

Neurodevelopmental outcome after neonatal hypoglycaemia: a multi-centre randomised controlled trial comparing intensive treatment versus expectant glucose monitoring in 'high risk' newborns

Submission date 23/08/2007	Recruitment status No longer recruiting	<input checked="" type="checkbox"/> Prospectively registered
		<input type="checkbox"/> Protocol
Registration date 23/08/2007	Overall study status Completed	<input type="checkbox"/> Statistical analysis plan
		<input checked="" type="checkbox"/> Results
Last Edited 06/02/2020	Condition category Pregnancy and Childbirth	<input type="checkbox"/> Individual participant data

Plain English summary of protocol
Not provided at time of registration

Study website
<http://www.studies-obsgyn.nl/hypoexit.nl>

Contact information

Type(s)
Scientific

Contact name
Ms Debbie Nuytemans

Contact details
Department of Neonatology
Emma Children's Hospital
Academic Medical Centre (AMC)
P.O. Box 22660
Amsterdam
Netherlands
1100 DD
+31 (0)20 566 3477
HYPO-EXIT@AMC.nl

Additional identifiers

EudraCT/CTIS number

Nil known

IRAS number**ClinicalTrials.gov number**

Nil known

Secondary identifying numbers

ZonMW Doelmatigheid 80-007022-98-07406

Study information

Scientific Title

Neurodevelopmental outcome after neonatal hypoglycaemia: a multi-centre randomised controlled trial comparing intensive treatment versus expectant glucose monitoring in 'high risk' newborns

Acronym

HYPO-EXIT

Study objectives

Current clinical practice varies widely, especially for infants with 'moderate' hypoglycaemia, due to lack of methodological sound studies. This leads to both over- and under-treatment of hypoglycaemic infants.

This study-protocol is directed at the comparison of two accepted management strategies at both ends of the current treatment-spectrum of moderate hypoglycaemia in 'high risk' newborns: an intensive treatment versus an expectant monitoring strategy.

Ethics approval required

Old ethics approval format

Ethics approval(s)

Ethics approval received from the local medical ethics committee

Study design

Multicentre, randomised, single-blinded, active controlled, parallel group trial

Primary study design

Interventional

Secondary study design

Randomised controlled trial

Study setting(s)

Hospital

Study type(s)

Treatment

Participant information sheet

Health condition(s) or problem(s) studied

Developmental-disabilities, blood-glucose, hypoglycaemia

Interventions

In the intensive treatment arm the aim is to increase the glucose concentration above 2.5 mmol /l within three hours by increasing the carbohydrate intake by oral nutrition and/or intravenous glucose administration.

In the expectant arm the aim is to maintain the glucose concentration above 2.0 mmol/l by the usual oral nutrition protocol.

Intervention Type

Drug

Phase

Not Specified

Drug/device/biological/vaccine name(s)

Glucose

Primary outcome measure

Primary outcome is neurodevelopment at 18 months, assessed with the Bayley Scales of Infant Development.

Secondary outcome measures

Current secondary outcome measures as of 04/09/2019:

1. Costs for medical treatment and hospital admission until 18 months of age:
 - 1.1 costs for diagnostic tests and treatment of the infant (glucose measurements, supplemental feeding, tube-feeding, intravenous glucose administration), and hospitalization costs for both the infant and mother
 - 1.2 costs for medical consumption related to neurodevelopmental impairment until the age of 18 months (visits to healthcare professionals and hospital admission after the neonatal period)
2. Plasma glucose concentrations and carbohydrate intake (breastfeeding, oral or enteral feeding and intravenous glucose)
3. Frequency of treatment failure, defined as infants who become severely hypoglycaemic despite the treatment they received (frequency and severity of hypoglycaemia episodes after randomization).

Previous secondary outcome measures:

Secondary outcomes are costs for medical treatment and hospital admission until 18 months of age.

