# STUDY OF CHILDREN WITH RECURRENT PNEUMONIA ADMITTED IN A TERTIARY HOSPITAL

Rijal P, Lama L, Shrestha S, Kakshapati P, Nayak R

Department of Pediatrics, Nepal Medical College Teaching Hospital, Attarkhel, Gokarneshwor-8, Kathmandu, Nepal.

### ABSTRACT

Recurrent pneumonia usually occurs due to an underlying disorder that negatively affects local or systemic defence mechanisms. The aim of this study was to find out the prevalence and risk factors associated with recurrent pneumonia in children and to determine possible strategic plan needed for better clinical outcome. Children between 2 months to 15 years old who had a history of 2 or more episodes of pneumonia per year, or 3 or more episodes in a life time were investigated prospectively at Nepal Medical College Teaching Hospital. Out of 653 children admitted for pneumonia, 74 (11.3 %%) met the criteria for recurrent pneumonia. Among 74 children with recurrent pneumonia, underlying risk factors was demonstrated in 65 patients (87.8%). Most common underlying diseases were aspiration syndrome in 21.6% patients, congenital heart disease in 13.5% patients and bronchial asthma in 12.1% children. No predisposing illness could be demonstrated in 12.1% patients. Approximately 1 in 9 children with pneumonia in our hospital had recurrent pneumonia. Aspiration syndrome was the most common underlying illnesses for undiagnosed recurrent pneumonia in children.

#### **KEYWORDS**

Aspiration syndrome, bronchiectasis, immune deficiency disorders, recurrent pneumonia

#### **CORRESPONDING AUTHOR**

Dr. Prashant Rijal, Assistant Professor, Department of Pediatrics, Nepal Medical College Teaching Hospital, Attarkhel, Gokarneshwor-8, Kathmandu, Nepal. Email: prashant\_rijal@hotmail.com

#### **INTRODUCTION**

Pneumonia is an acute inflammation of the lung parenchyma that develops due to both infectious and non-infectious causes. Pneumonia is a major problem in children; the World Health Organization (WHO) estimated that pneumonia occurred in approximately 156 million children (151 million in developing countries and 5 million in developed countries).<sup>1</sup> Recurrent pneumonia (RP) (at least 2 pneumonia episodes in 1-year period or more than 3 episodes at any time with radiographic clearing between episodes) was diagnosed in 6.4–9.0% of children with pneumonia.<sup>2-4</sup> Recurrent pneumonia usually result from deficiencies in the local pulmonary or systemic host defences or from underlying disorders that modify the lung defence. Therefore, paediatricians should investigate this possible underlying causes.<sup>5</sup> There are limited data on the underlying diseases predisposing to recurrent pneumonia in children. In addition, few studies have addressed this problem in the developing world.<sup>6-8</sup> The potential burden of RP on the health care system, in terms of frequent hospitalizations, is significant. Early recognition and management of the underlying cause is expected to reduce the number of pneumonia related hospital admissions, morbidity and mortality. Especially in the developing countries, increasing studies of epidemiology to recurrent pneumonia will ease preventative and therapeutic approach. The purpose of this study is to determine the prevalence and risk factors associated with recurrent pneumonia in Nepalese children in a tertiary care centre. Hence, we can make strategic plan to reduce the morbidity and mortality due to recurrent pneumonia.

## **MATERIALS AND METHODS**

It is an observational cross-sectional study conducted at Nepal Medical College Teaching Hospital, Attarkhel, Kathmandu, Nepal. The hospital is situated at Gokarneshwor-8, Nepal about 11 km. northeast of Kathmandu city. All patients with diagnosis of recurrent pneumonia admitted in pediatric ward of NMCTH during July 2013 to June 2018 were included in the study.

Patients who met the diagnostic criteria for RP were included in the study. Pneumonia was diagnosed based on the presence of tachypnea (with a respiratory rate of  $\geq 60$ /min for infants aged <2 months,  $\geq$ 50/min for infants aged 2-11 months,  $\geq$ 40/min for infants aged 12-60 months) and ≥30 in children more than 5 years of age, laboured breathing, bronchial breath sounds or crackles on chest auscultation and/ or radiological findings (pulmonary consolidation or infiltrate).<sup>9</sup> Children with persistent pneumonia and outpatient department (OPD) patients were excluded from the study. Patient demographics (gender, age at diagnosis of RP, age at first episode, number of episodes per year, and number of episodes prior to diagnosis of RP) and details pertaining to the aetiology of and risk factors for RP (presenting symptoms, history of mechanical ventilation, foreign

object aspiration, frequent infections, prematurity, vaccination history, number of household members, duration of breast feeding, parental consanguinity, family history of atopy or other chronic disorders, and exposure to tuberculosis) were recorded for each patient on forms. Physical examination findings and the results of routine and specific diagnostic tests (chest X-ray, purified protein derivatives (PPD) skin test, hemosiderin-laiden macrophages in broncho alveolar lavage, immunological evaluation, barium meal, gastroesophageal reflux scintigraphy, thoracic CT, and acid-fast staining of fasting gastric aspirate) were also recorded. Ethical clearance was taken from the Institutional Review Committee of Nepal Medical College Teaching Hospital. Data were taken from the patients' pro forma files. The data was analysed using SPSS version 16.

