# Study Of The Physiology Of Nasal CPAP In An Animal Model And Clinical Impact Analysis In Newborns With Respiratory Distress

A Orozco- Gutiérrez, A D Martínez-Oñate, A T Castillo, M Pineda-Leyte, A S Martínez, E N Abinader, C R Alcocer-Arreguin, R C Gil

# Citation

A Orozco-Gutiérrez, A D Martínez-Oñate, A T Castillo, M Pineda-Leyte, A S Martínez, E N Abinader, C R Alcocer-Arreguin, R C Gil. *Study Of The Physiology Of Nasal CPAP In An Animal Model And Clinical Impact Analysis In Newborns With Respiratory Distress*. The Internet Journal of Pediatrics and Neonatology. 2013 Volume 16 Number 2.

#### **Abstract**

Background: The continuous positive airway pressure (CPAP) is used to treat respiratory failure in infants with excellent results, pneumothorax occurs from 1.4% to 10.3% of children and mortality varies between 6.5% and 48%. We reported a study in pigs to determine if the cough and nasal seal affect these results. We reported a study in pigs to determine if the cough and nasal seal affect these results.

Method: 4 pigs were anesthetized and their intrabronchial pressure was measured by applying CPAP with sealed and unsealed prongs. Then cough was caused and the intrabronchial pressure changes were measured. Results: With CPAP unsealed disappears negative inspiratory pressure and the average pressure in the airway (MAP) increases. By using sealed CPAP the findings would increase. Then cough was caused and the intrabronchial pressure changes were measured.

Results: With CPAP unsealed disappears negative inspiratory pressure and the average pressure in the airway (MAP) increases. By using sealed CPAP the findings would increase. By using CPAP sealed the findings would increase.

With the cough, the CPAP without seal decreases all pressures, with seal the PE increases and the PI is more negative. In either case the measurement of the external pressure CPAP detected intrabronchial changes. In either case the measurement of the external pressure CPAP detected intrabronchial changes.

Conclusions: The nasal CPAP with leakage increases the PI and eliminates the negative expiratory pressure, decreasing the alveolar collapse, cough increases the negative pressure and probably cause barotrauma

The nasal seal increases the PI and PMVA, increasing the oxygen diffusion gradient and improving oxygenation

With cough the pressures increase the risk of barotrauma. Monitoring in children and does not detect the intrabronchial changes of pressure in the airway.

# **INTRODUCTION**

The continuous positive airway pressure (CPAP) has been used since 1971, when Gregory and his colleagues showed that CPAP endotracheal with a bag of anesthesia improved oxygenation and reduced mortality in preterm infants with respiratory distress syndrome subsequently different devices and strategies were developed to provide it [1-5].

In the following years, the mechanical ventilation replaced CPAP as the most common form of ventilatory support until

it practically fell into disuse, was not until 1989 that Dr. Avery published a classic study by mentioning that the use of CPAP generated a lower risk of injury Chronic lung ventilation compared to mechanical ventilation, this ventilation technique was reintroduced in the treatment of these children. [6-10].

In recent years, acceptance of CPAP has increased and various articles evaluating the efficacy and safety of the procedure have been reported [11-23].

Despite the widespread use of nasal CPAP there are few studies on the physiological bases [24,25] and little is known about the variables that affect the response of patients [26]. This lack of physiological knowledge has led to very different clinical outcomes that have tried to standardize developing a large number of devices for daily treatment of neonates [27-30].

Despite the variety of available interfaces for applying CPAP [31,32] nasal prongs are the most common method and the general consensus is that they should be as broad as possible to reduce resistance and fit comfortably without air leaks or damage tissue [33].

However, there are some gray areas in their application [34-37]. It is generally accepted that to deliver intra points of pressure CPAP of bubble is represented by the depth of immersion of the expiratory tube [38-40]. However, Chilton and Brooks and De Paoli reported that pharyngeal pressure is reduced by approximately 50% in infants with open mouth [41,42].

