RESEARCH ARTICLE

Comparison of the Four Proposed Apgar Scoring Systems in the Assessment of Birth Asphyxia and Adverse Early Neurologic Outcomes

Hosein Dalili^{1,2}, Firouzeh Nili¹, Mahdi Sheikh^{1,2}*, Amir Kamal Hardani¹, Mamak Shariat^{1,2}, Fatemeh Nayeri^{1,2}

1 Maternal, Fetal and Neonatal Research Center, Vali-asr Hospital, Tehran University of Medical Sciences, Tehran, Iran, 2 Breastfeeding Research Center, Vali-asr Hospital, Tehran University of Medical Sciences, Tehran, Iran

* mahdisheikh@gmail.com

Abstract

Objectives

To compare the Conventional, Specified, Expanded and Combined Apgar scoring systems in predicting birth asphyxia and the adverse early neurologic outcomes.

Methods

This prospective cohort study was conducted on 464 admitted neonates. In the delivery room, after delivery the umbilical cord was double clamped and a blood samples was obtained from the umbilical artery for blood gas analysis, meanwhile on the 1-, 5- and 10- minutes Conventional, Specified, Expanded, and Combined Apgar scores were recorded. Then the neonates were followed and intracranial ultrasound imaging was performed, and the following information were recorded: the occurrence of birth asphyxia, hypoxic lschemic Encephalopathy (HIE), intraventricular hemorrhage (IVH), and neonatal seizure.

Results

The Combined-Apgar score had the highest sensitivity (97%) and specificity (99%) in predicting birth asphyxia, followed by the Specified-Apgar score that was also highly sensitive (95%) and specific (97%). The Expanded-Apgar score was highly specific (95%) but not sensitive (67%) and the Conventional-Apgar score had the lowest sensitivity (81%) and low specificity (81%) in predicting birth asphyxia. When adjusted for gestational age, only the low 5-minute Combined-Apgar score was independently associated with the occurrence of HIE (B = 1.61, P = 0.02) and IVH (B = 2.8, P = 0.01).



G OPEN ACCESS

Citation: Dalili H, Nili F, Sheikh M, Hardani AK, Shariat M, Nayeri F (2015) Comparison of the Four Proposed Apgar Scoring Systems in the Assessment of Birth Asphyxia and Adverse Early Neurologic Outcomes. PLoS ONE 10(3): e0122116. doi:10.1371/ journal.pone.0122116

Academic Editor: Jörn-Hendrik Weitkamp, Vanderbilt University, UNITED STATES

Received: January 2, 2015

Accepted: February 20, 2015

Published: March 26, 2015

Copyright: © 2015 Dalili et al. This is an open access article distributed under the terms of the <u>Creative Commons Attribution License</u>, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Data Availability Statement: Due to legal restrictions, data are available upon request. Interested researchers may access the data by contacting the data access committee of maternal, fetal and neonatal research center, Tehran University of Medical Sciences (Email: <u>mfnhrc@yahoo.com</u>).

Funding: The authors have no support or funding to report.

Competing Interests: The authors have declared that no competing interests exist.

Conclusions

The newly proposed Combined-Apgar score is highly sensitive and specific in predicting birth asphyxia and also is a good predictor of the occurrence of HIE and IVH in asphyxiated neonates.

Introduction

Asphyxia is a damaging condition of impaired blood gas exchange, leading, if it persists to progressive hypoxemia, hypercapnia, and tissue oxygen debt, which can cause serious central nervous system, respiratory, cardiovascular, and renal complications.[1]

The prevalence of fetal asphyxia is increasing; in 1990s the incidence of asphyxia was reported as 5.4 per 1000 live births,[2] while in the 2006 the American College of Obstetricians and Gynecologists (ACOG) reported an incidence of 25–73 per 1000 live births.[1]

Early newborn assessment, prediction of the neonatal complications, and prompt intervention is crucial, to prevent the progression of asphyxia and also not to loose the window of therapeutic opportunity for minimizing its complications.[3]