## **RESULTS**

Among the 653 patients with a diagnosis of pneumonia during the study period, 74 (11.3%) were diagnosed as RP. In this group, 47 (63.5%) were male and 27 (36.4%) were female (Fig. 1).



Fig. 1: Sex-wise distribution of recurrent pneumonia

The most common presenting symptom was cough in 98.3%, followed by fever 63.5%, fast breathing in 52.7% patients. On examination 52.7% had tachypnoea, 52.7% cases had crackles, 32.4% cases had wheezing, 51.3% cases had decreased breath sound, 10.8 cases had clubbing, and 1.3% case had hepatosplenomegaly. Chest X-rays was done in all cases and in 49 (66.2%) cases chest CT-scans was done. Chest X-ray showed consolidation in 43 (58.1%), infiltration 17 (22.9%) and lymphadenitis in 5.4% cases. CT scan chest revealed infiltration in 19 (25.6%) cases, consolidation in 36 (48.6%), bronchiectasis in 8 (10.8%), air trapping in 7 (9.4%), atelectasis in 5 (6.7%), pleural effusion in 3 (4.0%), interstitial lung disease in 1 (1.3 %%) cases. Here, 16 (21.6%) patients had aspiration syndrome out of which 6 (37.5%) children had GERD and each 5 (31.2%) children had cerebral palsy and foreign body aspiration.10 (13.5%) patients had congenital heart disease, out of which 6 (60.0%) children had VSD, 2 (20.0%) patients had ASD and one of each had PDA

and complex heart disease. Nine (12.1%) patients had bronchial asthma. In each 8 (10.8%) patients, they had immune deficiency disorders, bronchiectasis and congenital anomalies of respiratory system. In primary immune deficiency disorders 4 (50.0%) children had common variable immunodeficiency, T-cell deficiency in 2 (25.0%) patients and natural killer cell deficiency in 2 (25.0%) patients. Congenital anomalies are an important underlying illness for recurrent pneumonia in children. Among congenital anomalies of respiratory system, 3 (37.5%) patients had tracheoesophageal fistula, 1 (12.5%) of each had tracheal stenosis, tracheal bronchus, agenesis of right pulmonary artery, hypoplasia of right lung, congenital cystoadenomatoid malformation. One (1.3%) of each patient diagnosed as interstitial lung disease and pulmonary nocardiosis. No predisposing illness could be demonstrated in the remaining 9 (12.1%) patients (Table-1).

Table-1: Distribution of patients with RP according to underlying disorder/ risk factors.		
77 1 1 · . 711	<b>Recurrent Pneumonia</b>	
Underlying Illness	No. of cases	% of cases
Bronchial Asthma	9	12.1
Bronchiectasis	8	10.8
Aspiration Syndrome		
GERD	6 (37.5%)	
Cerebral Palsy	5 (31.25%)	21.6
Foreign Body Aspiration	5 (31.25%)	
Total	16	
Immunodeficiency		
Primary		
Common Variable	8 (66.66%)	
Immunodeficiency	4 (33.33%)	16.2
T Cell deficiency	2 (16.66%)	10.2
NK Cell deficiency	2 (16.66%)	
Secondary (HIV)	4 (33.33%)	
Total	12	
Congenital Heart Disease		
VSD	6 (60%)	
ASD	2 (20%)	
PDA	1 (10%)	13.5
Complex Heart Disease	1 (10%)	
Total	10	
Congenital Anomalies of Respiratory system		
Tracheo-esophageal Fistula	3 (37.5%)	
Agenesis of right pulmonary artery	1 (12.5%)	
Hypoplasia of Right lung		
Tracheal Bronchus	1 (12.5%)	10.8
Tracheal Stenosis	1 (12.5%)	
Congenital cystoadenomatoid malformation	1 (12.5%)	
Total	1 (12.5%)	
Pulmonary Nocardiosis	1	1.35
Interstitial Lung Disease	1	1.35
Unidentified Cause	9	12.1
Total cases	74	100