Khan and Cols found that airway pressure during CPAP is variable and depends on the interaction of the depth of immersion and flow [43]. Another concern regarding the use of nasal CPAP is the presentation of pneumothorax as an often complication without a clear cause and tested from this accident [44].

We have not found any studies that measured the pressure changes associated with crying or hiccups, these events are common in children with CPAP and produce elevations on intrathoracic pressure in combination with treatment may be a perfect combination to cause pneumothorax.

Pneumothorax frequency associated with the use of CPAP in children with respiratory distress often varies from a low of 1.6% to 10.3% and the reported mortality is as low as 6.5% up to 48% in children with severe respiratory distress without any documented explanation for this variability [6,7,10,16,20,22,35,45,46].

The COIN study [45] and the SUPPORT study [46] two randomized controlled clinical trials have shown that CPAP can be used in young infants with high success rates, but have presented discrepancies in the risk of pneumothorax, COIN trial showed a high incidence of pneumothorax (9%) while the SUPPORT study reported a frequency of 6.8%. This was attributed to the fact that the COIN essay utilizes a CPAP of 8 cms. H2O opposed to a CPAP of 6 cms. H2O

used in the SUPPORT essay.

In 2010 we initiated a line of research trying to better understand the physiology of CPAP and achieve increased safety of the procedure, first we developed a mechanical simulator to study the physiology of CPAP and its response to common phenomena such as coughing and crying this study showed that nasal seal and common physiological phenomena have an impact on intrabronchial pressure, which affects the safety and efficacy of the procedure [48]. The study in animals intends to corroborate the findings of work in the simulator and provide a basis for further clinical studies.

# **HYPOTHESIS**

The hypothesis is that the hermetic seal of the nasal passages with nasal prongs modifies airway pressure by increasing the diffusion gradient and may experience increases or decreases during physiological phenomena such as crying or opening the mouth.

# **MATERIALS AND METHODS**

Surgical preparation.

We used 4 Yorkshire pigs with 6 days of birth weight of 2,500 grs. were anesthetized with acepromazine (0.75 mg/kg intramuscular) and sodium pentobarbital (6 mg/kg).

On sedated pig was applied 2% lidocaine and a tracheotomy was performed by placing a catheter in the secondary bronchus and measuring the intrabronchial pressure using an aneroid manometer. Performing 20 measurements for each variable in each pig.

We measured basal pressure of the animal spontaneously breathing, nasal CPAP tipped leaking and sealed and then the measurements were repeated causing coughing.

Nasal CPAP was applied with pressure of 7 cms. H20 using prongs sized to the nares of the pig and measured pressure in the system with a pressure gauge at the entrance of the prongs with the same system used in CPAP treatment in Children.

In all cases we measured ventilatory pressure for 20 cycles (inspiration and expiration), we obtained the average inspiratory and expiratory pressure in every situation, the average pressure was obtained by subtracting the average of the inspiratory pressure to the average of the expiratory pressure.

# **RESULTS**

Basal Determination without CPAP.

The average inspiratory and expiratory pressure in the bronchus without CPAP was (10/-10.29 cms. H20) and remained stable. The average pressure was 0.29.

Coughing increased the expiratory pressure (12.93 cms. H20) and inspiratory (-6 cms. H20), the average pressure was raised to 6.93.

Measuring the pressure in the system CPAP was stable varying from 2.5 cms. H20 inspiration to 7.5 cms. H20 expiration.

Nasal CPAP with unsealed ends

The average inspiratory and expiratory pressure in the bronchus with air leaking on nasal prongs in the nostrils was (9.70/0 cm. H20) range (0/12 and stable), with cough expiratory pressure was maintained but increased negative inspiratory pressure (9.85/-15 cms. H20), the average pressure decreased to -6.15.

We should mention that the range of negative inspiratory pressure was -10 to -20 cms. H2O.

Nasal CPAP with sealed edges

The average inspiratory and expiratory pressure with prongs without air leak through nostril was the 12.29/-2.5 cms. H2O, the average pressure rose to 9.79 cms. H2O.

The range of negative pressure during inspiration was from 0 to -5 cms. H2O.

