6-minute timed walk, timed up and down stairs, triple hop and vertical jump tests at baseline, 12 and 24 weeks
- 3. Physical activity measured using daily step counts, sedentary time, time spent in moderate and vigorous-intensity physical activity using a wearable activity tracker at baseline, 12 and 24 weeks
- 4. Quality of life and QALY measured using the EuroQol-5D (EQ-5D) and Child Health Utility 9D (CHU9D) at baseline, 12 and 24 weeks
- 5. Safety and adherence assessed using adverse events, pain and exercise counts at 12 weeks 6. Intervention practicalities assessed using semi-structured interviews at the end of the intervention at 12 weeks

#### Completion date

30/09/2023

# Eligibility

#### Kev inclusion criteria

- 1. Diagnosis of severe or moderate haemophilia and registered with a UK Haemophilia Centre
- 2. Aged 6 12 years
- 3. Inhibitor or non-inhibitor
- 4. Able to follow simple verbal instructions and provide informed consent

#### Participant type(s)

Patient

## Healthy volunteers allowed

No

#### Age group

Child

#### Lower age limit

6 years

#### Upper age limit

12 years

#### Sex

Male

#### Key exclusion criteria

1. Children 13 years or older (boys aged 13 years or older are excluded due to confounding effects of non-linear increases in neuromuscular maturation and muscle strength that occur in

boys during puberty and adolescence)

- 2. von Willebrand disease
- 3. Past history of fracture or trauma to the lower limb
- 4. Past history of orthopaedic surgery of the lower limb
- 5. Past history of acquired brain injury
- 6. Past history of any other disturbance of the central nervous system
- 7. Joint or muscle bleed in the lower limb in the past 6 weeks
- 8. Presence of lower limb pain
- 9. Participants who are unable to fully comply with verbal instructions

#### Date of first enrolment

01/01/2021

#### Date of final enrolment

01/03/2023

## Locations

#### Countries of recruitment

**United Kingdom** 

England

# Study participating centre

Kent Haemophilia Centre

East Kent Hospitals University NHS Foundation Trust Canterbury United Kingdom CT1 3NG

# Sponsor information

#### Organisation

East Kent Hospitals University NHS Foundation Trust

#### **ROR**

https://ror.org/02dqqj223

# Funder(s)

#### Funder type

Government

#### **Funder Name**

Research for Patient Benefit Programme - NIHR201588

#### Alternative Name(s)

NIHR Research for Patient Benefit Programme, Research for Patient Benefit (RfPB), The NIHR Research for Patient Benefit (RfPB), RfPB

#### Funding Body Type

Government organisation

#### **Funding Body Subtype**

National government

#### Location

United Kingdom

## **Results and Publications**

#### Individual participant data (IPD) sharing plan

The dataset will not be made available. The study involves participants with a rare condition, and although the data will be anonymised, some participant-level data may result in deanonymisation. Data will be held on a secure database by the sponsor.

#### IPD sharing plan summary

Not expected to be made available

#### Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
HRA research summary			28/06/2023	No	No
Participant information sheet	Participant information sheet	11/11/2025	11/11/2025	No	Yes