Sucrose for analgesia in newborn infants undergoing painful procedures

Vu Thi Hieu Neonatal department Children Hospital No2

Overview

- Pain in newborns- pain assessment
- Painful procedures
- Pain relief- "analgesia"
- Sucrose in painful procedures
- Analgesia practice in NICU and Neonatal department

Pain in newborns



We think the newborns DON'T feel pain...

Neonatal pain vs adult pain





... or less than adults

KIẾN THỰC VÀ THỰC HÀNH ĐIỀU TRỊ GIẨM ĐAU KHI THỰC HIỆN THỦ THUẬT Ở TRỂ SƠ SINH CỦA ĐIỀU DƯỚNG TẠI KHOA SƠ SINH VÀ HỎI SỨC SƠ SINH BỆNH VIỆN NHI ĐỒNG 2

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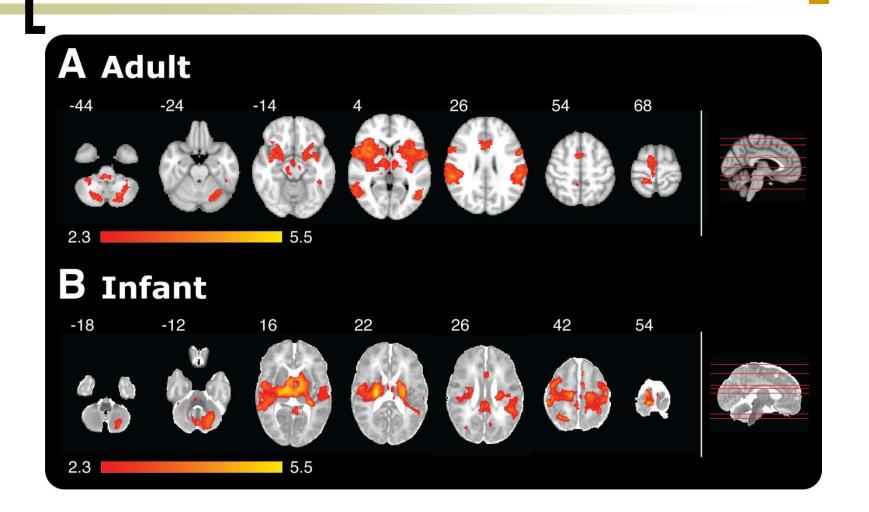
TÓM TẮT

Mục tiêu: Xác đinh tỉ lệ nhân viên khoa Sơ Sinh (SS) và Hồi Sức Sơ Sinh (HSSS) có kiến thức, và thực hành đúng trong thực hành điều trị giảm đau ở trẻ sơ sinh tại khoa SS và HSSS Bệnh viện Nhi Đồng 2. So sánh mối tương quan giữa kiến thức đúng và thực hành đúng.

Phương pháp nghiên cứu: Cắt ngang mô tả

Kết quả: Qua nghiên cứu, có sự tương đồng về phân bố nhân sự giữa hai nhóm ≤ 5 năm và > 5 năm kinh nghiệm.

Khi tiến hành khảo sát trên 106 điều dưỡng đang làm việc tại khoa SS và HSSS, trên 96% nhận định việc kiểm soát đau ở trẻ SS là quan trọng. Tuy nhiên, 93,4% cho rằng trẻ SS không đau hoặc đau ít hơn trẻ lớn, 52,8% cho rằng đau không ảnh hưởng ngắn hạn đến sinh lý và hành vi cũng như dài hạn đến sự phát triển của não và tâm lý trẻ sơ sinh. Tất cả điều đồng ý đặt Catherter, chích động - tĩnh mạch, CDTS là những thủ thuật gây đau cho trẻ, trên 90% đồng ý lấy máu, thay băng vết thương, lấy dextrostix, đặt NKQ cũng gây đau không kém. Trên 70% điều dưỡng nắm rõ các phương pháp giảm đau cao nhất là surcrose (98,1), emla (97,2%).



Pain assessment in newborns

Tool	Parameters	Score	Utility
Premature infant pain profile (PIPP) (24)	Gestational age, behavioral state, heart rate, oxygen saturation, brow bulge, eye squeeze, nasolabial furrow	Total: 0–21 each parameter scored 0–3; ≤6 minimal pain; >12 moderate to severe pain	Procedural and postoperative pain
FLACC (25)	Face, legs, activity, cry, consolability	Total: 0–10 each parameter scored 0–2; >4 moderate pain; >7 severe pain	Procedural and postoperative pain
COMFORT scale (behavioral and physiological parameters) (26)	Alertness, calmness, respiratory distress, movement, muscle tone, facial tension, blood pressure, heart rate	Total: 8–40 each parameter scored 1–5; 17–26 adequate sedation; ≥27 inadequate sedation/analgesia	Pain and sedation in NICU
COMFORT behavior scale (27)	Alertness, calmness, respiratory response (ventilated neonate) or crying (not ventilated), movement, muscle tone, facial expression	Total: 8–30 each parameter scored 1–5; >17 moderate pain requiring intervention	Postoperative pain in NICU