Cough with inspiratory and expiratory pressure was 16/-19 cms. H20 and the average pressure was -2.4 cms. H2O.

The range of negative pressure during inspiration was from -19 to -20 cms. of water.

The pressure in the measurement system of CPAP was stable at 2.5 cms. H20 inspiration to 7.5 cms.

**Table 1**Pressure Intrabronchial without CPAP, CPAP with and

without nasal seal and changes provoked by cough.

Pressure in H2O cms.	Basal without CPAP	Nasal CPAP unsealed prongs	Nasal CPAP sealed prongs
Without Cough.			
Expiratory Pressure	10.29 (10/12)	9.70 (8 to 12)	12.29 (12 to 14)
Inspiratory Pressure.	-10 (-8 /-12)	0 (0)	2.5 (0-5)
Average pressure (pE-pl) Variation in cms. H2O	0.29	9.7	9.79
Expiratory		-0.59	2
Inspiratory		10	7.5
Average		9.41%	9.5
External pressure measured			
Expiratory	7.5	7.5	7.5
Inspiratory	2.5	2.5	2.5
With Cough			
Expiratory Pressure	12.93 (10-20)	8.85 (8-10)	16.6 (15-18)
Inspiratory Pressure	-6 (-8 to -12)	-15 (-10 to -20)	-19 (-18 to -20)
Average pressure (pE-pI) Variation in cms. H2O	6.93	-6.45	-2.4
Expiratory		-4.08	3.67
Inspiratory		-9	13.
Average		-13.8	-9.33
External pressure measured			
Expiratory	7.5	7.5	7.5
Inspiratory	2.5	2.5	2.5

# **ANALYSIS**

The average pressure on the airway is increased 9 times when using nasal CPAP with both sealed edges and leaking edges. This increases the alveolar-arterial oxygen gradient that explains the improvement of children with respiratory distress.

Expiratory pressure remains almost unchanged and the inspiratory pressure loses its negativity becoming neutral (0) in the case of leaking edges and positive (2.5) when using sealed edges, so with CPAP it lowers the alveolar collapse reducing the need for exogenous surfactant and bronchopulmonary dysplasia.

Under normal conditions (without CPAP) while coughing the average pressure increases and becomes positive increasing from 0.29 to 6.93, the expiratory pressure increases 2.73 and the inspiratory pressure loses 4 cms. H2O to rise from -10 to -6 cms. H2O. However it remains within relatively narrow ranges.

When using CPAP with or without nasal seal the average pressure becomes negative (-6.45 and -2.4 with seal) probably due to inspiratory flow limitation.

Using CPAP without seal the expiratory pressure decreases in relation to the basal pressure from 12.93 to 8.85 (-33%) by the same flow limitation and by placing the nasal seal increases from 12.93 to 16.6 (28.3%) as the system becomes

closed and further allows increments in pressure with the same volume.

The inspiratory pressure with CPAP and cough becomes more negative in the case of CPAP with leak varies from -6 to -9 (50%) with nasal CPAP sealed the negative pressure changes from -6 to -13 (116%) this negative pressure with respect to this calculated flow for a basal pulmonary volume, at a higher volume sued by coughing and found this limited the negative pressure is increased significantly and can be the cause of pneumothorax frequently encountered in clinical practice.

# **DISCUSSION**

Our work verifies the results obtained by Khan [45] Chilton [39] and De Paoli [40], who demonstrated that the pressure in the airway when employed CPAP is not constant, is influenced by several factors and is not reflected in measuring the pressure in the CPAP circuit. We complement these studies, and we showed that there are secondary pressure changes to common physiological phenomena as cough, events that are common in infants receiving CPAP.

Traditionally, studies have used pressure gauges as close as possible to the nasal cannula as a measure of intrapulmonary pressure [46,47], in agreement with our results from the pressure near the nasal edges is different from the intrapulmonary pressure.

In our study, the use of CPAP with sealed edges increases the pressure and can improve oxygen gradient making the technique more efficient, but when coughing significantly increases the negative pressure, probably by decreasing inspiratory flow and this factor may be causing barotrauma making it more risky.