Apgar scoring system which was described in 1950s, is the oldest and most commonly used assessment tools for the evaluation of the newborn and the need for interventions in the delivery room.[4] In addition some studies used the Apgar score for the assessment of birth asphyxia and the prediction of adverse neonatal outcomes.[5-7]

The ACOG in the committee opinion on Apgar score, stated that a low Apgar score beyond 5 minutes is a suggestive criterion for asphyxia, however, there are several limitations with the Apgar score that make it inappropriate to be used alone for establishing the diagnosis of asphyxia, or for predicting the adverse neonatal outcomes.[8] Apgar score is influenced by the gestational age, neonate's maturity, drugs, etc. In addition the Apgar score that is assigned during resuscitation and intubation does not give a precise assessment of the newborn's situation. [9] These limitations made the researchers think of an alternative to the conventional Apgar score. Therefore the Specified-Apgar and later the Expanded-Apgar scores were suggested to allow the assessment of the newborn's condition independent of the interventions and the gestational age. [8,10,11] Despite these advances, there was still a need for a more comprehensive and precise scoring system that could predict the occurrence of adverse neonatal outcomes. Thus the Combined-Apgar score was proposed by Rüdiger et al. that consists of both the Specified and the Expanded Apgar scores to allow a more detailed description of neonate's postnatal condition.[12]

Despite all the attempts, there is still a considerable gap in this area of research; by our extensive search we could not find published studies evaluating the four proposed scoring systems and comparing them in the assessment of the newborns. No study has evaluated the sensitivity, specificity and the cutoffs of these scoring systems in assessing birth asphyxia and adverse neonatal outcomes especially after the corrections made to the limited conventional Apgar score. To date there is no accepted standards for evaluating the newborns (especially the preterm and the resuscitated newborns) under clinical conditions in the delivery room.[<u>8,13</u>]

We therefore conducted this study to compare the sensitivity and specificity of the four proposed Apgar scoring systems in the assessment of birth asphyxia and in predicting adverse early neurologic outcomes in the neonates especially after asphyxia and the hemodynamic instability of the newborns in the delivery room.



Table 1. The Conventional-Apgar scoring system as introduced by Virginia Apgar in 1953.

Sign	The Score					
	0	1	2			
Heart rate	Absent	Less than 100	More than 100			
Respiration	Absent	Slow, irregular	Good, Crying			
Muscle tone	Limp	Some flexion	Active motion			
Reflex Irritability	No response	Grimace	Cough, sneeze, cry			
Color	Blue or pale	Pink body, blue extremities	Completely pink			

doi:10.1371/journal.pone.0122116.t001

Materials and Methods

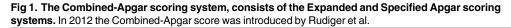
Study population and study design

This prospective cohort study was conducted on the newborns admitted to a tertiary referral hospital with an annual birth rate of 2200 births, between September 2012 and February 2014. Inclusion Criteria were: live birth at a gestational age > 25 weeks, birth within the study center, and the requirement for hospital admission. A total of 596 newborns were eligible to participate in the study, and 132 were excluded due to the following criteria: gestational age at \leq 25 weeks, birth out of the study center, major congenital anomalies, death in the delivery room, and missing parental informed consent. A written informed consent was taken from the parents before the delivery. A total of 464 neonates born at the gestational ages of 26–40 weeks completed the study which was approved by the Research Deputy and the Ethics Committee of the Tehran University of Medical Sciences.