Overall study start date

01/10/2007

Completion date

01/04/2013

Eligibility

Key inclusion criteria

Current participant inclusion criteria as 04/09/2019:

Infants greater than or equal to 35 weeks gestational age and greater than or equal to 2000 g with one of the four major risk factors for neonatal hypoglycaemia:

1. Small-for-Gestational-Age infants (SGA, birth-weight-for-gestational-age less than P10)
2. Large-for-Gestational-Age infants (LGA, birth-weight-for-gestational-age greater than P90)
3. Near-term infants 35 0/7 to 36 6/7 weeks gestational age with a birth weight greater than 2000 g
4. Infants of Diabetic Mothers (IDM)

Birth-weight-for-gestational-age is defined according to the growth charts of the Perinatale Registratie Nederland (PRN).

Previous participant inclusion criteria:

Infants greater than or equal to 35 weeks gestational age and greater than or equal to 2000 g with one of the four major risk factors for neonatal hypoglycaemia:

1. Small-for-Gestational-Age infants (SGA, birth-weight-for-gestational-age less than P10)
2. Large-for-Gestational-Age infants (LGA, birth-weight-for-gestational-age greater than P90)
3. Near-term infants 35 0/7 to 36 6/7 weeks gestational age with a birth weight greater than 2000 g
4. Infants of Diabetic Mothers (IDM)

Birth-weight-for-gestational-age is defined according to the Kloosterman growth charts.

Participant type(s)

Patient

Age group

Neonate

Sex

Both

Target number of participants

800

Total final enrolment

689

Key exclusion criteria

Current participant exclusion criteria:

Infants with serious co-morbidity will be excluded, because their co-morbidity can also affect neurodevelopment:

1. Very preterm infants (less than 34 6/7 weeks gestational age)
2. Severe perinatal asphyxia - presence of at least three of the next criteria:
 - 2.1. Signs of intrauterine asphyxia, like late decelerations on Cardiotocogram (CTG) or meconium stained amniotic fluid
 - 2.2. Arterial umbilical cord pH less than 7.10
 - 2.3. Delayed initiation of spontaneous respirations greater than 5 minutes after birth
 - 2.4. Five minute Apgar score less than 5

2.5. Multi-organ failure

3. Severe perinatal infection: requiring support of vital functions (infants without clinical signs of infection who are treated with antibiotics because of suspected perinatal infection can be included)
4. Respiratory insufficiency requiring respiratory support
5. Severe hypotension requiring vasopressor support
6. (Strong suspicion of) a syndrome or major congenital malformations

Other exclusion criteria:

7. Intravenous glucose administration before randomization
8. (Strong suspicion of) inborn error of metabolism
9. (Strong suspicion of) hyperinsulinism, except infants of diabetic mothers
10. No informed consent

Previous participant exclusion criteria:

Infants with serious co-morbidity will be excluded, because their co-morbidity can also affect neurodevelopment:

1. Very preterm infants (less than 34 6/7 weeks gestational age)
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Other exclusion criteria:

7. (Strong suspicion of) inborn error of metabolism
8. (Strong suspicion of) hyperinsulinism, except infants of diabetic mothers
9. No informed consent

Date of first enrolment

01/10/2007

Date of final enrolment

01/04/2011

Locations

Countries of recruitment

Netherlands

Study participating centre

Department of Neonatology
Amsterdam
Netherlands
1100 DD

Sponsor information

Organisation

Academic Medical Centre (AMC) (The Netherlands)

Sponsor details

Emma Children's Hospital
P.O. Box 22660
Amsterdam
Netherlands
1100 DD
+31 (0)20 566 2131
emma@amc.uva.nl

Sponsor type

Hospital/treatment centre

Website

<http://www.amc.uva.nl/>

ROR

<https://ror.org/03t4gr691>

Funder(s)

Funder type

Research organisation

Funder Name

The Netherlands Organisation for Health Research and Development (ZonMw) (The Netherlands)

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?
Results article	results	06/02/2020	06/02/2020	Yes	No