Pain assessment in newborns

NIPS

Neonatal Infant Pain Scale									
NIPS	0 point	1 point	2 points						
Facial expression	Relaxed	Contracted	-						
Cry	Absent	Mumbling	Vigorous						
Breathing	Relaxed	Different than basal							
Arms	Relaxed	Flexed/stretched	-						
Legs	Relaxed	Flexed/stretched							
Alertness	Sleeping/calm	Uncomfortable	-						

Pain assessment in newborns

N-PASS: Neonatal Pain, Agitation and Sedation Scale

Assessment	Sed	ation	Normal	Pain / Agitation			
Criteria	-2	-1	0	1	2		
Crying Irritability	No cry with painful stimuli	Moans or cries minimally with painful stimuli	Appropriate crying Not irritable	Irritable or crying at intervals Consolable	High-pitched or silent-continuous cry Inconsolable		
Behavior State	No arousal to any stimuli No spontaneous movement	Arouses minimally to stimuli Little spontaneous movement	Appropriate for gestational age	Restless, squirming Awakens frequently	Arching, kicking Constantly awake or Arouses minimally / no movement (not sedated)		
Facial Expression	Mouth is lax No expression	Minimal expression with stimuli	Relaxed Appropriate	Any pain expression intermittent	Any pain expression continual		
Extremities Tone	No grasp reflex Flaccid tone	Weak grasp reflex ↓ muscle tone	Relaxed hands and feet Normal tone	Intermittent clenched toes, fists or finger splay Body is not tense	Continual clenched toes, fists, or finger splay Body is tense		
Vital Signs HR, RR, BP, SaO₂	No variability with stimuli Hypoventilation or apnea	< 10% variability from baseline with stimuli	Within baseline or normal for gestational age	↑ 10-20% from baseline SaO ₂ 76-85% with stimulation – quick recovery ↑	↑> 20% from baseline SaOz ≤ 75% with stimulation - slow recovery ↑ Out of sync with vent		

The NPASS is a pain scale to assess pain in **neonates**.

- A score < 4 = mild pain (requires non-pharmacologic comfort measures).
- A score > 5 = moderate to severe pain (most likely requires pharmacologic intervention in conjunction with comfort measures)

Painful procedure

- 'can cause skin damage or mucosal damage by inserting or removing foreign bodies, and disturbing the body integrity of a neonate through therapeutic or diagnostic methods'.
- Ex: oral suctioning, tracheal suctioning, venipuncture, IV catheter insertion, heel lancing, wound treatment, ROP exam...

The frequency of painful procedures

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- 145 preterm baby, mean GA 35.4 ws, mean BW 2326g.
- An average 105.6 painful procedures were performed in each neonate during 2 weeks, and 7.5 daily.
- The number of painful procedures increases in lower GA babies.



PAIN

Acute effects

- Physiology and behavioral responses
 - Changes in stress hormones

Long-term effects

- Changing levels of neural activity can alter the normal development of the central nervous system
- Poorer cognitive and motor scores, impairments of growth, reduced white matter and subcortical gray matter maturation, altered corticospinal tract structure

Procedural pain and brain development in premature newborns

Objective: Preterm infants are exposed to multiple painful procedures in the neonatal intensive care unit (NICU) during a period of rapid brain development. Our aim was to examine relationships between procedural pain in the NICU and early brain development in very preterm infants.

Methods: Infants born very preterm (N = 86; 24–32 weeks gestational age) were followed prospectively from birth, and studied with magnetic resonance imaging, 3-dimensional magnetic resonance spectroscopic imaging, and diffusion tensor imaging: scan 1 early in life (median, 32.1 weeks) and scan 2 at term-equivalent age (median, 40 weeks). We calculated N-acetylaspartate to choline ratios (NAA/choline), lactate to choline ratios, average diffusivity, and white matter fractional anisotropy (FA) from up to 7 white and 4 subcortical gray matter regions of interest. Procedural pain was quantified as the number of skin-breaking events from birth to term or scan 2. Data were analyzed using generalized estimating equation modeling adjusting for clinical confounders such as illness severity, morphine exposure, brain injury, and surgery.