Data and specimen collection

Upon enrollment, a standardized questionnaire was completed for every neonate through medical records and physical examinations. The questionnaire contained inquiries regarding the gestational age at birth as calculated based on the ultrasound imaging, type of delivery, birth weight, and neonatal gender. In the delivery room, after the neonate was delivered, the umbilical cord was double clamped, and immediately a blood sample was obtained from the umbilical artery using a heparinized syringe for arterial blood gas analysis, which was performed by a blood gas analyzer (Nova Biomedical, Waltham, MA), meanwhile on the 1-, 5- and 10- minutes the Conventional, Specified, Expanded, and Combined Apgar scores were recorded by educated physicians according to Table 1 and Fig. 1. [4,8,12] The conventional Apgar score ranges from 0–10, the Specified Apgar score ranges from 0–10, the Expanded Apgar score ranges from 0-7, and the Combined Apgar score ranges from 0-17. The physicians were all educated in the delivery room prior to the beginning of the study to assure the consistency and avoid interpersonal biases of calculating the four proposed Apgar scores. Then the neonates were followed by a neonatologist until discharge and the following information were recorded: the occurrence of birth asphyxia (that was diagnosed based on the following criteria: 1. a sentinel (signal) hypoxic event occurring immediately before or during labor, 2. evidence of a metabolic acidosis in fetal umbilical cord arterial blood obtained at delivery $(pH \le 7.00 \text{ and base deficit} \ge 12 \text{ mmol/L})$, 3. onset of multisystem involvement (central nervous, renal, pulmonary, cardiovascular, gastrointestinal systems) within 72 hours of birth, 4. exclusion of other identifiable etiologies, such as trauma, coagulation disorders, infectious conditions, or genetic disorders), [14,15] the occurrence and severity of Hypoxic Ischemic Encephalopathy (HIE) (that was diagnosed based on the following criteria: 1. a sentinel (signal) hypoxic event occurring immediately before or during labor, 2. evidence of a metabolic acidosis

		Minutes		\$
		1	5	10
С	Continuous Positive Airway Pressure a			
0	Oxygen			
M-B	Mask and Bag Ventilation b			
Ι	Intubation and Ventilation			
Ν	Neonatal Chest Compression			
Е	Exogenous Surfactant			
D	Drugs			
	Sum of Expanded Apgar			
	Scoring Each Item:			L
	0 = Intervention was performed ; $1 =$ No intervention was performed			
	a : score 0 if "Mask and Bag" or "Intubation and Ventilation" is score 0			
	b : score 0 if "Intubation and Ventilation" is scored 0			
А	Appearance (Skin Color)			
	2 = Completely pink			
	1 = Centrally pink with acrocyanosis			
	0 = Centrally blue or pale			
Р	Pulse (HR)			
	2 => 100 beats per minute			
	1 = < 100 beats per minute			
	0 = No heart beat			
G	Grimacing (Reflex)			
	2 = Appropriate for gestational age			
	1 = Reduced for gestational age			
	0 = No reflex response			
А	Activity (Muscle Tone)			
	2 = Appropriate for gestational age			
	1 = Reduced for gestational age			
	0 = No reflex response			
R	Respiration (Chest Movement)			
	2 = Regular chest movement			
	1 = Small of irregular chest movement			
	0 = No chest movement			
	Sum of Specified Apgar		-	



doi:10.1371/journal.pone.0122116.g001

in fetal umbilical cord arterial blood obtained at delivery (pH \leq 7.00 and base deficit \geq 12 mmol/L), 3. early onset of neonatal encephalopathy as detected by structured neurologic examination performed serially by a single investigating neonatologist at approximately 12, 36, and 72 hours after birth. 4. exclusion of other identifiable etiologies, such as trauma, coagulation disorders, infectious conditions, or genetic disorders), also the severity of HIE was recorded based on Sarnat chart (Table 2) [1,14–16], and the occurrence of neonatal seizure that was documented by an abnormal electroencephalogram (EEG) performed in case of clinical seizures and in every comatose infant at 16–48 hours after birth, which was not due to electrolyte imbalances or any identifiable infectious, hypoglycemic, or traumatic causes.[15] In addition intracranial ultrasound imaging was performed on all the neonates by a single person during the



Cavarity Chara 1 (Mild) Chara 2 (Madavata) Chara						
Severity	Stage 1 (Mild)	Stage 2 (Moderate)	Stage 3 (Severe)			
Level of consciousness	Hyperalert	Lethargic or Obtunded	Stupor or coma			
Activity	Normal	Decreased	Absent			
Neuromuscular Control						
Muscle Tone	Normal	Mild hypotonia	Flaccid			
Posture	Mild distal flexion	Strong distal flexion	Intermittent decerebration			
Stretch Reflexes	Overactive	Overactive	Decreased or absent			
Complex or primitive reflexes						
Suck	Weak	Weak or absent	Absent			
Moro (Startle)	Strong	Weak	Absent			
Tonic neck	Slight	Strong	Absent			
Autonomic Function						
Pupils	Mydriasis	Miosis	Variable			
Heart Rate	Tachycardia	Bradycardia	Variable			
Seizures	None	Common	Uncommon			

Table 2. The Sarnat chart for the staging of the severity of hypoxic ischemic encephalopathy as introduced by Sarnat H.B and Sarnat M.S. in 1976.

doi:10.1371/journal.pone.0122116.t002

first postnatal week to detect the occurrence of intraventricular hemorrhage (IVH) and its severity.

