SMALL BUT MIGHTY WARRIORS



XXIV. NEONATOLOGICKE SETKANI XVIII. HANAKOVY DNY

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DisclosureOptimism is just a lack of information

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"He's right, when you look at it that way, it's not so bad!"

- 1935 AAP preterm infant < 2500g (standard adopted in Europe in 1919)
- In the mid 20th century gestational age and crown-to-heel length advocated as markers of viability
- 1950's RDS described to the RCOG and identified as a principal cause of death in infants younger than 37/40
- This led to distinction between premature and growth restricted infants

- 60's start of positive pressure ventilation, in 70's improving outcomes in infants less than 1800g
- $1973 \text{viability in US defined as } 28^{+0}/40$
- $1978 1^{st}$ infant below 750g ventilated
- 80's survival of infants born between 24⁺⁰ and 25⁺⁶/40 expected possibility in regional units and limits of viability moving towards 24 weeks of gestation
- 90's surfactant and antenatal steroids survival improved in infants 24 to 32 weeks of gestation, long term outcomes more and more important

- Late 90's and 21st century— reports of survival at 23/40 and indeed at 22/40 weeks of gestation
- Infants too immature to survive, and thus **provision of intensive care is** unreasonable at <23 weeks and <500 g, respectively (Seri I, Evans J. J Perinatol. 2008)
- Infants born at > or =25 weeks' gestation and with a birth weight of > or =600 g are warrant initiation of intensive care (the majority of these patients survive, and at least 50% do so without severe long-term disabilities) (Seri I, Evans J. J Perinatol. 2008)
- For infants born between 23(0/7) and 24(6/7) weeks' gestation and with a birth weight of 500 to 599 g, survival and outcome are extremely uncertain ('Grey Zone')
- 'Grey zone' of infant viability, the line between patient autonomy and medical futility is blurred, and medical decision-making is complex and needs to embrace careful consideration of several factors. (Seri I, Evans J. J Perinatol. 2008)

- There is no sharp limit of development, age, or weight at which a human fetus automatically becomes viable
- A baby's chances for survival increases 3-4% per day between 23 and 24 weeks of gestation and about 2-3% per day between 24 and 26 weeks of gestation
- After 26 weeks the rate of survival increases at a much slower rate because survival is high already
- The **limit of viability** was the gestational age at which a prematurely born fetus/infant has a **50% chance** of long-term survival outside its mother's womb
- Currently the limit of viability is considered to be around 24 weeks in the most developed countries

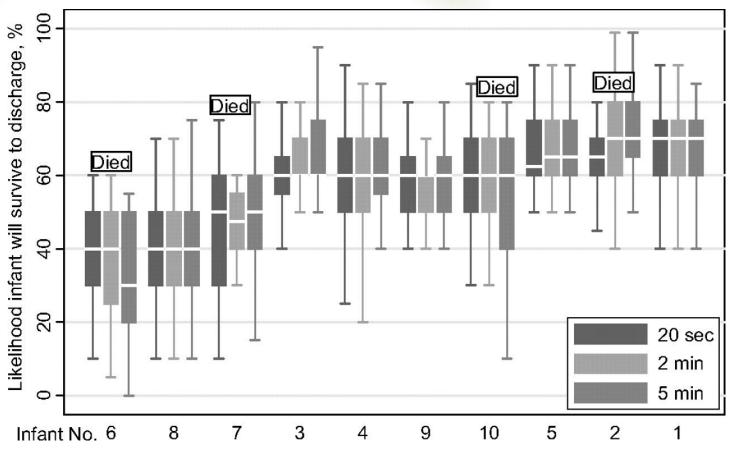
Limits of Viability – Call the 'Paeds'





Limits of Viability - Call the 'Paeds'

Predicted chance of survival to discharge (%) by attending neonatologists and fellows (combined group) at 20 seconds, 2 minutes, and 5 minutes for all infants, shown in ascending order of median.



