# Effectiveness of a sildenafil citrate suspension for pulmonary hypertension in children: a randomised pragmatic trial

Submission date	Recruitment status	Prospectively registered
30/01/2008	No longer recruiting	[_] Protocol
Registration date	Overall study status	[] Statistical analysis plan
09/05/2008	Completed	[_] Results
Last Edited	Condition category	Individual participant data
09/05/2008	Circulatory System	[] Record updated in last year

#### Plain English summary of protocol

Not provided at time of registration

## **Contact information**

**Type(s)** Scientific

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## Additional identifiers

EudraCT/CTIS number

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers N/A

## Study information

#### Scientific Title

Sildenafil citrate suspension versus sildenafil citrate powder paper for reduction of pulmonary hypertension in children during cardiac catheterisation: a randomised pragmatic trial

#### **Study objectives**

In Brazil, sildenafil citrate is the most accessible option for the treatment of PH and it is widely used in most hospitals. When licensing sildenafil for treatment of PH, the regulating agency kept it forbidden for people under 18 years old. In the absence of an appropriate pharmaceutical formulation for children, the staff usually crush the tablets to be added to liquid food. However, modifying a commercially available medication may lead to increased toxicity, undesirable side effects, decreased efficacy, poor patient compliance because of the medications taste and potential hazards to health care workers. Our study hypothesis is that a compounding suspension made by a pharmacist in controlled conditions will do better than the powder papers used in routine.

#### Ethics approval required

Old ethics approval format

#### Ethics approval(s)

The project was submitted to the Ethics Committee of the Oswaldo Cruz Foundation (ref: CAAE - 0068.0.011.000-07) and will also be submitted to the Ethics Committee of every participating hospital.

#### **Study design** Randomised pragmatic double-blind multicentre trial

**Primary study design** Interventional

Secondary study design 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

Pulmonary hypertension

#### Interventions

Power calculations: 60% of patients reduce at least 20% the mean pulmonary arterial pressure after using powder papers of Sildenafil citrate (SC). To demonstrate that at least 80% of patients would present 20% reduction after using SC suspension with an alpha error of 5% and power of 80% it would be necessary to recruit 182 patients. The pilot intends to recruit 20 patients.

#### Disease information:

Pulmonary hypertension (PH) is a haemodynamic and clinic disease characterised by a progressive increase of pulmonary vascular resistance and a decrease of pulmonary vascular capacitance leading to right ventricular failure and death. It may occur in isolation (idiopathic pulmonary arterial hypertension) or it may be associated to a variety of systemic disorders (scleroderma, lupus, human immunodeficiency virus [HIV] infection) or cardiopulmonary pathologies such as congenital heart disease. PH is a rare condition and secondary pulmonary arterial hypertension prevalence is less than 0.0001% (statistics about secondary pulmonary hypertension, 2007). Although severe pulmonary arterial hypertension is uncommon, the prognosis of these patients is life threatening and treatment options are limited.

#### Interventions:

A sildenafil citrate suspension developed from the crushed tablets will be compared with the powder papers usually given to the children.

 Sildenafil citrate suspension: the suspension will be prepared from both commercially available tablets and sildenafil citrate powder and the formulation must be easy to prepare at hospital pharmacies. The physical, chemical and microbiology stability of this suspension will be accessed simulating in-use conditions and the ones established by Brazilian drug laws.
Sildenafil citrate powder papers: the powder papers will be prepared from the crushed commercial tablets and addition of a diluent. The tablets will be sent to a private pharmacy to compound sildenafil citrate 5 mg powder papers (sachets) as its already done as a routine.

Both arms will receive sildenafil citrate 0.2 mg/kg as a single dose during cardiac catheterisation.

#### Intervention Type

Drug

**Phase** Not Specified

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

Sildenafil citrate (SC)

#### Primary outcome measure

The proportion of patients in each group that reduce pulmonary arterial hypertension in at least 20% twenty minutes after the administration of sildenafil citrate 0.2 mg/kg, during cardiac catheterisation.

#### Secondary outcome measures

1. At 20 minutes measured by cardiac catheterisation after administration of sildenafil citrate:

- 1.1. Pulmonary vascular resistance
- 1.2. Pulmonary vascular resistance/systemic vascular resistance
- 1.3. Oxygenation index or oxygen requirement

1.4. Cardiac output

- 1.5. Mean systemic arterial pressure and surgery indication based on the response to sildenafil
- 2. At 24 hours: any adverse events
- 3. At 2 months: cardiac surgery and death

## **Overall study start date** 01/05/2008

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**Completion date** 01/11/2008

## Eligibility

#### Key inclusion criteria

1. Children from 0 to 18 years old, either sex

2. Diagnosed as having PH associated with congenital heart disease

3. Needing cardiac catheterisation with sildenafil to evaluate the response to the drug as a decision criteria to cardiac surgery

**Participant type(s)** Patient

**Age group** Child

**Lower age limit** 0 Years

**Upper age limit** 18 Years

**Sex** Both

**Target number of participants** 182

**Key exclusion criteria** Children will be excluded if: 1. The clinician believes that sildenafil represents an additional risk for the patient 2. The carrier refuses to sign the informed consent

Date of first enrolment 01/05/2008

Date of final enrolment 01/11/2008

## Locations

**Countries of recruitment** Brazil **Study participating centre National Instutute of Quality Control in Health** Rio de Janeiro Brazil 21040-900

### Sponsor information

**Organisation** National Institute of Quality Control in Health - Oswaldo Cruz Foundation (Brazil)

**Sponsor details** Av. Brasil 4365 Manguinhos Rio de Janeiro Brazil 21040-900

**Sponsor type** Research organisation

Website http://www.incqs.fiocruz.br

ROR https://ror.org/04jhswv08

## Funder(s)

**Funder type** Research organisation

**Funder Name** Oswaldo Cruz Foundation (Brazil)

**Funder Name** National Institute of Cardiology (Brazil)

## **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