

Risk Factors for Severe Lower Respiratory Tract Infections among Infants in Tripoli-Libya: A Case-Control Study

ELHassan Dawi¹, Monia Suweadi¹ and Marwa Mahfoud²

¹Department of Paediatrics, Tripoli Children Hospital, Faculty of Medicine, University of Tripoli, Tripoli - Libya

²Tripoli Children's Hospital, Tripoli -Libya

Received 5 April 2017 / Accepted 25 May 2017

ABSTRACT

Acute Lower respiratory tract infections (ALRTI) are major cause of mortality and morbidity among children worldwide and particularly children in developing countries

The objective of this case control- study is to evaluate the risk factors for hospitalization with severe ALRTI in Tripoli-Libya. The study was conducted from November 2012 to March 2013.

Cases were recruited from all infant aged 1 to 12 months admitted to Tripoli children's hospital with severe ALRTI and who met the inclusion criteria during that period. Controls were recruited from vaccination centers in different area inside Tripoli. Mothers were interviewed for potential risk factors using pre-designed questionnaire includes infant's age, sex, mother age, maternal education level, occupation, parity, pattern of feeding in the first four months of life, number of house hold, number of persons sleeping with the infant in the same bed room and passive smokers. 154 cases and 194 controls were enrolled in the study.

In logistic regression model, significantly increased risk of hospitalization was associated with, young infant less than 3 months of age (odds ratio (OR) = 2.2, 95% CI 1.3-3.8), formula feed infant (OR=2.3, 95% CI 1.1-4.8), large number of house hold (OR=1.3, 95% CI 1.1-1.6) and increased number of person sleeping with infant at the same room (OR=1.9, 95% CI 1.4-2.4).

On another hand highly educated mothers (OR= 0.42, 95% CI 0.2-0.8) and mother's age above 30 years (OR = 0.19, 95% CI 0.08-0.4) were protective factors.

In this study several risk factors for severe ALRTI were identified, some factors are amenable to interventions.

Key words- Risk factors; Case-control study; Acute lower respiratory tract infections; Infants

INTRODUCTION

Acute lower respiratory tract infections (ALRI), such as pneumonia and acute bronchiolitis, are the leading cause of morbidity and mortality in children under five years of age. According to recent estimates, every year about 120-156 million cases of ALRI occur globally with approximately 1.4 million resulting in death. More than 95% of these deaths occur in developing countries.¹

ALRI are major reason for admission to Tripoli children's hospital especially in winter season.

ALRI are caused by a number of infective agents, with *Streptococcus pneumoniae* being generally the most frequently identified bacterial agent, and Respiratory Syncytial Virus being the most frequent viral agent.²

A large number of factors determine whether the contact with an etiologic agent will result in a severe episode of ALRI. These factors are related to the child, disease, environment, the family and its socio-economic status, the health system and type of care.^{3,4}

The child's age has been shown to be an important determinant of severity of ALRI; the rates of hospitalization which are marker of severity, were up to 10 times higher for infants comparing with children aged 1 to 4 years.⁵

In spite of the high incidence of acute respiratory infections, there is very little information on the risk factors for these infections. Epidemiological studies of LRTI in developing countries identified low birth weight, malnutrition, vitamin A deficiency, lack of breastfeeding and passive smoking as risk factors for LRTI.⁶ Recent studies have added other risk factors to the list including poor socioeconomic status, large family size, and family history of bronchitis, advanced birth order, crowding, young age, inadequate ventilation and air pollution.^{7,8} The identification of risk factors associated with severe LRTI especially these amenable to corrective measures is vital for the formulation of more effective policies and strategies to improve health globally.⁹

This case-control study was conducted to identify the risk factors for severe ALRTI among hospitalized infant at Tripoli-Libya.



MATERIALS AND METHODS

Unmatched Case-control study conducted between November 2012 to March 2013 at Tripoli Children Hospital, Tripoli-Libya, a tertiary care hospital situated in northern west part of Libya.