These intrabronchial pressure changes are not reflected on the gauge system of CPAP, which casts doubt on the effectiveness of surveillance systems of the pressure due to their limited value to prevent acute lung injury by CPAP.

Unexpected increases related to intrabronchial pressure with sealed nostrils and normal physiological events may be an important factor for the wide variability in the results of studies on the efficacy and safety of nasal CPAP and have to be considered in future work on this ventilatory technique.

This study explains the results published in 1973 by Gregory and corroborated in 2009 [11] using tracheal CPAP. In this technique the patient is intubated and connected to a CPAP

with an elastic anesthesia bag, which has a high volume that prevents inspiratory pressure to drop and being elastic absorbs pressure surges to maintain alveolar stability and a gradient alveolar-arterial stable.

#### CONCLUSIONS

The use of nasal CPAP has been widely disseminated and has proven an effective procedure for the treatment of premature infants with respiratory distress with an efficacy similar to mechanical ventilation. This is a very well accepted and safe in the treatment of lung diseases.

This study supports our previous findings on a mechanical simulator [48]

The hermetic nasal seal in nostrils either tight ends, secretions or mucosal edema produces unexpected changes in intra bronchial pressure, affecting the efficacy and safety of nasal CPAP. Reducing air leaks the CPAP with nasal seal is more effective. Very young children may need more respiratory support with stable and higher pressures, however the use of nasal CPAP with seal can increase the risk of barotrauma.

In older children or those with less severe lung disease the use of CPAP with leak on nasal edges may be sufficient to maintain respiratory homeostasis and alveolar stability.

Inadvertent changes secondary to nasal seal may explain variability in efficacy and safety results observed in previous studies, and can be influenced by local events routines and uncontrolled (techniques for fixing the points of the same size, edema of the nasal mucosa, nasal bleeding with clots and scabs and cleaning techniques).

The hermetic seal on the nostril during the use of nasal CPAP is a factor, which influences the pressure in the airways and should be taken into account during clinical trials to ensure more consistent and reliable results.

Maintaining stable figures in the surveillance system for pressure in the CPAP does not exclude the presence of sudden intrabronchial variations herefore not useful to reduce the risk of barotrauma.

### References

- 1.- Gregory GA, Kitterman JA, Phibbs RH, Tooley WH, Hamilton WK.Treatment of the idiopathic respiratory distress syndrome with continuous positive airway pressure. N Engl J Med 1971, 284:1333-1340.
- 2.- Kattwinkel J, Fleming D, Cha CC, Fanaroff AA, Klaus