Statistical analysis

All statistical analyses were performed using SPSS statistical software (version 18.0.0: PASW, Chicago, IL). Chi-squared analysis, Fisher's exact test, independent-samples T test, One-way Anova, Pearson correlation coefficient, and ROC curve were used to analyze the correlations and relationships between variables. Multivariate logistic regression was used to evaluate the dependency of the obtained results. Sample size was calculated for a power of 80% and an alpha error of 0.05. Estimated odds ratios (ORs) with 95% confidence intervals (95% CIs) and P value < 0.05 were used to evaluate the statistical significance of the associations and correlations between variables.

Results

Descriptive statistics

This prospective cohort study was conducted on 464 admitted neonates who were born at gestational ages > 25 weeks. At enrollment the mean \pm standard deviation (SD) for gestational age at birth was 35.9 ± 3.2 weeks; birth weight was 2586 ± 866 grams; cord blood pH was 7.25 ± 0.09 ; cord blood bicarbonate was 19.2 ± 4 mEq/l; cord blood base deficit was 2.2 ± 4 mEq/l; Conventional-Apgar score at 5- minute was 8.8 ± 1.1 (Range 3–10); Specified-Apagr score at 5- minute was 8.6 ± 1.6 (Range 3–10); Expanded-Apgar score at 5-minute was 5.7 ± 1.6 (Range 0–7); and for Combined-Apgar score at 5- minute was 15 ± 3.1 (Range 4–17).

Of 464 neonates who completed the study, 246 were preterm (born at a gestational age < 37 weeks) (53%), 209 had low birth weight (birth weight < 2500 grams) (45%), 43 had birth asphyxia (9.2%), 29 had hypoxic ischemic encephalopathy (6.25%) that was mild in 6 neonates (0.9%), moderate in 14 (3%) and severe in 9 neonates (1.9%). IVH was detected in 41 of the neonates (8.8%), and seizure with an abnormal EEG occurred in 35 neonates (7.5%).



The Scoring System	рН	PO2	PCO2	HCO3	Base Defici
Conventional Apgar					
	r = 0.13	r = 0.07	r = - 0.11	r = 0.14	r = - 0.23
	P = 0.008 *	P = 0.14	P = 0.03 *	P = 0.7	P = 0.000 *
Expanded Apgar					
	r = 0.18	r = 0.13	r = - 0.11	r = - 0.05	r = - 0.29
	P = 0.000 *	P = 0.01 *	P = 0.03 *	P = 0.27	P = 0.000 *
Specified Apgar					
	r = 0.24	r = 0.11	r = - 0.16	r = - 0.04	r = 0.31
	P = 0.000 *	P = 0.02 *	P = 0.002 *	P = 0.37	P = 0.000 *
Combined Apgar					
	r = 0.31	r = 0.12	r = - 0.2	r = 0.02	r = 0.32
	P = 0.000 *	P = 0.01 *	P = 0.000 *	P = 0.61	P = 0.000 *

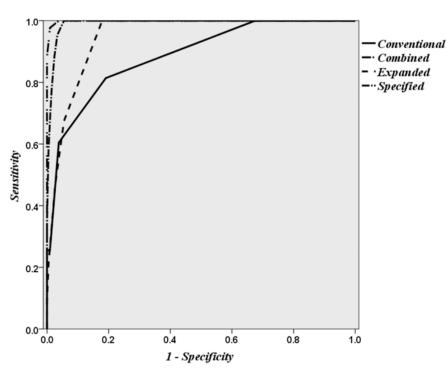
Table 3. Pearson correlation coefficient showing the association between different Apgar scores and cord blood gas parameters.