All infants aged between 1 month and 12 months hospitalized with severe ALRTI and who met the inclusion criteria during the study period were included in the study as cases. The inclusion criteria include, infant's age between 1 to 12 months, full term, with normal birth weight and free from other chronic illness like congenital heart disease, chronic diarrhea, etc.

Hospitalization due to severe ALRTI was defined as "in-hospital care for ≥ 24 hours, when the infant presented with chest in drawing and/or tachypnea in the presence of cough or difficult breathing, complemented by information on other clinical signs (crepitation, wheezing) and/or radiological evidence of lung disease".^{10,11}

Healthy infants between 1 month and 12 months, attending primary care center for routine vaccination in different areas inside Tripoli city were included in the study as controls.

The data was obtained from the mothers in hospital for cases and in vaccination clinics for controls, using pre-designed questionnaire.

The questionnaire encloses the potential risk factors including the infant age, sex, feeding pattern in the first 4 months of life, maternal age, parity, education level, occupation, family size and number of persons sleeping with the infant at the same room and passive smoking.

Statistical analysis was conducted using Stata® version 12, Odds ratio estimated with 95% CI using logistic regression to adjust for confounders.

RESULTS

There were 154 infant admitted with severe ALRTI and met the inclusion criteria to Tripoli Children's Hospital during the study period. The cases were matched to 194 controller.

The cases and controls did not differ significantly in age, the mean infant age was 4 ± 2.6 months for cases and 3.9 ± 2 months for controls, and there were more boys in cases (59.5%) as compared to controls (49%).

Infant's age of less than three months was associated with increased risk of severe ALRTI, OR=2.2, 95% CI 1.3-3.8 (Table 1).

The mean maternal age was almost the same in both groups 30 ± 5 years for cases and 31 ± 5 years for controls.

The risk of developing severe ALRTI was lower among infants of mothers with higher level of education (OR=0.42, 95% CI 0.21-0.82) when compared with lower level of maternal education. Increased maternal age (above 30 years) was associated with lower risk of severe ALRTI, OR = 0.19, 95% CI 0.08-0.44, (Table 2).

Increased number of house hold or the number of persons sleeping with infant at the same room were associated with increased risk of severe ALRTI (Table 3).

Concerning the four types of feeding during the first four months of life, this study found that, formula only group had a higher risk of severe ALRTI compared with other groups, OR=2.3, 95% CI 1.1- 4.8) (Table

Table 1: Distribution of cases and controls according to infant age and sex with respective odds ratios (OR), confidence intervals (95% CI) and P value.

Characteristics	Cases (%) no	Control (%) No	Adjusted OR (CI 95%)	P value
<i>Infant age</i>				
months 3 ≤	(51.3%)79	(58.8)114	1	
months 3 >	(48.7%)75	(41.2)80	(1.3-3.8)2.2	0.004
<i>Infant sex</i>				
Male	(59%)91	(49)95	1	
Female	(41%)63	(51)99	(0.53-1.50)0.88	0.63



Table 2: Distribution of cases and controls according to maternal characteristics with respective OR, 95% CI and *P* value.

Characteristics	(%) Case No	(%) Control no	Adjusted OR (CI 95%)	<i>P</i> value
<i>Maternal education level</i>				
Less than high school	(20.8)32	(11.9)23	1	
High school or university	(79.2)122	(88.1)171	(0.21-0.82)0.42	0.016
<i>Maternal age</i>				
Less than 25 y	(18.2)28	(9.8)19	1	
y 25-30	(28.6)44	(27.8)54	(0.17-0.90)0.39	0.027
More than 30 y	(53.2)82	(62.4)121	(0.08-0.44)0.19	0.001>
<i>Maternal work</i>				
House wife	(68.2)105	(61.9)120	1	
Employ	(31.8)49	(38.1)74	(0.48-1.18)0.76	0.22
<i>(Parity)</i>	O.R per child 1.1(0.9-1.30)			0.058

Table 3: Distribution of cases and controls according to house condition and smoking with respective OR, 95% CI and *P* value.