- MH. A device for administration of continuous positive airway pressure by the nasal route. Pediatrics 1973, 52:131-4.
- 3.- Dunn. PM. -Respiratory distress syndrome. Continuous positive airway pressure (CPAP) using the Gregory box. Proc R Soc Med. 1974 April, 67 (4): 245-247.
- 4.- Krouskop RW, Brown EG, Sweet AY. -The early use of continuous positive airway pressure in the treatment of idiopathic respiratory distress syndrome. J Pediatr 1975, 87:263-7.
- 5.- Wung JT, Driscoll JM Jr, Epstein RA, Hyman AI. A new device for CPAP by nasal route. Crit Care Med 1975, 3:76
- 6.- Avery ME, Tooley WH, Keller JB, Hurd SS, Bryan MH, Cotton RB et al, Is Chronic Lung Disease in Low Birth Weight Infants Preventable? A Survey of Eight Centers. Pediatrics, Jan 1987, 79: 26-30.
- 7.- Latini G.-Minimal handling and bronchopulmonary dysplasia in extremely low-birth-weight infants. Eur J Pediatr. 2003 Apr, 162 (4):227-9.
- 8.- Liptsen E, Aghai ZA, Pyon KE, Saslow JG, Nakhla T, Long J. -Work of Breathing During Nasal Continuous Positive Airway Pressure in Preterm Infants: A Comparison of Bubble vs. Variable-Flow Devices. Journal of Perinatology 2005, 25:453-458
- 9.- Higgins RD, Richter SE, Davis JM. Nasal continuous positive airway pressure facilitates extubation of very low birth weight neonates. Pediatrics 1991, 88 (5):999-1003.
- 10.- De Klerk AM, De Klerk RK. Nasal continuous positive airway pressure and outcomes of preterm infants. J Paediatr Child Health 2001, 37:161 7.
- 11.- Orozco A, Estrada RM: Orotracheal Continuous Positive Airway Pressure Compensation With a Bag: A Preliminary Report About a New Technique for Assisted Ventilation in Premature Infants. The Internet Journal of Pediatrics and Neonatology. 2009 Volume 10 Number 2
- 12.- Kugelman A. Feferkorn I. Riskin A. I Chistyakov. Kaufman B. Bader D. -Nasal Intermittent Mandatory Ventilation Versus Nasal Continuous Positive Airway Pressure for Respiratory Distress Syndrome: A Randomized, Controlled, Prospective Study. Pediatr Respir Rev 2003, 4:2-8.
- 13.- Nowadzky T, Pantoja A. Britton JR. -Bubble Continuous Positive Airway Pressure, A Potentially Better Practice Reduces the Use of Mechanical Ventilation Among Very Low Birth Weight Infants With Respiratory Distress Syndrome. Pediatrics, 2009, 123: 1534-1540
- 14.- Narendran V, Donovan EF, Hoath SB, Akinbi HT, Steichen JJ, Jobe AH. Early Bubble CPAP and Outcomes in ELBW Preterm Infants. Journal of Perinatology 2003, 23:195-199
- 15.- Dani C, Bertini G, Pezzati M, Cecchi A, Caviglioli C, Rubaltelli FF. -Early Extubation and Nasal Continuous Positive Airway Pressure After Surfactant Treatment for

- Respiratory Distress Syndrome Among Preterm Infants <30 Weeks' Gestation. Pediatrics 2004, 113: e560-e563
- 16.- Polin A. R -Bubble CPAP: A Clash of Science, Culture, and Religion. The Journal of Pediatrics. -2009,54:5:633-634.
- 17.- Courtney E, Barrington S, Keith J. -Continuous Positive Airway Pressure and Noninvasive Ventilation. Clin Perinatol 34 (2007) 73-92
- 18.- Codazzi D, Nacoti M, Passoni M, Bonanomi E, Sperti LR, Fumagalli R. Continuous positive airway pressure with modified helmet for treatment of hypoxemic acute respiratory failure in infants and a preschool population: A feasibility study. Pediatr Crit Care Med 2006, 7(5):455-460. doi:10.1097/01.PCC.0000235246.68050.3A
- 19.- Morley CJ, Lau R, De Paoli A, Davis PG. Nasal continuous positive airway pressure: Improve bubbling does gas exchange?. Arch Dis Child Fetal Neonatal Ed 2005 (90) F343-F344.
- 20. AARC Guideline: Neonatal CPAP. Application of Continuous Positive Airway Pressure to Neonates via Nasal Prongs, Nasopharyngeal Tube, or Nasal Mask -2004 Revision & Update. Respiratory Care.- 2004 (49) 9: 1101-1108
- 21.- Morley C, Davis P. Continuous positive airway pressure: current controversies. Curr Opin Pediatr. April 2004 (16) 2:141-5.
- 22.- Diblasi RM. Nasal continuous positive airway pressure (CPAP) for the respiratory care of the newborn infant. Respir Care. 2009 Sep, 54(9):1209-35.
- 23.- Dunn MS, Reilly MC. Approaches to the initial respiratory management of preterm neonates. Paediatr Respir Rev. 2003 Mar, 4(1):2-8.
- 24.- Wild M, Alagesan K. PEEP and CPAP. British Journal of Anaesthesia. CEPD Reviews (2001) 1 (3): 89-92. doi: 10.1093 / bjacepd/1.3.89
- 25.- Pillow JJ, Hillman N, Moss TJ, et al. Bubble continuous positive airway pressure enhances lung volume and gas exchange in preterm lambs. Am J Respir Crit Care Med 2007, 176:63-9.
- 26.- De Paoli AG, Morley C, Davis PG. Nasal CPAP for neonates: what do we know in 2003? Arch Dis Child Fetal Neonatal Ed 2003, 88(3):F168-F172 doi: 10:1136/fn.88.3.F168
- 27.- Pandit PB, Courtney SE, Pyon KH, et al. Work of breathing constantant During variable-flow nasal continuous positive airway pressure in preterm neonates. Pediatrics 2001, 108:3:682-685
- 28.- Boumecid H, Rakza T, Abazine A, Klosowski S, Matran R, and Storme L. Influence of three nasal continuous positive airway pressure devices on breathing pattern in preterm infants. Arch Dis Child Fetal Neonatal Ed 2007, 92:F298-F300 Published Online First: 6 November 2006 doi:10.1136/adc.2006.103762
- 29.- Gutiérrez Laso A, Sáenz González P, Izquierdo Macián