doi:10.1371/journal.pone.0122116.t003

Birth asphyxia, umbilical cord blood gas and the proposed scores

There was a significant linear correlation between all the proposed Apgar scores and the cord blood pH, and base deficit, and an inverse correlation with PCO2 levels. However the Combined-Apgar score had the strongest correlations with cord blood gas parameters.(<u>Table 3</u>)

Fig. 2 illustrates the sensitivity and specificity of the 5-minute proposed scores in predicting birth asphyxia by using a receiver operating characteristic (ROC) curve. Area under the curve





doi:10.1371/journal.pone.0122116.g002



	Sensitivity	Specificity	PPV	NPV	LR +	LR -	AUC	95%CI
Conventional Apgar score (cutoff less than 7)	81%	81%	73.3%	92.8%	4.26	0.23	0.887	0.83–0.94
Expanded Apgar score (cutoff less than 4)	67%	95%	55.7%	96.5%	13.4	0.34	0.949	0.92-0.97
Specified Apgar score (cutoff less than 7)	95%	97%	79.1%	98.7%	31.6	0.05	0.991	0.98–0.99
Combined Apgar score (cutoff less than 10)	97%	99%	93.1%	99.7%	97	0.03	0.999	0.99–1.00

Table 4. Comparison of the four proposed Apgar scores in predicting birth asphyxia.

PPV: Positive Predictive Value

NPV: Negative Predictive Value

LR +: Likelihood Ration for positive results

LR-: Likelihood Ration for negative results

AUC: Area Under the Curve

95%CI: 95% Confidence Interval

doi:10.1371/journal.pone.0122116.t004

for Conventional-Apgar was 0.88, for Expanded-Apgar was 0.94, for Specified-Apgar was 0.99 and for Combined-Apgar was 0.99.(Fig. 2)(Table 4)

Also we calculated the best cutoff for each 5- minute calculated scores in predicting birth asphyxia with the observed sensitivity, specificity, positive predictive value (PPV), and negative predictive value (NPV). (<u>Table 4</u>)

As illustrated in <u>Fig. 2</u>, and <u>Table 4</u>. the Combined-Apgar score has the highest sensitivity, specificity, PPV and NPV in predicting birth asphyxia.

15 neonates (3.2%) had a 5- minute Conventional-Apgar score < 7, 52 neonates (11.2%) had a 5-minute Expanded-Apgar score < 4, 46 neonates (9.9%) had a 5- minute Specified-Apgar score < 7, and 48 neonates (10.3) had a 5- minute Combined-Apgar score < 10.

The proposed scores in predicting adverse early neurologic outcome

In the asphyxiated neonates, among the proposed scores, the low 5- minute Combined-Apgar score (less than 10) (P = 0.01) and low 5- minute Expanded-Apgar score (less than 4) (P = 0.02) were the only scores associated with the occurrence of HIE, however neither could predict the severity of HIE. The low 5- minute Combined- Apgar score (P = 0.001) and low 5- minute Expanded-Apgar score (less than 4) (P = 0.01) were the only scores associated with the occurrence of IVH, however neither could predict the severity of IVH. None of the low 5- minute scores was associated with the occurrence of neonatal seizure. When adjusted for gestational age by using logistic regression model, only low 5-minute Combined-Apgar score remained significantly associated with HIE (B = 1.61, P = 0.02) and IVH (B = 2.8, P = 0.01).

Discussion

In the current study the newly proposed Combined-Apgar score was highly sensitive and specific in predicting birth asphyxia, with a high PPV and NPV. In addition a low 5- minute Combined-Apgar score was significantly associated with major adverse early neurologic outcomes in the asphyxiated neonates, which was independent of gestational age.

