

## MEDICINE

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### SEVERE BRONCHOPULMONARY DYSPLASIA AND PEDIATRIC ACUTE RESPIRATORY DISTRESS SYNDROME. RESEMBLANCE AND CONTRAST

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#### Abstract

The leading causes of death among children under five still stay a respiratory insufficiency, preterm birth complications, pneumonia, intrapartum-related complications, diarrhea, and congenital abnormalities. Acute respiratory distress syndrome (ARDS) is the most dramatic complication of lung system failure. Similar clinical manifestation has a severe bronchopulmonary dysplasia (BPD). Both of these pathologies lead to severe respiratory insufficiency and require intensive therapy, respiratory support in paediatric intensive care unit (PICU). Manifestation of severe BPD similar to ARDS, but it is not the same and management has some nuances. In this article we had tried to analyze resemblance between BPD and PARDS and determine contrasts.

**Keywords:** *BPD. ARDS. Chronic lung disease. Intensive therapy. Respiratory support.*

**Introduction:** Bronchopulmonary dysplasia (BPD) was known since 1967 year as a chronic lung disease most commonly seen in premature infants who required mechanical ventilation and oxygen therapy for acute respiratory distress but can also occur in neonates that had a less severe respiratory course [1,2]. BPD is the most common chronic respiratory disease

in infants. BPD is the leading cause of chronic lung disease in children [3]. Main causes of BPD these are lung tissue immaturity, mechanical ventilation, oxygen toxicity, infection and inflammation, genetic predisposition. Mortality at children with severe BPD until 2 years old about 25%[4].

**Materials and methods:** We had provided literature review of up to dated information about BPD and PARDS, further more comparing analysis of collected data was done. Our own observing study was provided in the City hospital #1 Astana, Kazakhstan. We had provided observation of clinical manifestations and laboratory data collection at 18 children. The groups were balanced by age they were studied during hospitalisation at PICU. Age of the patients from 28 days till 3 years old, 7 of them with PRDS and 11 with severe BPD in paediatric intensive care unit. All patients had a respiratory insufficiency of the 3rd degree, due to pneumonia. They were on invasive ventilation, orotracheal intubation, with the same modes of ventilation. Management of all patients was equal includes antibacterial therapy, fluid management, cardiovascular support, diuretics and artificial lung ventilation.

ARDS represents the most severe form of acute lung injury (ALI) and is characterized by alveolar leukocyte infiltration and protein-rich pulmonary oedema [5]. Mortality from paediatric acute respiratory distress syndrome (PARDS) varies widely from 15% and up to 45% [6,7].

**Results:** The same Biomarkers in BPD and PARDS were detected in numerous investigations. Various biomarkers detected in different biological fluids have been proposed for early identification of BPD predisposed newborns (8). Among these that have been implicated both in BPD and PARDS (*vide infra*) are the following. KL-6 (a lung injury marker), interleukin (IL)-6, interleukin-8, sICAM-1, angiopoietin-2, and matrix metalloproteinase-8,9. [9-13].

Pathogenesis of BPD and PARDS. BPD is caused due to an interaction between genetic and environmental factors (hyperoxia, invasive mechanical ventilation, and sepsis [14]. Immature lung tissue impacted by external factors: infections, high concentration of oxygen, long time ventilation, barotrauma, volutrauma, or atelectotrauma, which initiates a cascade of inflammation reaction involving cytokines. This activates the cell death pathways. Damage of immature lungs is followed by resolution of injury to close to normal lung architecture or repair, and leads to fibrosis [15]. On the other hand PARDS pathogenesis consists of cascade mechanism after direct pulmonary tissue damage resulting into the destruction of alveolar-capillary unit. Damage of alveolar-capillary barrier resulted in increased permeability of big molecules such as protein rich oedema fluid into the alveolar lumen, dysfunction of surfactant production, and impaired fluid clearance from alveolar cells. These changes are ARDS pathophysiology chain and accompanied by dysregulated inflammation from dysfunctional leukocytes and influx of cytokines [13].

Pathogenesis of BPD and PARDS has a lot of resembling features and contrast points. But one pathology cannot exclude another one. Moreover, children with BPD are at risk for development ARDS. [16].

Data of patients with PARDS and BPD. Tab.1.

	PARDS	BPD
Total amount	7	11
Mean day of PICU hospitalisation	27	31
Type of respiratory support	Invasive ventilation	Invasive ventilation
Regiment of ventilation	SIMV	SIMV
Mean days on ventilation.	12.3(±4.2)	21.4(± 6.1)
Mean days on noninvasive respiratory support.	6.4(±3.8)	7.8(±7.2)
Dead	1(14.3%)	3 (27.3%)

Findings indicate that children with bronchopulmonary dysplasia have longer duration of hospitalisation in PICU than children with respiratory distress syndrome. Quantity of days on respiratory support in BPD group was higher. Furthermore, in the group of patients with severe BPD mortality rate was higher than in group with ARDS. At the same moment management was consimilar. During observing study of patients with severe respiratory insufficiency we emphasised what patients with BPD had high titre of cytomegalovirus infection, but this issue was not included in our study, potentially it could have impact to results. This item should be examined more carefully.