#### DISCUSSION

The respiratory tract is protected by its local and general body defences. Nevertheless, respiratory tract infections constitute a major health problem all over the world.<sup>10</sup> The etiology of RP varies depending upon the location of pneumonia, which can involve a single or various lobes or lung regions. Recurrent pneumonia involving a single lobe or segment, is caused by localized pathology, such as local compression, malformation or inflammation, whereas, pneumonias affecting more than one lobe, suggest a more generalized abnormality, such as mucociliary clearance dysfunction, aspiration, immunodeficiency or asthma.<sup>11</sup> In our study, 11.3% of admission to the hospital with a diagnosis of pneumonia met the definition of recurrent pneumonia. It is slightly higher than other studies where 6.4%-9.0% of patients met the criteria for recurrent pneumonia.<sup>12,13</sup>

An underlying illness for recurrent pneumonia was found in 87.8% of our patients. Similar to Weigl *et al* data where cause for recurrent pneumonia was found in 83.0% of children.<sup>14</sup> We did not identify the cause for RP in the remaining 12.1% of our patients. Comparatively, unknown aetiology of patients with recurrent pneumonia was found in 13.3% by Cabezuelo *et al.*<sup>15</sup> The incidence and mortality rate of pneumonia in children are 10 times higher in developing countries than developed countries. Additionally, it is important to note that lack of epidemiological report from developing countries make it difficult to design global strategies for prevention, diagnosis, and treatment of RP.<sup>16</sup>

The most frequent underlying illness was aspiration syndrome found in 21.6% patients followed by congenital heart disease in 13.5% and bronchial asthma in 12.1% patients at our centre. Study done by Hovsing *et al* showed recurrent aspiration in 25.7% patients, lung disease (airway stenosis, bronchiectasis, middle lobe syndrome or tracheoesophageal fistula) in 16.1% patients and immune deficiency in 16.1% patients.<sup>17</sup>

Among the aspiration syndrome, gastroesophageal reflux was seen in 37.5% patients. Each 31.5% patients had oropharyngeal incoordination due to cerebral palsy and foreign body aspiration. These illnesses were confirmed by barium esophagography, milk scan and flexible bronchoscopy. Bolursaz et *al*<sup>18</sup> showed aspiration syndrome in 51.7% patients among which GERD in 90.1% patients, cerebral palsy in 7.8% patients and foreign body aspiration in 3.9% patients. Here, in 13.5% of patients with RP had congenital heart disease where Ventricular septal defect was most commonly diagnosed in 60.0% of children. Data analysed by Lodha et al<sup>19</sup> revealed that 9.0% of children with RP had underlying congenital heart disease which is lower than our study. Bronchial asthma is an important underlying illness for recurrent pneumonia in children. In this study, 12.1% of children with bronchial asthma had RP. Despite recent advances in healthcare, many children referred with recurrent pneumonia are

shown to have undiagnosed asthma. Asthma has been reported as an important underlying cause of recurrent pneumonia in children.<sup>20</sup> In our study, 10.8% of the patients were diagnosed with primary immune deficiency disorders, including common variable immunodeficiency in 50.0% patients, T-cell deficiency in 25.0% patients and natural killer cell deficiency in 25.0% patients. Immune deficiency should be suspected in children with infections that are especially severe and recurrent, that are caused by unusual organisms, or that involve multiple sites in addition to the lungs. A study done by Capanoglu et al<sup>21</sup> showed that 4.0% children with RP had primary immunodeficiency which slightly lower than our data. This may be due to more number of referral cases in our centre.

Bilateral interstitial pneumonias on chest radiograph films and/or the presence of opportunistic infections were paramount in diagnosing HIV infection as an underlying factor in recurrent pneumonia. Similarly, HIV infection was found in 5.4% of children.<sup>22</sup>

Bronchiectasis was present in 10.8% cases. Past history of treatment for pulmonary tuberculosis was present in 62.5% cases while in 37.5%. In cases predisposing factor could not be identified. A data of 60.7% cases of bronchiectasis were analysed by Pifferi *et al*<sup>23</sup> which is very high as compared to our study. This may be due to less number of cases in our study.

Congenital anomalies are considered as an important underlying illness for recurrent pneumonia in children. We confirmed congenital anomaly of respiratory system in 10.8% of children. Three patients had tracheoesophageal fistula, one of each had tracheal stenosis, tracheal bronchus, agenesis of right pulmonary artery, hypoplasia of right lung, congenital cystoadenomatoid malformation. Congenital anomaly of respiratory system usually manifest during infancy as described by in this study.<sup>24</sup> Patria *et al*<sup>25</sup> analysed that the significant risk factors for recurrent pneumonia were GERD, a history of atopy and/or allergy, and asthma. In our study, aspiration syndrome and congenital heart disease were the important risk factors for recurrent pneumonia.

Our study had some limitations. As we included only hospitalized patients, the true number of patients with this condition may have been understimated.

In conclusion, recurrent pneumonia is still a diagnostic challenge in Pediatrics. Many children with RP need thorough investigations. In undiagnosed recurrent pneumonia, bronchial asthma, immune deficiency disorders, congenital heart disease, and gastroesophageal reflux should be considered in the differential diagnosis. Early treatment of the child's underlying condition is crucial in order to stabilize lung disease and thus prevent progressive pulmonary function deterioration.