Characteristics	Case (%)No	Control (%)No	Adjusted OR (CI 95%)	<i>P</i> value
<i>House crowding</i>				
House hold size			(OR Per person 1.3(1.1-1.6)	0.002
No sleeping with child in same room			(OR Per person 1.9(1.4-2.4)	0.001>
<i>Smoking</i>				
Smoking	(49.3)76	(55.1)107	1	
No smoking	(50.7)78	(44.9)87	(0.82-2.40)1.40	0.20

Table 4: Distribution of cases and controls according to the type of feeding in first 4 months with respective OR, 95% CI and *P* value.

Type of feeding	Cases (%)No	Control (%)No	(Adjusted OR 95% CI	<i>P</i> value
Exclusive breast	(23)36	(23)45	1	
Breast with formula≤3 times day	(12)19	(32)63	(0.19-0.74)0.38	0.01
Breast with formula>3 times day	(34)52	(28)54	(0.67-2.2)1.2	0.87
Formula	(31)47	(16)32	(1.1-4.8)2.3	0.033



4).

DISCUSSION

This study had provided an optimal framework to assess the risk factors of severe ALRTI among infant 1 to 12 months of age in Tripoli, in particular the modifiable risk factors. We consider this study is the first study to examine risk factors for severe ALRTI in this area.

Young infants, young mothers, lower maternal education, bottle feeding, and increased number of house hold and the number of children sleeping at the same bed room have shown significant association with severe ALRTI.

More boys were effected compared with girls in this study, although this was not statistically significant. However, a significant association has been observed in some other studies.^{12,13}

Infant's age was a significant risk factor for severe lower respiratory tract infections. The highest risk was found among children aged 1 to 3 months. It is well known that the age between 6 and 18 months has indeed been termed as the period of vulnerability due to degradation of maternal antibodies and immaturity of immune system.¹⁴ The finding of the opposite relative risk of severe lower respiratory tract infections for children aged 1 to 3 months in our study may be explained by yet the risk of transmission of infectious agents may be smaller in the youngest age group, but when infected this group experiences severe symptoms.

Concerning maternal factors, higher level of maternal education (high school and university level) appeared to reduce the risk of severe ALRTI compared with a lower level of education (primary and secondary level). The relationship between better maternal education and lower risk of infection is widely known in the literature.¹⁵⁻¹⁷ Mothers with better education are more likely to have better socioeconomic positions enabling for better nutrition, better housing environment, better access to health and social care.

In our study, the association between higher education and lower risk of severe ALRTI was significant.

Regarding maternal age, older maternal age was protective factor. The protective effect of older maternal age on children's respiratory health has been noted previously.^{18,19} and was further proved in this study. Most authors explain this association based on common sense arguments i.e. the older the mother, the more experienced she should be in caring for her child.

Despite that literatures shows that maternal work and parity have significant negative effect on child health²⁰⁻²¹ our study does not demonstrate any statistically significant risk of these two factors.

In relation to the risk of severe ALRTI and type of feeding in the first four months of life. We found a protective effect of exclusive breastfeeding or more breast feeding (3 bottle or less per day) compared with bottle feeding or more bottle feeding (more than 3 bottles per day).

A similar protective effect of exclusive breast feeding has been observed in previous studies from other parts of the world.²²

Moreover we noticed that, the most protective factor is more breast feeding (3 bottles or less per day group), the reason of why this factor is the most protective even when compared with exclusive breast needs further research.

Household crowding and Night time crowding were risk factors for severe ALRTI due to enhancement of exposure to infectious agents especially when sharing the bedroom.^{23,24} Our study showed a significant increase in the risk of severe ALRTI due to crowding.

It is well known that houses in Libya are large and that housing standards have improved, but many families have the habit of sleep in one bed room. Recognizing of this fact by public may reduce the number of persons sleeping in the same rooms and consequently reduce the risk of ALRTI.

Passive smoking is a recognized risk factor for ALRTI.²⁵ The mechanism behind this is the known inhibitory effects of smoking on ciliary activity and other local protective mechanisms.²⁶ However our risk estimate for lower respiratory tract infections associated with smoking was nearly the same among cases and controls.

