**Conclusions** The mean number of painful procedures per NICU patient per day declined and analgesic treatment changed to a more tailored or individualized approach. Non-pharmacological pain- or stress reducing strategies like NIDCAP and sucrose were fully embedded in our pain management. As further reduction of the number of painful procedures is unlikely we should explore newer pharmacological agents and apply non-pharmacological interventions more frequently.

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## PREDICTING THE EFFICACY OF ORAL SUCROSE IN REDUCING PAIN DURING OPHTALMOLOGICAL EXAMINATION FOR RETINOPATHY OF PREMATURITY: A PROSPECTIVE RANDOMISED STUDY

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**Background and aim** Retinopathy of prematurity (ROP) is one of the major morbidity among preterm infants. Although, local anesthetics reduce pain to some extent, eye examination still remains as a painful procedure. We aimed to evaluate the effect of oral sucrose combined with local anesthetics for pain relief during ophthalmological examination.

**Method** A total of forty patients under 32 weeks of gestational age were included in the study. Infants were randomly assigned to receive either oral sucrose solution (Group-1; n=21) or sterile water (Group-2; n=19) combined with topical proparacaine hydrochloride two minutes before examination. Pacifier was used in all patients as non-nutritive sucking during the study. Pain score was evaluated by premature infant pain profile (PIPP) scale. Each infant was video-recorded during and after the procedure.

**Results** Both groups were similar in terms of gestational age, birth weight, postnatal age and actual weight. There was no significant difference between groups in behavioral state, heart rate and oxygen saturation before the examination. At speculum insertion, heart rate variability was similar in both groups whereas oxygen desaturation was apparent in Group-2 (Group-1: 1.7±0.8 and Group-2: 2.5±0.6, p=0.001) and PIPP scores were also lower in Group-1 (Group-1: 14.5±1.8 and Group-2: 17.2±1.7, p=0.001). Total time of crying was significantly shorter in Group-1 (Group-1: 58.8±12.1 and Group-2: 96.3±24, p=0.001).

**Conclusion** Procedural pain is known to have acute and even long term negative. behavioral and developmental effects in neonates. In our study, use of sucrose in addition to local anesthetics during ophthalmological examination is shown to attenuate pain.

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## MORPHINE PREMEDICATION FOR INTUBATION IN PRETERM INFANTS - A PHARMACOKINETIC AND PHARMACOGENETIC REPORT

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**Background and aims** Morphine is used in preterm infants, but data are scars on pharmacokinetics (PK), pharmacodynamics (PD) and pharmacogenetics (PG). We aimed to study PK and PD/PG relation of morphine in infants included in a RCT comparing morphine with short-acting analgosedation as premedication for intubation.

**Methods** 17 preterm infants with a median (IQR) gestational age of 26.6 (25.1–28.7) w, birth weight 924 (721–1240) g and postnatal age 136 (17.5–322) h were randomized to receive morphine (0.3 mg/kg).

Blood samples for morphine, M6G and M3G concentrations were collected before administration, 20 min, 6 and 24 h after intubation. DNA was isolated from salivary swabs to genotype 18 polymorphisms in 12 genes using Taqman assays. Pain assessment (ALPS-0 and EDIN scales) was performed and additional morphine boluses were offered accordingly. The morphine level/pain score relation and the genotype influence on time to achieve a low pain score was calculated.

**Results** In infants receiving no additional doses, clearance was 1.5–3.3 ml/kg/min in 5 infants of 5–34 h and 9.9 in one infant of 332 h postnatal age.

Both morphine and morphine+M6G/5 correlated with mean ALPS-0 score at 6h (p=0.02 and 0.04) and 24 h (p=0.01 and 0.02). The COMT rs4680G>A (Val158Met) SNP was significantly correlated with time to reach the lowest pain score. COMT rs4680A patients experienced a faster response to opioids compared to rs4680GG patients in both groups (p=0.001 and p=0.072).

**Conclusions** Morphine clearance is dependent on postnatal age in premature infants. Genotyping would improve individual dosing of opioids during NICU-care.

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## ANALGESIA MODULATES CORTICAL RESPONSE TO PAINFUL PROCEDURES IN EXTREMELY PRETERM NEWBORN INFANTS

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**Introduction** Proper pain management in the neonatal intensive care unit (NICU) is essential. The clinical pain assessment depends on objective measurement of indirect behavioural and physiological pain indicators. Cortical pain processing has been observed in preterm infants from 24 weeks gestational age but no study has focused exclusively on cortical pain response in extremely premature infants ( $\leq$ 28 weeks gestational age).

**Aims** This study aimed to demonstrate cortical pain processing in extremely preterm infants. Furthermore, the study aimed to investigate the impact of analgesic drugs on cerebral haemodynamics and the relationship between behavioural and cortical pain responses.

Material and Methods A clinical study was performed in the NICU, including preterm infants ≤28 weeks gestational age or with a birth weight ≤1500 grams. Patients with severe on-going intraventricular haemorrhage or hydrocephalus were excluded. The infants were studied during routinely performed venepunctures and endotracheal tube suctions. Near-infrared spectroscopy was used for the study of cortical activity, parallel to observation of systemic haemodynamics and pain assessment with the *Premature Infant Pain Profile* and *Échelle Douleur Inconfort Nouveau-Né*.

**Results** During the procedures significant increases in the cerebral concentration of oxygenated haemoglobin were observed bilaterally during venepuncture and unilaterally during endotracheal tube suctions. Simultaneously, minor alterations in systemic haemodynamics occurred. The cerebral pain response was significantly reduced by analgesia.

**Conclusions** These results indicate the existence of cortical pain processing in very premature infants. Analgesia reduces this cortical response.

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## IMAGING TOOLS TO ASSESS THE DEVELOPMENT OF CORTICAL MORPHOLOGY AND CONNECTIVITY

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