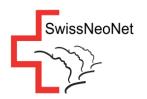
Mark Adams, PhD Swiss Neonatal Network & Follow-up Group

Patent Ductus Arteriosus Ligation – What is the ideal rate?

WIC Fall Research Meeting Oxford, October 2-4, 2019







#### Patent Ductus Arteriosus (PDA)

2 Options:

- 1. Medical: Indomethacin, Ibuprofen, Paracetamol
- 2. Surgical ligation

"Despite ... research and clinical experience ... over 6 decades, there is still uncertainty and controversy about the significance, evaluation and management of PDA in preterm infants."

AAP Committee on Fetus and Newborn

Pediatrics Dec 2015. doi: 10.1542/peds.2015-3730







#### **PDA Ligation in Swiss Neonatal Network**

All 9 Swiss NICUs

- 2012 2016
- < 28 weeks gestation</li>
- N = 1143

A: 5 NICUs <= 10% Ligation (range: 0% – 9.8%)

B: 4 NICUS > 10% Ligation

(range: 13.8% – 23%)







For illustration purposes only:

the real unit distribution is disclosed

#### Compare outcome group A versus B:

Odds of group A for:

- mortality
- severe morbidity (NEC Bell's stage >= 2, Moderate to severe BPD, ROP stage >= 3)
- moderate to severe NDI at 2 years of age
- Risk adjustment is made for:
  - Gestational age, birth weight z-score, male sex, outborn, early onset sepsis, severe IVH







# Outcome comparison group A versus B:

Adjusted OR for adverse outcome in units with lower proportion of PDA ligation

		All infants			Infants with treated PDA only			
Outcome	OR	95% CI	AUC	OR	95% CI	AUC		
Child died at any time	1.007	(0.711-1.425)	0.82	0.992	(0.628-1.568)	0.82		
Death or major morbidity	0.737	(0.563-0.965)	0.78	0.659	(0.465-0.934)	0.75		
Major morbidity	0.695	(0.517-0.936)	0.72	0.604	(0.416-0.877)	0.69		
Moderate to severe NDI at 2 years	1.135	(0.708-1.818)	0.67	1.372	(0.732-2.574)	0.73		

Association of outcome with PDA ligation?

- Did group B have sicker infants requiring PDA ligation?
- Did group B have sicker infants because of PDA ligation?







# Outcome of infants with PDA ligation:

Adjusted OR for adverse outcome of infants with PDA ligation

	All infants			Infants with treated PDA only			
Outcome	OR	95% CI	AUC	OR	95% CI	AUC	
Child died at any time	0.253	(0.133-0.48)	0.82	0.291	(0.15-0.563)	0.81	
Death or major morbidity	2.075	(1.356-3.175)	0.78	1.824	(1.176-2.83)	0.75	
Major morbidity	3.219	(2.088-4.962)	0.72	2.591	(1.652-4.065)	0.69	
Moderate to severe NDI at 2 years	2.151	(1.186-3.899)	0.67	2.471	(1.268-4.816)	0.73	

Association of outcome with PDA ligation?

- Are infants with PDA ligation sicker requiring PDA ligation?
- Are infants with PDA ligation sicker because of PDA ligation?







#### Outcome of infants with PDA ligation:

Adjusted OR for adverse outcome of infants with PDA ligation

		All infants			Infants with treated PDA only			
Outcome	OR	95% CI	AUC	OR	95% CI	AUC		
Child died at any time	0.564	(0.265-1.201)	0.78	0.495	(0.227-1.076)	0.78		
Death or major morbidity	2.69	(1.759-4.114)	0.74	2.175	(1.4-3.38)	0.71		
Major morbidity	3.219	(2.088-4.962)	0.72	2.591	(1.652-4.065)	0.69		
Moderate to severe NDI at 2 years	2.151	(1.186-3.899)	0.67	2.471	(1.268-4.816)	0.73		

Subgroup analysis of all infants that survived until day 10 of life.