The possible explanation is that in our study almost all of the fathers smoke outdoors and none of the mothers were smokers.

This may explain the different results of this study when compared with other study in which maternal smoking is quite high, reach up to 66% in some studies.²⁷

A Similar results to our study has been obtained in Alaska study.²⁸ Which reveal that smoking by a household member was not associated with increased hospitalization risk, in spite that smoking is common (39%) but smoking in house was rarely reported (1%).

All children may be exposed to tobacco smoke, although educational efforts may have reduced smoking inside homes and subsequently reduced the risk of passive smoking as what was noticed in the present study.

Limitations of the present study

Selection of controls, as selection of appropriate controls for cases of severe ALRTI possess several difficulties.

Selection of patients admitted to the same hospital for other diseases as controls may result in biased comparisons as described by Berkson²⁹, subjects selected from the community would have the advantage of being more representative of the general population, but for logistic reason that was impossible in our study.

Therefore we selected controls from vaccination clinics in Tripoli, as the vaccination coverage in Libya is very high³⁰, infants presented to vaccination clinics should be representative for infants in the community.

CONCLUSION

Young infants, young less educated mother, bottle feeding, increased number of households, nighttime crowding, and bottle feeding were the main risk factors for severe lower respiratory tract infections.

RECOMMENDATIONS

Encouragement of breast feeding especially in first 4-6



months by increasing the awareness about benefits of breast feeding during antenatal visits, improving maternal education level, reducing the number of persons sleeping in the same room whenever possible especially with children less than five years of age, finally smoking in the household should be strongly discouraged. Taken together, these measures may reduce the burden of severe ALRTI.

ACKNOWLEDGMENTS

To NCDC (National Center for Disease Control and prevention -Libya) and LEPIDC (Libyan European Partnership for Infectious Disease Control in Libya) team for their support in conducting this study and especially for Dr Paul M. for his enormous advice and assistance in statistical analysis.