- I, Fernández Gilino C, Gimeno Navarro A, Moreno M Gormaz, Torres Palomares D, Morcillo Sopena F, Roqués Serradilla V. -Nasal continuous positive airway pressure in preterm infants: comparison of two low-resistance models. An Pediatr (Barc). 2003 Apr, 58(4):350-6.
- 30.- KM Chan, HB Chan. The Use of Bubble CPAP in Premature Infants: Local Experience. HK J Paediatr 2007, 12:86-92
- 31.- J. Hammer. Nasal CPAP in preterm infants does it work and how? Intensive Care Medicine 2001 Nov, 27(11):1689-91. Epub 2001 October 12.
- 32.- Courtney SE, Pyon KH, Saslow JG, et al. Lung recruitment and breathing pattern varying during versus continuous flow nasal continuous positive airway pressure in premature infants: an evaluation of three devices. Pediatrics 2001, 107 (2):304-308.
- 33.- Louis P. Halamek, Colin Morley. Continuous Positive Airway Pressure During Neonatal Resuscitation. Clin Perinatol 33 (2006) 83-98
- 34.- De Paoli AG, Davis PG, Faber B, Morley CJ. Devices and pressure sources for administration of nasal continuous positive airway pressure (NCPAP) in preterm neonates. Cochrane Database of Systematic Reviews 2008, Issue 1. Art No.: CD002977. DOI: 10.1002/14651858.CD002977.pub2
- 35.- Polin RA, Sahni R. New Experience with CPAP. Semin Neonatol 2002, 7(5):379-389.
- 36.- Simpson JH, Ahmed I, McLaren J, Skeoch CH. Use of nasal continuous positive airway pressure during neonatal transfer. Arch Dis Child Fetal Neonatal Ed 2004, 89(4):F374-F375 doi: 10.1136/adc.2003.033159
- 37.- Elgellab A, Riou And Abbazine A, et al. Effects of nasal continuous positive airway pressure (NCPAP) on breathing pattern in spontaneously breathing premature newborn infants. Intensive Care Med. Volume 27, Number 11, 1782-1787.
- 38.- Tsyr-Yuh Ho, Shan-Fu Ou, Shih-Hui Huang, Chi-Ning Lee, Luo-Ping Ger, Kai-Sheng Hsieh, Hui-Ying Cheng, Wei Yang Lee, Ken-Pen Weng. Effects of Flow Rate on Delivery of Bubble Continuous Positive Airway Pressure in an In Vitro Model. Neonatol Pediatr 2010, 51(4):214-8
- 39.- J. Jane Pillow, Noah Hillman, Timothy JM Moss,