Limits of Viability – Call the 'Paeds'

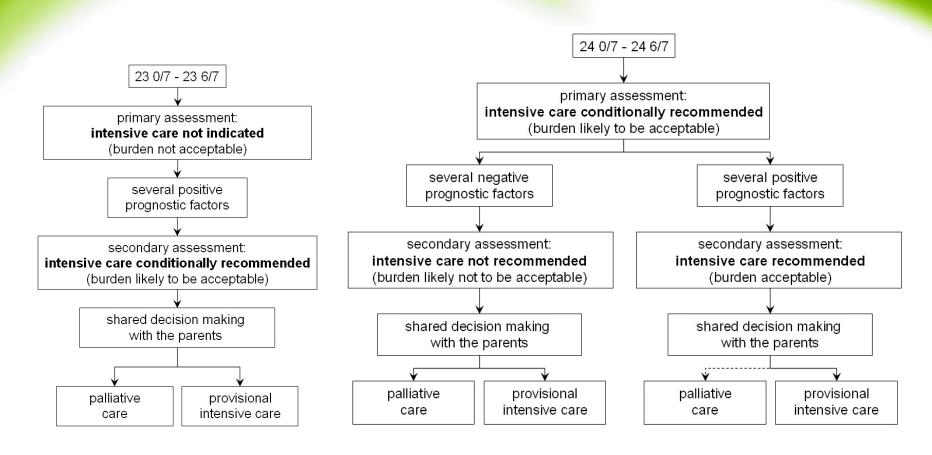
Message from the 'Paeds'

We do not have a clue about gestation or survival when called to extremely preterm delivery

But we can provide either intensive or comfort care based on best obstetric and parental information



Limits of Viability - Guidelines



Swiss Society of Neonatology (2011)

My name is Tommy.

I was born at 24 weeks and weighed 11b 9oz.

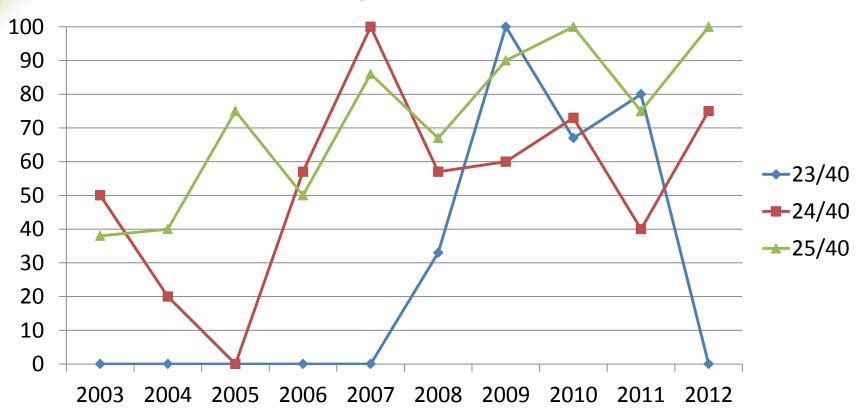
When I grow up, I wan to be a rugby player!





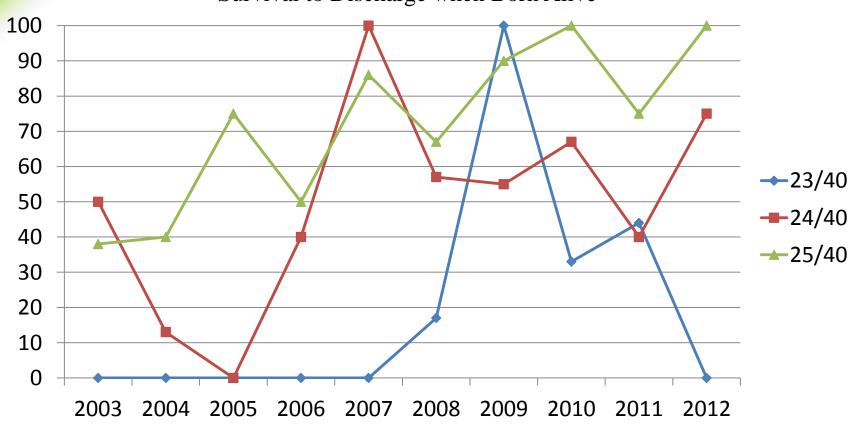
CWIUH – Results in Grey Zone Inhorn Infants

Survival to discharge when Intensive Care Started



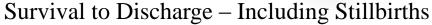
CWIUH – Results in Grey Zone Inhorn Infants