#### REFERENCES

- 1. World Health Organization. Global Tuberculosis Report 2016. Available from URL: http://www.who. int/tb/publications/global\_report/en (Accessed: March 2017).
- 2. Wald E. Recurrent and non-resolving pneumonia in children. *Semin Respir Infect* 1993; 8: 46-58.
- 3. Lodha R, Puranik M, Natchu UC, Kabra SK. Recurrent pneumonia in children: Clinical profile and underlying causes. *Acta Paediatr* 2002; 91: 1170-3.
- 4. Owayed AF, Campbell DM, Wang EE. Underlying causes of recurrent pneumonia in children. *Arch Pediatr Adolesc Med* 2000; 154: 190-4.
- 5. Sheares BJ. Recurrent pneumonia in children. *Pediatr Ann* 2002; 31: 109-14.
- Kumar M, Biswal N, Bhuvaneswari V, Srinivasan S. Persistent Pneumonia: Underlying Cause and Outcome. *Indian J Pediatr* 2009; 76: 1223-6.
- Ozdemir O, Sari S, Bakirtas A, Zorlu P, Ertan U. Underlying diseases of recurrent pneumonia in Turkish children. *Turk J Med Sci* 2010; 40: 25-30.
- 8. Lodha R, Puranik M, Chandra U, Natchu M, Kabra SK. Persistent pneumonia in children. *Indian Pediatr* 2003; 40: 967-70.
- 9. McIntosh K. Community-acquired pneumonia in children. *N Engl J Med* 2002; 346: 429-37.
- 10. Verhagen LM, de Groot R. Recurrent, protracted and persistent lower respiratory tract infection: A neglected clinical entity. *J Infect* 2015; 1: 106-11.
- 11. Mantzouranis E, Mathianaki K, Fitrolaki MD, Mihailidou E, Paspalaki P. A boy with recurrent pneumonia *Breathe* 2011; 7: 367-70.
- 12. Couriel J. Assessment of the child with recurrent chest infections. *Br Med Bull* 2002; 61: 115-32.
- 13. Çiftçi E, Güneş M, Köksal Y, İnce E, Doğru Ü. Underlying causes of recurrent pneumonia in Turkish children in a university hospital. *J Trop Pediatr* 2003; 49: 212-5.
- 14. Weigl JA, Bader HM, Everding A, Schmitt HJ. Population-based burden of pneumonia before school entry in Schleswig-Holstein, Germany. *Eur J Pediatr* 2003; 162: 309–16.

- 15. Cabezuelo HG, Vidal MS, Abeledo GA, FronteraIzquierdo P. Underlying causes of recurrent pneumonia. *Ann Pediatr* 2005; 63: 409-12.
- 16. Khaled S, Sherif AM, Kotb AM. Recurrent/Persistent pneumonia among Children in Upper Egypt. *Mediterr J Hematol Infect Dis* 2013; 5: 28.
- 17. Hoving MF, Brand PL. Causes of recurrent pneumonia in children in a general hospital. *J Paediatr Child Health* 2013; 49: 208–12.
- 18. Bolursaz MR, Lotfian F, Ghaffaripour HF, Hassanzad M. Underlying causes of recurrent and persistent pneumonia in children at a pulmonary referral hospital in Tehran, Iran. *Arch Iran Med* 2017; 20: 266-9.
- 19. Lodha R, Kabra SK. Recurrent/persistent pneumonia. *Indian Pediatrics* 2000; 37: 1085-92.
- Eigen H, Laughlin JJ, Homrighausen J. Recurrent pneumonia in children and its relationship to bronchial hypereactivity. *Pediatri* 1982; 70: 698– 704.
- 21. Çapanoglu M, Zorlu P, Sari E, Senel S. The etiology of recurrent pneumonia with onset during infancy, and the effect of risk factors on age at first episode and episode frequency. *Turkish J Pediatr Dis* 2017; 4: 243-7.
- 22. Beard LJ, Maxwell GM, Thong YH. Immunocompetence of children with frequent respiratory infections. *Arch Dis Child* 1981; 56: 101-5.
- 23. Pifferi M, Caramella D, Ragazzo V, Cangiotti AM, Macchia P, Boner AL. Bronchiectasis in children with recurrent pneumonia: an immunopathological damage associated with secondary ciliary dysmotility. *Int J Immunopathol Pharmacol* 2008; 21: 215-9.
- 24. Rubin BK. The evaluation of the child with recurrent chest infections. *Pediatr infect Dis J* 1985; 4: 88-98.
- 25. Patria F, Longhi B, Tagliabue C *et al*. Clinical profile of recurrent community- acquired pneumonia in children. *BMC Pulm Med* 2013; 10: 13-60.