REFERENCES

- Nair H, Simoes EA, Rudan I, Gessner BD, Azziz-Baumgartner E, *et al.* (2013) Global and regional burden of hospital admissions for severe acute lower respiratory infections in young children in 2010: a systematic analysis, *Lancet* **381**, 1380-390.
- Kabra SK1, Lodha R, Broor S, Chaudhary R, Ghosh M and Maitreyi RS (2003) Etiology of acute lower respiratory tract infection, *Indian J Pediatric* **70** (1)33-36.
- Jackson S, Mathews KH, Pulanic D, Falconer R, Rudan I, *et al.* (2013) Risk factors for severe acute lower respiratory infections in children: a systematic review and meta-analysis, *Croat Med J* **54**,110-121.
- Wonodi CB, Deloria-Knoll M, Feikin DR, DeLuca AN, Driscoll AJ, *et al.* (2012) Evaluation of risk factors for severe pneumonia in children: The pneumonia etiology research for child health study, *Clinical Infectious Diseases* **54**(2)124-131.
- Izurieta HS, Thompson WW, Kramarz P, *et al* (2000). Influenza and the rates of hospitalization for respiratory disease among infants and young children, *N Engl J Med*, **342**(4), 232-239.
- Behrman S (1991) Epidemiology of acute respiratory infection in children of developing countries, *Rev Infect Dis* **13** (6),454-462.
- Hussey GD, Apolles P, Arendse Z, Yeates J, Robertson A, Swingler G *et al* (2000) Respiratory syncytial virus infection in children hospitalized with acute lower respiratory tract infections, *Afr Med J* **90**,509-512.
- Agrawal PB, Shendurnikar N, Shastri NJ (1995) Host factors and pneumonia in hospitalized children, *J Indian Med Assoc* **93**, 271-272.
- Global Health Risks: Mortality and Burden of Disease Attributable to Selected Major Risks (2009). WHO, Geneva, *World Health Organization*.
- Technical Basis for WHO Recommendations on the Management of Pneumonia in Children at First Level Health Facilities (1991) *WHO/ARI/91.20 Geneva, World Health Organization*.
- Management of the Child With Serious Infection or Severe Malnutrition: Guidelines for Care at the First Referral Level in Developing Countries (2001) WHO Geneva, *World Health Organization*.
- Selwyn BJ (1990) The epidemiology of acute respiratory tract infection in young children: comparison of findings from several developing countries. Coordinated Data Group of BOSTID Researchers, *Rev Infect Dis*. **12**(1 8) ,870-888.
- Graham NM (1990) The epidemiology of acute respiratory infections in children and adults: a global perspective, *Epidemiol Rev*. **12**,149-178.
- Anders Koch, Per Sørensen, Preben Homøe, Kåre Mølbak, Freddy Karup Pedersen *et al.*(2002) Population-Based Study of Acute Respiratory Infections in Children, Greenland, *Emerg Infect Dis* **8**(6),586-593.
- Hajizadeh M, Nandi A and Heymann J (2014) Social inequality in infant mortality: What explains variation across low and middle income countries? *Soc Sci Med* **101**, 36-46.
- Houweling TAJ and Kunst AE (2009) Socio-economic inequalities in childhood mortality in low- and middle-income countries: a review of the international evidence, *British Medical Bulletin* **93**,7-26.
- Wang H, Liddell CA, Coates MM, Mooney MD, Levitz CE, Schumacher AE, *et al.* (2014) Global, regional, and national levels of neonatal, infant, and under-5 mortality during 1990–2013: a systematic analysis for the Global Burden of Disease Study, *Lancet* **384**, 957- 979.
- Victoria CG, Fuchs SC, Flores JAC, Fonseca W and Kirkwood B (1994) Risk factors for pneumonia among children in a Brazilian metropolitan area, *Pediatrics* **93**, 977- 985.
- Prietsch SOM, Fischer GB, César JA *et al.* (2008) Acute lower respiratory illness in under-five children in Rio Grande, Brazil: prevalence and risk factors, *Cadernos de Saúde Pública* **24**,1429-1438.
- Abbi rita, Parul Christian, Sunder Gujral, and Tara Gopaldas (1991) The impact of maternal work status on the nutrition and health status of children, *Food and Nutrition Bulletin* **13**(1), 20-25.
- Sonneveldt E, DeCormier Plosky W and Stover J (2013) Linking high parity and maternal and child mortality: what is the impact of lower health services coverage among higher order births?, *BMC Public Health* **13**(3),7.
- Ratageri VH, Kabra SK, Dwivedi SN and Seth V (2000) Factors associated with severe asthma, *Indian Pediatr* **37**, 1072-1082.
- Kovesi T, Gilbert NL, Stocco C, *et al.* (2007) Indoor air quality and the risk of lower respiratory tract infections in young Canadian Inuit children, *CMAJ* **177**(2), 155-160.
- Banerji A, Greenberg D, White LF, *et al.* (2009) Risk factors and viruses associated with hospitalization due to lower respiratory tract infections in Canadian Inuit children: a case-control study, *Pediatr Infect Dis J* **28** (8), 697-701.
- Wright AL, Holberg C, Martinez FD, *et al.* (1991) Relationship of parental smoking to wheezing and no wheezing lower respiratory tract illnesses in infancy, *J Pediatr* **118**, 207-214.
- Monto AS and Ross H (1977) Acute respiratory illness in the community: effect of family composition, smoking, and chronic symptoms, *Br J Prev Soc Med* **31**(12),101-108.
- Koch. A., *et al.* (2003) Risk factors for acute respiratory tract infections in young Greenlandic children, *Am J Epidemiol* **158**(4), 374-384.
- Lisa R. Bulkow, Rosalyn J. Singleton, Carolyn DeByle, *et al.*(2012) Risk Factors for Hospitalization With Lower Respiratory Tract Infections in Children in Rural Alaska, *Pediatrics* **129** (5),1220-1227.
- Berkson J (1946) Limitations of the application of fourfold table analysis to hospital data, *Biometrics* **2**(3),47-53.
- Libya WHO and UNICEF estimates of national immunization coverage (2015) , pages 1-14.