This study is among the very first studies that compare the four proposed scoring systems in assessing the newborns in the delivery room. In this study the Combined-Apgar score had the highest sensitivity and specificity in predicting birth asphyxia, followed by the Specified-Apgar score that was also highly sensitive and specific. The Expanded-Apgar score was highly specific but not sensitive and the Conventional-Apgar score had the lowest sensitivity and low specificity in predicting birth asphyxia. The low sensitivity and specificity of the Conventional-Apgar

scoring system in assessing birth asphyxia was recognized in many studies since 1980s; Skyes et al. in a study of 1210 newborn stated that because the Apgar score doesn't usually reflect the degree of acidosis at delivery, its value in asphyxial assessment must be questioned. [17] In addition due to the serious limitations of the Conventional-Apgar score especially in premature and resuscitated neonates, many papers criticized the use of the Apgar score as the gold standard in assessing asphyxia and neonatal outcomes. [18,8,9,13,14] Even a Lancet paper had called for the Apgar score to be pensioned off.[18] Neil Marlow in his paper indicated that in the absence of prematurity and neonatal resuscitation a low Apgar score may identify adverse neonatal outcome, therefore suggested using Apgar score with caution while continue to search for an alternative for the Apgar score that could be used in all the newborns.[19] To solve the limitations of the Coventional-Apgar score, two additional scoring systems were suggested; Rudiger et al. suggested the Specified-Apgar score, that contained the same but slightly changed items of the Conventional-Apgar score, it describes the condition of the newborn regardless of the medical interventions or the gestational age.[10] Further, for more precise neonatal evaluation the ACOG and AAP introduced the Expanded-Apgar score that counts the medical intervention given to the newborn in the delivery room.[8] In 2012 Rudiger et al. introduced the Combined-Apgar scoring system that is a combination of both Specified and Expanded Apgar scores, and documented that the Combined-Apgar score was superior to Specified and Expanded Apgar scores alone in predicting neonatal mortality in preterm infants.[12]

Based on the results of the current study for the first time cutoff points were suggested for the 5- minute Combined, Expanded and Specified Apgar scoring systems for the assessment of birth asphyxia and adverse neonatal outcome. A 5- minute Conventional-Apgar score < 10, Expanded-Apgar score < 4 and Specified-Apgar score < 7 were associated with birth asphyxia and severe neonatal acidosis.

In our study only a low 5-minute Combined-Apgar score could independently predict the occurrence of HIE and IVH in asphyxiated neonates, however it could not predict the severity of HIE or HIV. None of the Conventional, Specified, or Expanded Apgar scores alone could independently predict the adverse neurologic outcomes in asphyxiated neonates. This was in accordance with the ACOG and AAP committee opinion on the Conventional-Apgar score and also other studies that indicated a Conventional-Apgar score at 5- minute correlates poorly with future neurologic outcomes and therefore should not be used as a predictor of neurologic or adverse neonatal outcomes.[7,8,14,18–21]

Conclusions

This study shows that the newly proposed Combined-Apgar score that was introduced in 2012 by changing and combining the Conventional, Specified and Expanded Apgar scores, to solve the limitations of using each of these scores alone, is highly sensitive and specific in predicting birth asphyxia with a high PPV and NPV, and also is a good predictor of the occurrence of HIE and IVH in asphyxiated neonates.

Acknowledgments

The authors thank Dr. Elaheh Amini and Dr. Tahere Esmaeilnia for their contribution in data gathering.

Author Contributions

Conceived and designed the experiments: HD AKH M. Sheikh M. Shariat. Performed the experiments: HD AKH M. Sheikh M. Shariat F. Nili F. Nayeri. Analyzed the data: HD M. Sheikh

M. Shariat. Contributed reagents/materials/analysis tools: HD M. Sheikh M. Shariat AKH. Wrote the paper: HD M. Sheikh M. Shariat F. Nili F. Nayeri.