**Conclusion:** despite of numerous resembling points in aetiology, pathogenesis and biomarkers diagnostics PARDS and BPD is not the same, because of outcomes are divers. Group of patients with respiratory insufficiency on the background of BPD had worse results than previously well fitted patients with ARDS. Variability in reported data presents challenges in tactics of intensive care of patients with BPD. Patients with severe BPD should be managed in another way. And issues at management of severe respiratory failure in the background of BPD in PICU should be more enlightened.

**References:**

- [1] Northway W.H., Jr., Rosan R.C., Porter D.Y. // Pulmonary disease following respiratory therapy of hyaline-membrane disease. Bronchopulmonary dysplasia. *New Engl. J. Med.* 1967. 276:357–368. doi: 10.1056/NEJM196702162760701. [PubMed] [Cross Ref]

- [2] Jobe A.H., Bancalari E. // Bronchopulmonary dysplasia. *Am. J. Respir. Crit. Care Med.* 2001;163:1723–1729. doi: 10.1164/ajrccm.163.7.2011060. [PubMed] [Cross Ref].
- [3] Bhandari A, Bhandari V. // Biomarkers in bronchopulmonary dysplasia. *Paediatr Respir Rev* (2013) 14(3):173–9. 10.1016/j.prrv.2013.02.008 [PubMed] [Cross Ref].
- [4] S. Walker et al // Mortality Estimates for Severe Bronchopulmonary Dysplasia Complicated by Pulmonary Hypertension. Conference: 2011 PH Professional Network Symposium.
- [5] Erickson S, Schibler A, Numa A. et al. // Acute lung injury in pediatric intensive care in Australia and New Zealand: a prospective, multicenter, observational study. *Pediatr Crit Care Med* (2007) 8(4):317–23. 10.1097/01.PCC.0000269408.64179.FF [PubMed] [Cross Ref].
- [6] Kneyber MC, Brouwers AG, Caris JA, Chedamni S, Plotz FB. // Acute respiratory distress syndrome: is it underrecognized in the pediatric intensive care unit? *Intensive Care Med* (2008) 34(4):751–4. 10.1007/s00134-008-1029-4 [PubMed] [Cross Ref].
- [7] Lopez-Fernandez Y, Azagra AM, de la Oliva P, Modesto V, Sanchez JI, Parrilla J, et al. // Pediatric acute lung injury epidemiology and natural history study: incidence and outcome of the acute respiratory distress syndrome in children. *Crit Care Med* (2012) 40(12):3238–45. 10.1097/CCM.0b013e318260caa3 [PubMed] [Cross Ref].
- [8] A. Bhandari, C. Carroll, V. Bhandari // BPD Following Preterm Birth: A Model for Chronic Lung Disease and a Substrate for ARDS in Childhood. *Front Pediatr.* 2016; 4: 60. Published online 2016 Jun 15. doi: 10.3389/fped.2016.00060
- [9] Kim DH, Kim HS, Shim SY, Lee JA, Choi CW, Kim EK, et al. // Cord blood KL-6, a specific lung injury marker, correlates with the subsequent development and severity of atypical bronchopulmonary dysplasia. *Neonatology* 2008. 93(4):223–9. 10.1159/000111100 [PubMed] [Cross Ref]
- [10] Ogihara T, Hirano K, Morinobu T, Kim HS, Ogawa S, Hiroi M, et al. // Plasma KL-6 predicts the development and outcome of bronchopulmonary dysplasia. *Pediatr Res* 2006. 60(5):613–8. 10.1203/01.pdr.0000242361.47408.51 [PubMed] [Cross Ref]
- [11] Bhandari V, et al. // Hyperoxia causes angiotensin 2-mediated acute lung injury and necrotic cell death. *Nat Med* 2006. 12(11):1286–93. 10.1038/nm1494 [PMC free article] [PubMed] [Cross Ref]
- [12] Aghai ZH et al. // Angiotensin 2 concentrations in infants developing bronchopulmonary dysplasia: attenuation by dexamethasone. *J Perinatol* (2008) 28(2):149–55. 10.1038/sj.jp.7211886 [PubMed] [Cross Ref]

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- [13] Sapru A, Flori H, Quasney MW, Dahmer MK. //Pediatric Acute Lung Injury Consensus Conference Group . Pathobiology of acute respiratory distress syndrome. *Pediatr Crit Care Med* 2015. 16(5 Suppl 1):S6–22.10.1097/PCC.0000000000000431 [PubMed] [Cross Ref]
- [14] Baraldi E, Filippone M. //Chronic lung disease after premature birth. *N Engl J Med* 2007. 357(19):1946–55.10.1056/NEJMra067279 [PubMed] [Cross Ref].
- [15] Bhandari V. //Bronchopulmonary dysplasia. E –book. (2016) ISSN 978-3-319-28486-6. USA.
- [16] Lal CV, Ambalavanan N.// Pulmonary hypertension in bronchopulmonary dysplasia. In: Bhandari V, editor. , editor. *Bronchopulmonary Dysplasia. Respiratory Medicine*. Switzerland: Springer; 2016. p. 259–80.