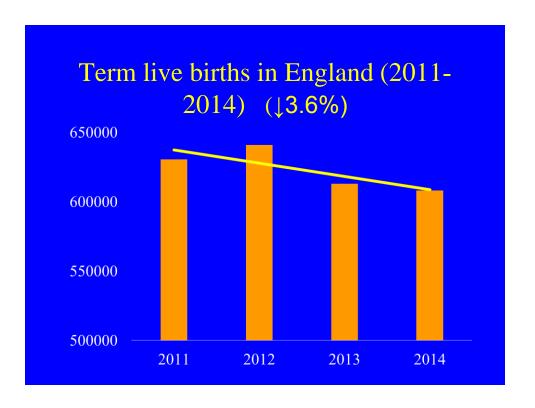


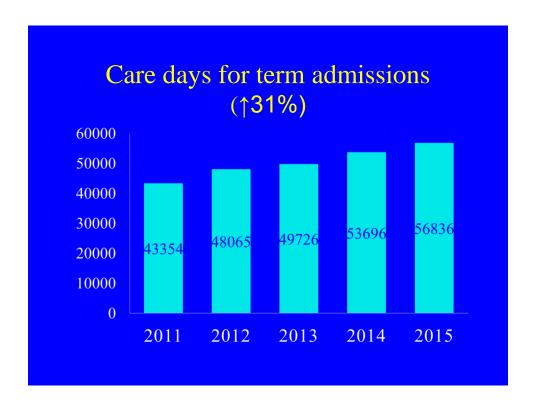


RESOURCE ALERT

Atain programme







Why is this important

RESUSCITATION AFTER FETAL HYPOXIA-ISCHAEMIA

- Staff present with appropriate competence
- **ABCD**
- Start with air
- When to stop
- Communication
- **Documentation**
- Simulation training



ROLE OF THERAPEUTIC HYPOTHERMIA

- Acute profound hypoxia-ischaemia
- Standard of care since 2010
- Early passive
- Active in specialist centre
- 30% reduction in adverse outcome

PERINATAL STROKE

- Associated with complications of labour and delivery but causal link unknown
- Rare blood disorder
- May be silent
- Clinical signs not specific



NEONATAL INFECTION

- NICE guidance 2012
- GBS, Gram negative, herpes simplex
- Antenatal, intrapartum, postnatal acquisition
- Early signs very subtle
- Associated pathology
- Narrow therapeutic window
- Low threshold to screen and treat

NEONATAL JAUNDICE

- NICE guidance 2010
- Gestation specific thresholds
- Clinical signs may be subtle
- Often narrow therapeutic window
- But placing by a window not effective
- Underlying pathology esp meningitis
- Kernicterus hearing, basal ganglia

NEONATAL METABOLIC ADAPTATION

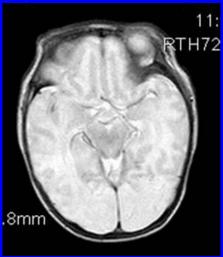
- 1. Blood glucose control
- 2. Alternative fuels lactate, ketone bodies

Hypoglycaemia and brain injury.

1920s	Insulin overdose in adults
1933	Insulin treatment of psychiatric disorder 30 mins hypoglycaemic coma
1950s	Neuronal necrosis demonstrated
Present	Insulin, OHAs - treatment error or self harm Insulin producing tumours and IEMs Failure of metabolic adaptation



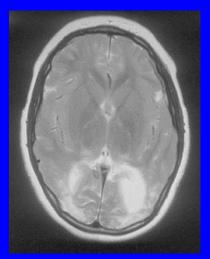
Imaging - acute (courtesy of Dr N Stoodley)



Term baby, 5 days, admitted at 3 days poor feeding, fitting, unrecordable blood

Increased parenchymal signal and loss grey-white matter differentiation in posterior parietal and occipital regions

Imaging - long term (courtesy of Dr N Stoodley)



Imaging on childhood following neonatal hypoglycaemia Atrophy, gliosis and ulegyria in posterior parietal and occipital regions



Adverse neurodevelopmental outcome of moderate neonatal hypoglycaemia

661 infants 6808 samples

Bwt 1337g (315) Gest 30.5 (2.7)

Age 48hrs - 8 weeks

Causal relationship with signs in 5

Lucas et al, 1998

Occurrence of hypoglycaemia

	Plasma glucose (mmol/l)		
	<0.6	<1.6	<2.6
Overall	10%	28%	66%*
On 1 day	8%	20%	32%
On 2 days	1%	4%	18%
On <u>></u> 3 days	1%	4%	16%**

Variation between centres: * 53-79%

** 4-31%

Lucas et al, 1998



Factors associated with hypoglycaemia

- Neonatal unit
- Apgar 5 < 5
- Bwt <1000g
- SGA

Lucas et al, 1998

Regression analysis

Bayley motor and developmental scales Dependent variables:

Independent variables: Days hypoglycaemia*

Sex

Birthweight <10th centile Gestation <30/40 Clinical complications*

SBR >170 Apgar 5 <5

Social and educational*

Lucas et al, 1998



Relative risk of neurodevelopmental impairment (Bayley score ≤70 or CP)

Days hypoglycaemia	Adjusted RR
0	1
1-2	1.1
3-4	2.2
> 5	3.5 (1.3-9.4)
	Lucas et al, 1998

Hyperinsulinaemic newborn monkeys

Pre and post delivery sub cut insulin by pump Not fed after birth

Tested 8 months:

Pre-training - emotionality and adaptability Matching to sample tasks Delayed matching to sample - memory

Schrier et al, 1990



Hyperinsulinaemic newborn monkeys

Results:

10 hr hypo -

more training difficulties more procedural alterations (adaptability) additional help from tutors (motivation)

No differences in -

completion of tasks (with extra attention) personality characteristics neurology

Schrier et al, 1990

Operational thresholds

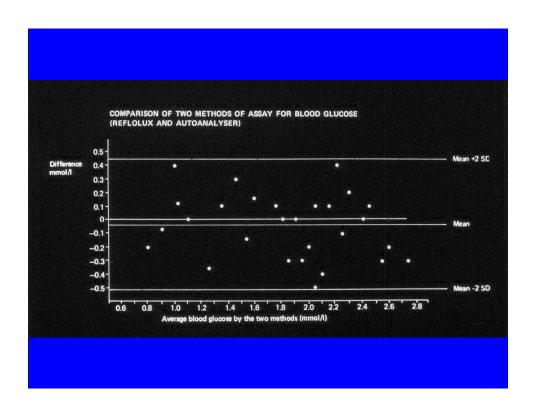
 The concentration of plasma or whole blood glucose at which intervention should be considered to increase the glucose level.



Operational thresholds

Infants with clinical signs: 2.5 mmol/l

Infants at risk: Persistently <2.0 mmol/l <1.0 mmol/l at any time





38 weeks

Labetalol in pregnancy and postnatal Fetal distress in labour, NVD Placenta small and gritty Good condition, no resuscitation Birthweight 2192g (2nd centile) Head circumference 33cm (25th-50th centile)

Neonatal hypoglycaemia?

1hr: paed informed of weight, "watch feeding and temperature"

11 hours: paed examination normal

14 hours: discharged

28 hours: MW visit, baby floppy and

unresponsive, cardioresp depression

28.5 hours: NNU, BM unrecordable, cardiac arrest, fits, renal impairment



Follow up:

MRI: parieto-occipital cortical and white matter injury

Global developmental delay

Autistic features

Neonatal hypoglycaemia?

37 weeks

Emergency CS for breech

Meconium stained liquor

Good condition, no resuscitation

Birthweight 2900g (50th-75th centile)

Head circumference 34cm (75th-91st centile)



Meconium obs normal, discontinued

Support to attach at breast

12 hours: unsettled not feeding, EBM by syringe

16 hours: lethargic, placed skin to skin, would

not suck, gasping, paed informed

16.5 hours: BM unrecordable, paed attended, NNU

17 hours: cardiorespiratory arrest x 3, fits, renal

impairment

Neonatal hypoglycaemia?

Follow up:

MRI: injury to basal ganglia, cortex, white matter Very long chain acyl coA dehydrogenase deficiency Microcephaly, epilepsy and severe disability



37 weeks

Diet controlled gestational diabetes **Emergency CS for CTG abnormalities** Good condition, no resuscitation Birthweight 2566g (25th centile) Head circumference 33cm (25th centile)

Neonatal hypoglycaemia?

Planned to breast fed 18 hours - not fed, sleepy BG 0.4mmol/l Took bottle slowly, remained sleepy BG 1.4mmol/l **Admitted NNU** IV glucose, infection screen



Age (hrs)	BG (mmol/l)
18	0.4
19	1.4
20	1.6
21	3.7
25	5.8

Neonatal hypoglycaemia?

1.8 mmol/l BG Ammonia 112 umol/l Cortisol 87 nmol/l **TSH** 3.27 mU/l Thyroxine 30.3 pmol/l Insulin <1.0 mU/l **BOH** 533 umol/l 234 umol/l Acac

Organic acids, amino acids, carnitine profile - normal

Infection screen negative



Maximum glucose requirement 6mg/kg/min

Remained sleepy until day 4

No further low BG

Day 2 - EBM tube feeds commenced

Day 3 - Successfully weaned from iv glucose

Day 4 - Fully breast/cup fed (pulled tube out)

Follow up - no neurodevelopmental concerns

Infants at risk of abnormal neonatal metabolic adaptation

- Altered maternal metabolism
 - -intrapartum administration of glucose
 - -maternal drug treatment
 - -diabetes
- Secondary to neonatal complications
 - -infection
 - -polycythaemia
 - -perinatal hypoxia-ischaemia
 - -hypothermia
 - -prematurity
- IUGR
- Neonatal hyperinsulinism transient or prolonged.
- Endocrine disorders
- Inborn errors of metabolism



Blood glucose and brain injury The evidence

Healthy full term None

Prolonged hypoglycaemia Preterm

Confounding factors

IUGR None direct

? ADD

Prolonged with signs Cortical loss, global delay

Clinical recommendations Healthy, term, AGA babies

- Support breast feeding
- No routine blood glucose monitoring



Clinical recommendations Very preterm or sick babies

- Regular glucose monitoring
- Milk or IV glucose to maintain BG >2.5mmol/l
- Expressed breast milk as tolerated

Clinical recommendations Small or vulnerable babies

- Support breast feeding
- Maintain rigorous clinical monitoring
- Expressed breast milk or formula supplements according to clinical signs and BG



And above all: Identify and document

- Risk factors
- Coexisting conditions
- Clinical signs/ normality
- Accurate blood glucose measurements
- Response to treatment
- Investigations for underlying pathology