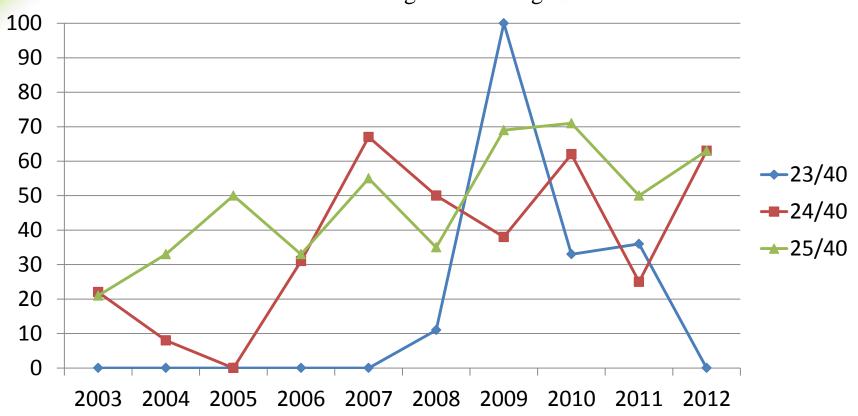
Survival to Discharge when Born Alive



CWIUH – Results in Grey Zone

Inborn Infants





CWIUH – Results in Grey Zone Inhorn Infants

Based on outcomes at 25/40

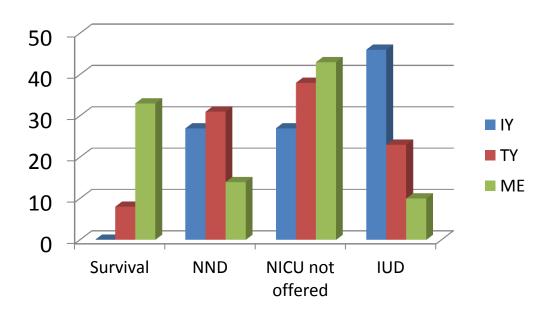
- Initial Years 2003 2005
- Transitional Years 2006 2008
- Modern Era 2009 2012



CWIUH – Results in Grey Zone

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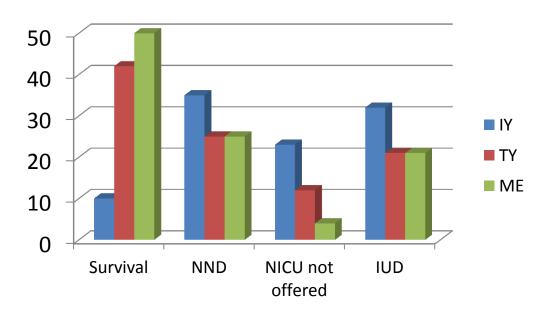
23/40	Initial Years	Transitional Years	Modern Era
Number of Deliveries	11	13	21
Survival (%)	0 (0)	1 (8)	7 (33)
Neonatal Deaths after Starting Intensive Care (%)	3 (27)	4 (31)	3 (14)
Intensive Care not Started (%)	3 (27)	5 (38)	9 (43)
Stillbirths (Intra/Ante partum deaths) (%)	5 (46)	3 (23)	2 (10)



CWIUH – Results in Grey Zone

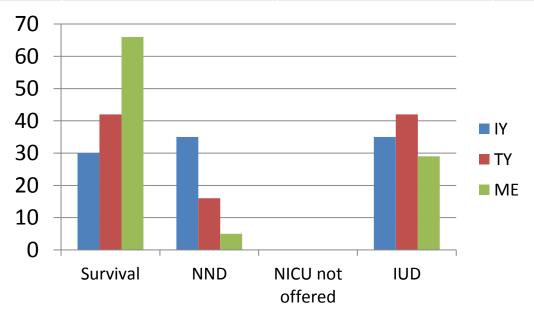
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24/40	Initial Years	Transitional Years	Modern Era
Number of Deliveries	31	24	56
Survival (%)	3 (10)	10 (42)	28 (50)
Neonatal Deaths after Starting Intensive Care (%)	11 (35)	6 (25)	14 (25)
Intensive Care not Started (%)	7 (23)	3 (12)	2 (4)
Stillbirths (Intra/Ante partum deaths) (%)	10 (32)	5 (21)	12 (21)



CWIUH – Results in Grey Zone Inborn Infants

25/40	Initial Years	Transitional Years	Modern Era
Number of Deliveries	26	31	41
Survival (%)	8 (30)	13 (42)	27 (66)
Neonatal Deaths after Starting Intensive Care (%)	9 (35)	5 (16)	2 (5)
Intensive Care not Started (%)	0 (0)	0 (0)	0 (0)
Stillbirths (Intra/Ante partum deaths) (%)	9 (35)	13 (42)	12 (29)