- Graeme Polglase, Geoff Bold, Chris Beaumont, Machiko Ikegami, and Alan H. Jobe. Bubble Continuous Positive Airway Pressure Enhances Lung Volume and Gas Exchange in Preterm Lambs. Am J Respir Crit Care Med 2007 July 1, 176 (1): 63-69.
- 40.- Lee KS, Dunn MS, Fenwick M, Shennan AT. A comparison of underwater bubble continuous positive airway manufacturer-ventilator-derived continuous positive airway pressure in premature neonates ready for extubation. Biol Neonate. 1998, 73(2):69-75
- 41.- Chilton HW, Brooks JG. Pharyngeal pressures in nasal CPAP. J Pediatr.1979 May, 94(5):808-10
  42.- De Paoli AG, Lau R, Davis PG, Morley CJ. Pharyngeal pressure in preterm infants Receiving nasal continuous positive airway pressure. Arch Dis Child Fetal Neonatal Ed 2005, 90:F79-F81.doi: 10.1136/adc.2004.052274
- 43.- Doron J. Kahn, Robert H. Habib and Sherry E. Courtney. Effects of Flow Amplitudes on Intraprong Pressures During Bubble Versus Ventilator-Generated Nasal Continuous Positive Airway Pressure in Premature Infants. Pediatrics 2008, 122:1009-1013 (doi: 10.1542/peds.2007-3416)
- 44.- Doron Kahn, Sherry Courtney, Andrew Steele, Robert Habib. -Unpredictability of Delivered Bubble Nasal Continuous Positive Airway Pressure: Role of Bias Flow Magnitude and Nares-Prong Air Leaks. Pediatr Res. 2007 Sep, 62(3):343-7.
- 45.- Makhoul IR, Smolkin T, Sujov P. Pneumothorax and nasal continuous positive airway pressure ventilation in premature neonates: a note of caution. ASAIO J 2002, 48(5):476-479.
- 46.- Morley CJ, Davis GP, Doyle LW, Brion LP, Ascote JM, Carlin JB. Nasal CPAP or Intubation at birth for very preterm infants. New England Journal of Medicine 2008, 358:700-708
- 47.- Support Study Group of the Eunice Kennedy Shriver NICHD Neonatal Research Network. Early CPAP versus Surfactant in Extremely Preterm Infants. N Engl J Med 2010, 362:1970-1979 May 27, 2010
- 48.- A. Orozco-Gutierrez, RM Estrada-Velazquez, C. Gil-Rosales: Analysis of distal airway pressure changes in a simulation model of continuous positive airway pressure (CPAP). The Internet Journal of Pediatrics and Neonatology. 2011 Volume 13 Number 1. DOI: 10.5580/192b

# **Author Information**

# Alberto Orozco- Gutiérrez

Research Department, Facultad Mexicana de Medicina de la Universidad La Salle (Mexican Faculty of Medicine, La Salle University), Tlalpan, Mexico; Department of Neonatology. Hospital Ángeles del Pedregal

Mexico City, Mexico

orozcogutierrezalberto@gmail.com

# Ariel de Jesús Martínez-Oñate

Department of Neonatology. Hospital Ángeles del Pedregal Mexico City, Mexico

# **Arturo Tovar Castillo**

Research Department, Facultad Mexicana de Medicina de la Universidad La Salle (Mexican Faculty of Medicine, La Salle University)

Tlalpan, Mexico

# Miriam Pineda-Leyte

Research Department, Facultad Mexicana de Medicina de la Universidad La Salle (Mexican Faculty of Medicine, La Salle University)

Tlalpan, Mexico

# Alan Salas Martínez

Research Department, Facultad Mexicana de Medicina de la Universidad La Salle (Mexican Faculty of Medicine, La Salle University)

Tlalpan, Mexico

# **Emilia Nuñez Flor Abinader**

Department of Neonatology. Hospital Ángeles del Pedregal

Mexico City, Mexico

# Christian Rodrigo Alcocer-Arreguin

Research Department, Facultad Mexicana de Medicina de la Universidad La Salle (Mexican Faculty of Medicine, La Salle University), Tlalpan, Mexico; Department of Neonatology. Hospital Ángeles del Pedregal Mexico City, Mexico

# **Rosales Cesar Gil**

Department of Neonatology. Hospital Ángeles del Pedregal Mexico City, Mexico