References

- Umbilical cord blood gas and acid-base analysis. ACOG Committee Opinion No. 348. American College of Obstetricians and Gynecologists. Obstet Gynecol 2006; 108:1319–22. PMID: <u>17077266</u>
- Thornberg E, Thiringer K, Odeback A, Milsom I. Birth asphyxia: incidence, clinical course and outcome in a Swedish population. Acta Paediatr. 1995; 84:927–32. PMID: <u>7488819</u>
- Whitelaw A, Thoresen M. Clinical trials of treatments after perinatal asphyxia. Curr Opin Pediatr. 2002; 14:664–8. PMID: <u>12436031</u>
- Apgar V. A proposal for a new method of evaluation of the newborn infant. Curr Res Anesth Analg. 1953; 32:260–267 PMID: <u>13083014</u>
- Misra PK, Srivastava N, Malik GK, Kapoor RK, Srivastava KL, Rastogi S. Outcome in relation to Apgar score in term neonates. Indian Pediatr. 1994; 31:1215–8. PMID: 7875781
- Ondoa-Onama C, Tumwine JK. Immediate outcome of babies with low Apgar score in Mulago Hospital, Uganda. East Afr Med J. 2003; 80:22–9. PMID: <u>12755238</u>
- 7. Padayachee N, Ballot DE. Outcomes of neonates with perinatal asphyxia at a tertiary academic hospital in Johannesburg, South Africa. South African Journal of Child Health. 2013; 7: 89–94
- The Apgar score. ACOG Committee Opinion No. 333. American Academy of Pediatrics; American College of Obstetricians and Gynecologists. Obstet Gynecol 2006; 107:1209–12. PMID: <u>16648434</u>
- 9. Lopriore E, van Burk GF, Walther FJ, de Beaufort AJ. Correct use of the Apgar score for resuscitated and intubated newborn babies: questionnaire study. BMJ. 2004; 329:143–4. PMID: <u>15208208</u>
- Rüdiger M, Wauer RR, Schmidt K, Küster H. The Apgar score. [Letter to the Editor] Pediatrics. 2006; 118:1314–1315. PMID: <u>16951034</u>
- Rüdiger M, Braun N, Gurth H, Bergert R, Dinger J. Preterm resuscitation I: clinical approaches to improve management in delivery room. Early Hum Dev. 2011; 87:749–53. doi: <u>10.1016/j.earlhumdev.</u> 2011.08.019 PMID: <u>21920678</u>
- 12. Rüdiger M, Aguar M. Newborn Assessment in the Delivery Room. Neoreviews 2012; 13;e336 doi: <u>10.</u> <u>1097/PCC.0b013e318253c945</u> PMID: <u>22791094</u>
- Rüdiger M, Kuster H, Herting E, Berger A, Müller C, Urlesberger B, et al. Variations of Apgar score of very low birth weight infants in different neonatal intensive care units. Acta Paediatr. 2009; 98:1433–6. doi: 10.1111/j.1651-2227.2009.01347.x PMID: 19558600
- Fahey J, King TL. Intrauterine asphyxia: clinical implications for providers of intrapartum care. J Midwifery Womens Health. 2005; 50:498–506. PMID: <u>16260364</u>
- Martin-Ancel A, Garcia-Alix A, Gaya F, Cabanas F, Burgueros M, Quero J. Multiple organ involvement in perinatal asphyxia. The Journal of pediatrics. 1995; 127:786–93. PMID: <u>7472837</u>
- Sarnat HB, Sarnat MS. Neonatal encephalopathy following fetal distress. A clinical and electroencephalographic study. Arch Neurol. 1976; 33:696–705. PMID: <u>987769</u>
- Sykes GS, Molloy PM, Johnson P, Gu W, Ashworth F, Stirrat GM, et al. Do Apgar scores indicate asphyxia? Lancet. 1982; 1:494–6. PMID: <u>6121150</u>
- 18. Is the Apgar score outmoded? Lancet. 1989; 1(8638):591–92. PMID: 2564115
- 19. Marlow N. Do we need an Apgar score? Arch Dis Child. 1992; 67:765–7. PMID: 1519971
- Behnke M, Eyler FD, Carter RL, Hardt NS, Cruz AC, Resnick MB. Predictive value of Apgar scores for developmental outcome in premature infants. Am J Perinatol. 1989; 6:18–21. PMID: <u>2462884</u>
- Socol ML, Garcia PM, Riter S. Depressed Apgar scores, acid-base status, and neurologic outcome. Am J Obstet Gynecol. 1994; 170:991–8; discussion 8–9. PMID: <u>8166220</u>