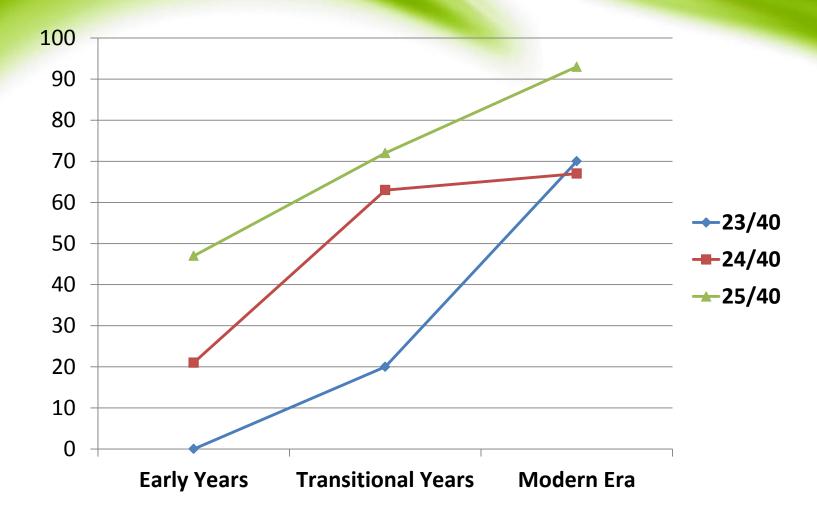
CWIUH – Results in Grey Zone Inborn Infants

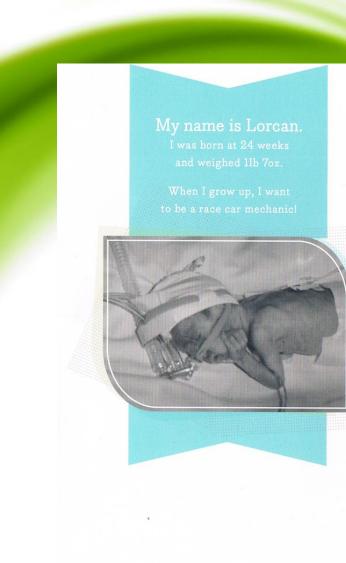
23/40	Initial Years	Transitional Years	Modern Era
Number of Infants NICU Started	3	5	10
Survival (%)	0 (0)	1 (20)	7 (70)

24/40	Initial Years	Transitional Years	Modern Era
Number of Infants NICU Started	14	16	42
Survival (%)	3 (21)	10 (63)	28 (67)

25/40	Initial Years	Transitional Years	Modern Era
Number of Infants NICU Started	17	18	29
Survival (%)	8 (47)	13 (72)	27 (93)

CWIUH – Results in Grey Zone Inborn Infants



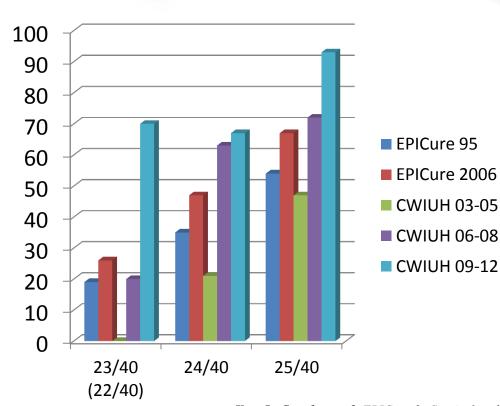




European Perspective (UK and Ireland)

EPICure study 1 and 2

GA at birth	22-23w	24w	25w
Admissions 1995 n	130	244	290
Survival (n) (%;95%CI)	25 (19 :12- 26)	85 (35 :29-41)	156 (54 :48- 59)
Admissions 2006 n	196	330	426
Survival (n) (%;95%CI)	51 (26 ;20-32)	155 (47 ;42- 52)	285 (67 ;62-71)
Survival change % (96%CI)	+7 (-2to16)	+12 (4-20)	+13 (6-21)
Chi-squared p	0.1	0.004	<0.001