Results: After comprehensively adjusting for multiple clinical factors, greater neonatal procedural pain was associated with reduced white matter FA ($\beta = -0.0002$, p = 0.028) and reduced subcortical gray matter NAA/ choline ($\beta = -0.0006$, p = 0.004). Reduced FA was predicted by early pain (before scan 1), whereas lower NAA/ choline was predicted by pain exposure throughout the neonatal course, suggesting a primary and early effect on subcortical structures with secondary white matter changes.

Interpretation: Early procedural pain in very preterm infants may contribute to impaired brain development.

ANN NEUROL 2012;71:385-396

Neonatal procedural pain analgesia

	Procedures	Side effects
Opioid	Tracheal intubation Venipuncture, arterial puncture, IM or SC injection, LP in some cases Post operative in NICU	-Respiratory depression -Risk was increased by preterm birth and intercurrent comorbid conditions
Paracetamol	Ineffective: heel lancing	Overdose and hepatotoxicity
Regional analgesia EMLA	Venipuncture, arterial puncture, IM or SC injection, CVC insertion, LP Ineffective: heel lancing	Local reactions.
Non pharmacological analgesia	Venipuncture, arterial puncture, CVC insertion, LP	

Non pharmacological pain relief: sucrose

- Sucrose has been widely recommended for routine use in procedural pain in newborns.
- Mechanism of action: not precisely understood, maybe the sweet taste increase endorphines release.

- Advantages: no side effects, easy to use.
- Disadvantages: less effective in prolonged/ more intensive pain procedures.
- Reduce acute effects of pain (behavior responses); long- term effects: not precisely understood.



Cochrane Database of Systematic Reviews

Sucrose for analgesia in newborn infants undergoing painful procedures (Review)

Stevens B, Yamada J, Ohlsson A, Haliburton S, Shorkey A.

Sucrose for analgesia in newborn infants undergoing painful procedures.

Cochrane Database of Systematic Reviews 2016, Issue 7. Art. No.: CD001069.

DOI: 10.1002/14651858.CD001069.pub5.

www.cochranelibrary.com

Figure 4. Forest plot of comparison: 6 Heel lance: Sucrose (24%) + NNS vs. water + NNS, outcome: 6.2 PIPP 30 s after heel lance (term and preterm infants).

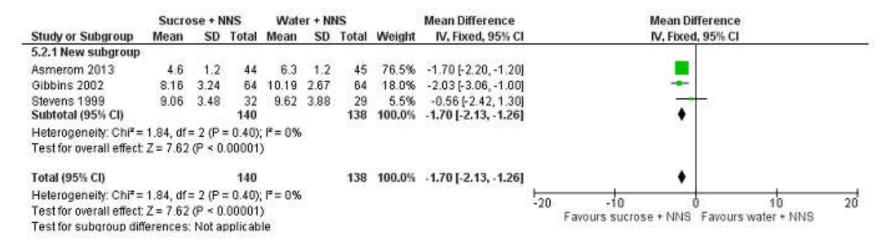


Figure 5. Forest plot of comparison: 6 Heel lance: Sucrose (24%) + NNS vs. water + NNS, outcome: 6.3 PIPP 60 s after heel lance.

	Sucro	se + N	NS	Wate	er + NI	NS		Mean Difference			Mean Difference	9	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% CI			IV, Fixed, 95% C	Ē.	
Gibbins 2002	8.78	4.03	60	11.2	3.47	59	79.3%	-2.42 [-3.77, -1.07]			and the second		
Stevens 1999	9.48	4.42	21	10.54	4.61	24	20.7%	-1.06 (-3.70, 1.58)			*		
Total (95% CI)			81			83	100.0%	-2.14 [-3.34, -0.94]			,		
Heterogeneity: Chi*= Test for overall effect					8				-100 Fa	-50 vours sucros	0 e + NNS Favour	50 s water + NNS	100

Figure 6. Forest plot of comparison: 18 Venipuncture: sucrose (24% to 30%) versus control (sterile water or no treatment), outcome: 18.1 PIPP score during venipuncture.

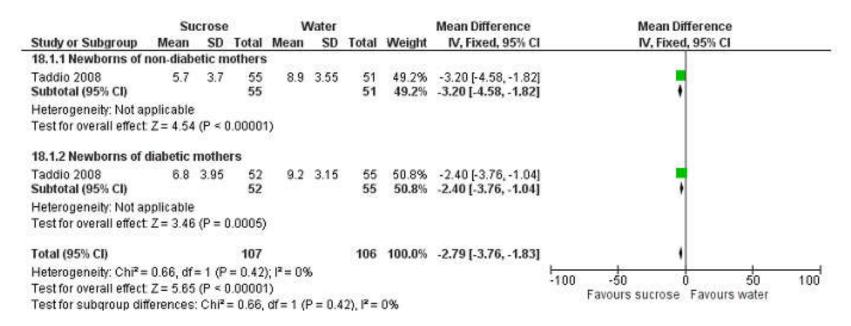


Figure 7. Forest plot of comparison: 23 Intramuscular injection (term infants): Sucrose (20-25%) vs. water or no intervention, outcome: 23.2 PIPP during IM injection (term infants).