# Summary

- Supply sensitive variation of PDA ligation (0% 23%)
- Infants born in group A have lower odds for major morbidities (0.5, CI: 0.33 to 0.75)
- Infants with PDA ligation have higher odds for major morbidities (2.6, CI: 1.58 to 4.25)
- Possible lower mortality in infants with PDA ligation (0.49, CI: 0.23 to 1.07)







# How to improve quality?

- Study of PDA ligation:
  - Current: differentiation into pre and post-ligation morbidities
  - Publication in peer-reviewed journal
- Raise awareness, discuss, monitor
  - First presentation of results in Spring meeting
  - In-depth analysis of procedures in Fall meeting:
    - Director of unit with lowest rate presents local approach
    - Director of unit with highest rate presents local approach modelling his/her presentation on that of unit with lowest rate
    - Local adjustment of SOP?
    - Long term monitoring in online evaluation and SwissNeoNet Annual Report







#### SWISSNEONET CORE

Network Coordinator Mark Adams

Follow-up Coordinator Giancarlo Natalucci

Steering Committee: Dirk Bassler Sven Schulzke Philipp Meyer

President past: Hans Ulrich Bucher Assistant: Tanja da Cunha Audit: Sue Behre

Institute for Biostatistics, Epidemiology and Prevention Zurich: Milo Puhan, PhD supervisor Viktor VonWyl, PhD advisor Julia Braun, statistics advisor







# SwissNeoNet Collaborators

Aarau: Cantonal Hospital Aarau, Children's Clinic, Department of Neonatology (Ph. Meyer, C. Anderegg), Department of Neuropaediatrics (A. Capone Mori, D. Kaeppeli); Basel: University of Basel Children's Hospital (UKBB), Department of Neonatology (S. Schulzke), Department of Neuropaediatrics and Developmental Medicine (P. Weber); Bellinzona: San Giovanni Hospital, Department of Pediatrics (G.P. Ramelli, B. Simonetti Goeggel); Berne: University Hospital Berne, Department of Neonatology (M. Nelle), Department of Pediatrics (B. Wagner), Department of Neuropaediatrics (M. Steinlin, S. Grunt); Biel: Development and Pediatric Neurorehabilitation Center (R. ); Chur: Children's Hospital Chur, Department of Neonatology (T. Riedel), Department of Neuropa di real di real di Killer); Fribourg: Cantonal Hospital Fribourg, Department of Neuropediatrics (K. Fuhrer); Lausake: rsinglespital (CHUV), Department of Neonatology (J.-F. Tolsa, M. Roth-Kleiner), Department of Development M. Bickle-Graz); Geneva: Department of child and adolescent, University Hospital (HUG) Ne al of Cate E. Pfister), Division of Development and Growth (P. S. Huppi, C. Borradori-Tolsa); Line him and Paediatric Intensive Care Unit (M. Stocker), Department of Neuropedia ic 1. 59//mt-Mechelke, F. Bauder); Lugano: Regional Hospital Lugano, Department of Pediatrics (V. Pezzoli) Muene Mugen: Cantonal Hospital Muensterlingen, Department of Pediatrics (B. Erkert A. Mueller); Neuchatel: Canton Tospital Neuchatel, Department of Pediatrics (M. Ecoffey); St. Gallen: Cantonal Hospital St. Gallen, Department of Neonatology (A. Malzacher), Children's Hospital St. Gallen, Neonatal and Paediatric Intensive Care Unit (J. P. Micallef), Department of Child Development (A. Lang-Dullenkopf); Winterthur: Cantonal Hospital Winterthur, Department of Neonatology (L. Hegi), Social Pediatrics Center (M. von Rhein); Zurich: University Hospital Zurich (USZ), Department of Neonatology (D. Bassler, R. Arlettaz), University Children's Hospital Zurich, Department of Neonatology (V. Bernet) and Child Development Centre (B. Latal, G. Natalucci).