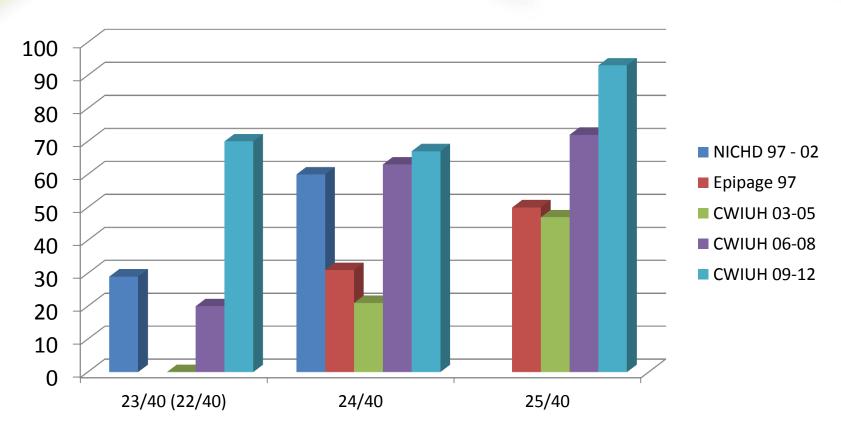


Kate L. Costeloe et.al. EPICure 2: Survival and Early Morbidity of Extremely Preterm Babies in England: Changes Since 1995. Pediatric Academic Societies and Asian Society for Pediatric Research Joint Meeting 2008, Hawaii, USA.

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International Perspective (France and USA)

Epipage Study and NICHD

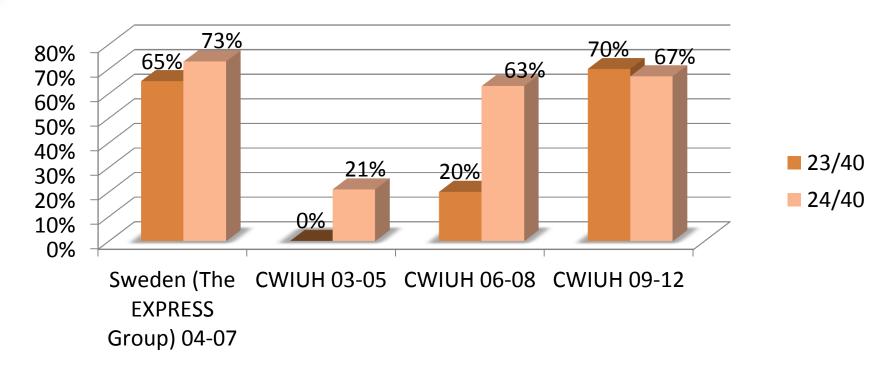


Larroque B et al. Survival of very preterm infants: Epipage, a population based cohort study. Arch Dis Child Fetal Neonatal Ed. 2004 Mar;89(2):F139-44

Fanaroff AA et al. Trends in neonatal morbidity and mortality for very low birthweight infants. Am J Obstet Gynecol. 2007 Feb;196(2):147.e1-8

Swedish Perspective

- CWIUH only inborn infants
- Sweden survival to 365 days
- All babies admitted to NICU (offered full intensive care)



One-Year survival of Extremely Preterm infants After Active Perinatal Care in Sweden, JAMA 2009 (The EXPRESS Group)

Swedish Perspective

- In infants born at 22 to 26 weeks of gestation, increased intensity of perinatal care reduced 1-year mortality in fetuses alive at the mother's admission for delivery
- Death or survival with NDI at 2.5 years' CGA was reduced in health care regions with higher intensity compared with regions with lower intensity of perinatal care
- Increased survival was not associated with increased neonatal morbidity or rate of NDI
- These findings, combined with the knowledge that survival cannot be predicted by initial appearance at birth, support a proactive approach to perinatal management of the EPT infant

Summary

- There is no exact threshold for 'limit of viability'
- Early ultrasound dating scan is better than any postnatal assessment
- Survival is improving in the extremely premature infants 23, 24, 25 weeks of gestation (even in the Irish level III centers CWIUH)
- Survival at 24/40 is consistently above 60%
- Survival at 25/40 is well above 70%, reaching 90%
- When care started at 23/40 survival seems to be above 50%
- Neurodevelopmental impairment at 2.5 yrs of age in infants between 22 to 26 weeks of gestation is 30% (based on Sweden data from 2004-2007 EXPRESS study)
- Proactive management and antenatal steroids are linked with improved survival without increase in NDI (and in fact with some decreased morbidity)
- It seems reasonable to offer NICU care to infants from 23+0/40 however that includes active perinatal care

Acknowledgements

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