	St	icrose	Ĕ	Water or	no interve	ntion		Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% CI	IV, Fixed, 95% CI
23.2.1 Infants of non	-diabetic	moth	ers	100000	2000000				
Taddio 2008 Subtotal (95% CI)	7.4	3.84	59 59	8.5	3.17	56 56	52.6% 52.6%	-1.10 [-2.38, 0.18] -1.10 [-2.38, 0.18]	
Heterogeneity: Not ap	pplicable	6							597-5870 P
Test for overall effect	Z=1.68	(P=0	0.09)						
23.2.2 Infants of diab	etic mo	thers							
Taddio 2008 Subtotal (95% CI)	6.2	3.84	59 59	7.2	3.62	58 58	47.4% 47.4%	-1.00 [-2.35, 0.35] -1.00 [-2.35, 0.35]	
Heterogeneity: Not ap	pplicable	š							
Test for overall effect	Z = 1.45	(P=	0.15)						
Total (95% CI)			118			114	100.0%	-1.05 [-1.98, -0.12]	-
Heterogeneity: Chi*=	0.01, df	= 1 (P	= 0.92)	$1^2 = 0\%$					- t - t - t - t - t - t - t - t - t - t
Test for overall effect				A10 - 1000					-4 -Z 0 Z 4
Test for subgroup dif				df = 1 (P = 0)	1.92), $P = 0$	%			Favours sucrose Favours water/no interv

Figure 8. Forest plot of comparison: 30 ROP examination: sucrose (24% to 33%) (sucrose or sucrose + NNS) versus control (water or water + NNS), outcome: 30.1 PIPP score during eye examination.

	Su	Sucrose Wat						Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% CI	IV, Fixed, 95% CI
30.1.1 Sucrose via s	yringe v	ersus	contro	l (sterile	wate	er via sy	yringe)		
Boyle 2006 Subtotal (95% CI)	14.3	1.6	10 10	15.3	1.9	10 10	21.7% 21.7%	-1.00 [-2.54, 0.54] -1.00 [-2.54, 0.54]	‡
Heterogeneity: Not as	pplicable	ğ							11
Test for overall effect	Z=1.27	(P = 0	0.20)						
30.1.2 Sucrose + pa	cifier ver	sus c	ontrol	(sterile	water	+ pacit	fier)		
Boyle 2006	12.1	3.4	11	12.3	2.9	9	6.7%	-0.20 [-2.96, 2.56]	+
Dilli 2014	13.7	2.1	32	16.4	1.8	32	55.9%	-2.70 [-3.66, -1.74]	
Mitchell 2004	8.8	2.71	15	11.4	2.32	15	15.7%	-2.60 [-4.41, -0.79]	*
Subtotal (95% CI)			58			56	78.3%	-2.47 [-3.27, -1.66]	• 1
Heterogeneity: Chi ² =	2.84, df	= 2 (P	= 0.24	$); 1^2 = 29$	%				
Test for overall effect	Z= 5.97	(P < (0.0000	1)					
Total (95% CI)			68			66	100.0%	-2.15 [-2.86, -1.43]	•
Heterogeneity: Chi ² =	5.56, df	= 3 (P	= 0.13); I ² = 46	%			Harris Harris Barbara Cherry Chi	20 10 0 10 20
Test for overall effect	Z = 5.88	(P < (0.0000	1)					-20 -10 0 10 20 Favours sucrose Favours water
Test for subgroup dif	ferences	Chi2	= 2.73.	df = 1 (F	$^{2} = 0.1$	$ 0\rangle, ^2 =$	63.3%		Lavoria artifas. Lavoria Matei

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Qua giám sát lâm sàng, 93% điều dưỡng tuân thủ thực hiện giảm đau khi CDTS, 26% có giảm đau khi chích động – tĩnh mạch và 29% lấy máu xét nghiệm. Tất cả các trường hợp đặt sonde dạ dày và lấy dextrostix đều không được giảm đau, 89% khi thay băng, 50% khi đặt catherter, 23% không giảm đau khi đặt NKQ. Tỉ lệ điều dưỡng tuân thủ thực hành kiểm soát giảm đau cho trẻ chiếm 30, 3%.

Conclusion

- Sucrose: effective analgesia in some neonatal painful procedures.
- Increase using pharmacological (regional analgesia: EMLA) and nonpharmacological (sucrose) during some painful procedures in newborns, esp preterms.

References

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