Clinical predictors of radiographic abnormalities among infants with bronchiolitis in a paediatric emergency department

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Abstract

Background: Acute viral respiratory exacerbation is one of the most common conditions encountered in a paediatric emergency department (PED) during winter months. We aimed at defining clinical predictors of chest radiography prescription and radiographic abnormalities, among infants with bronchiolitis in a paediatric emergency department.

Methods: We conducted a prospective cohort study of children less than 2 years of age with clinical bronchiolitis, who presented for evaluation at the paediatric emergency department of an urban general hospital in France. Detailed information regarding historical features, examination findings, and management were collected. Clinical predictors of interest were explored in multivariate logistic regression models.

Results: Among 410 chest radiographs blindly interpreted by two experts, 40 (9.7%) were considered as abnormal. Clinical predictors of chest radiography achievement were age (under three months), feeding difficulties, fever over 38°C, hypoxia under than 95% of oxygen saturation, respiratory distress, crackles, and bronchitis rales. Clinical predictors of radiographic abnormalities were fever and close to significance hypoxia and conjunctivitis.

Conclusion: Our study provides arguments for reducing chest radiographs in infants with bronchiolitis. For infants with clinical factors such as age less than three months, feeding difficulties, respiratory distress without hypoxia, isolated crackles or bronchitis rales, careful clinical follow-up should be provided instead of chest radiography.

Keywords: Bronchiolitis, Chest radiography, Radiographic pneumonia, Clinical decision rule

Background

Acute viral respiratory exacerbation is one of the most common conditions encountered in a paediatric emergency department (PED) during winter months [1]. It is often difficult to distinguish viral from bacterial cause of dyspnea with only clinical features. The lack of reliable clinical decision rule for management of bronchiolitis often leads to perform chest radiography (CR) in order to detect radiographic abnormalities inconsistent with this diagnosis, such as pneumonia or other cardio-respiratory disease. An American survey of clinical practices in the diagnosis and management of bronchiolitis showed a CR rate of 72% that lead to an increased likelihood of antibiotic therapy and length of stay in hospital [2]. Recent studies showed that most of these CR were read as negative, (ie consistent with a simple acute viral respiratory exacerbation), and that many of them might have been avoided, saving time, money, and children exposure to ionizing radiations [3,4].

The aims of our study were to investigate historical features and clinical examination findings in the evaluation of infants with bronchiolitis that conduct to the prescription of CR, and to determine clinical predictors of radiographic abnormalities.

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Methods

Study design and setting
The study was approved by the ethical committee CPP (Comité de Protection des Personnes) – Ile de France 3. We performed a prospective cohort study of children less than 2 years of age with clinical bronchiolitis, who presented for evaluation to the PED of an urban general hospital (Antoine Beclere – Clamart, Paris suburb, France) between October 2006 and February 2007. According to the American Academy of Pediatrics, bronchiolitis was defined as a constellation of clinical symptoms and signs including a viral upper respiratory prodrome followed by increased respiratory effort and wheezing in children less than 2 years of age [5]. All children who had clinical bronchiolitis on examination were eligible to the study. All physicians working in the PED were asked and accepted to participate. Physicians were oriented to the questionnaire before the start of the study and were continually informed of study details throughout the study period. They completed a questionnaire for each patient included. To prevent free text responses, fixed-choice format has been chosen.

Clinical data collection
Detailed information regarding historical features and examination findings at first evaluation were collected. Demographics data included age, gender, history of prematurity, known chronic illness, and previous wheezing episodes. Historical features of interest included duration of symptoms, history of fever, feeding difficulties, and antibiotics use before PED admission. Physicians were asked to report temperature, respiratory rate, retraction signs and wheezing, oxygen saturation, auscultation findings (bronchitis rales, crackles) and the presence of otitis, conjunctivitis or toxic appearing. Clinicians recorded the following management: CR, antibiotic prescription, and decision of hospitalisation.

Chest radiography interpretation
All CR were performed by using standard equipment and radiographic techniques. They were blindly and independently reviewed in digital format by 2 paediatric radiologists. CR was read as either normal or abnormal. Abnormalities were lobar consolidation, segmental or lobar atelectasia, or cardiomegaly. Others findings (prominent bronchial opacities, peribronchial infiltrates, hyperinflation or sub-segmental atelectasia) were considered as consistent with the diagnosis of bronchiolitis and therefore had no impact on therapeutic decision. For CR whose interpretation differed between both radiologists, we organized a consensus meeting.

Statistical analysis
Analyses were performed on the whole study population to assess the relationship between history and clinical findings and the decision to realise CR or not. Clinical predictors of radiographic abnormality were studied on patients that underwent CR. Continuous and ordinal variables were categorized: temperature was coded as \(<38°C, 38 – 39°C or \geq 39°C\), hypoxia was defined by an oxygen saturation \(<95%\), age-related respiratory rate was defined in accordance with Liu definition (Table 1) [6]. We created a global score of respiratory distress, using weighted retraction signs widely used and that showed a good inter-observer reproducibility (Tables 2 and 3) [7]. Variables with a \(p\)-value \(\leq 0.20\) in univariate analysis were considered candidate variables for inclusion in the multivariate model and \(p\)-values \(\leq 0.05\) were considered statistically significant.

Because CR is a frequent event (about 50%), oddsratio substantially overestimate the relative risk. We used then Poisson regression to estimate relative risk, first for performing CR, and then for radiographic abnormalities [8]. To enhance power of our analysis, we used multiple imputation for missing values [9-11].

All analyses were performed on Statata 11 Software (StataCorp. 2007. Stata Statistical Software: Release 11. College Station, TX: StataCorp LP).

Results

Demographics data and clinical findings
Study forms were completed for 821 patients (Table 4). One hundred–and-seventy infants (21%) were less than three months of age, 13% had prematurity history, and 60% had no wheezing history before this episode. The mean duration of symptoms was 4.1 days (SD 5.3 days), 38% of children had feeding difficulties and about 15% had antibiotic prescription before the emergency admission. Clinically, about half of children were febrile, 105 (14.1%) had oxygen saturation under 95%, 272 (33%) had bronchitis rales and 104 (12.7%) crackles. 427 (52%) of the children have had a chest radiography and 350 (42.6%) had been hospitalized. One hundred and four (12.7%) children had otitis, 27 (3.3%) conjunctivitis. 215 (26%) received antibiotics (13.5% of children without CR versus 38.8% of children with CR, \(p < 0.001\)).

Chest radiographs
Among the 427 chest radiographs performed, 410 were interpreted by both experts. Two chest radiographs were excluded because of their poor quality and 15 could not

| Table 1 Age related respiratory rate from Liu et al. [6] |
|------------------------------------------------------|
| **Respiratory rate (l/min)** | 1 point | 2 points | 3 points |
| Age < 2 months | ≤60 | 61-69 | ≥70 |
| Age 2 – 12 months | ≤50 | 51-59 | ≥60 |
| Age 12 – 24 months | ≤40 | 41-44 | ≥45 |
be retrieved (Figure 1). Consistency rate between both blinded radiologists was 84.3%.

Most infants had prominent bronchial opacities, peri-bronchial infiltrates and/or hyperinflation. These CR were considered to be consistent with the diagnosis of bronchiolitis or viral acute exacerbation. Forty (9.7%) CR were considered as abnormal: 39 with lobar or alveolar condensations and one with lobar atelectasia.

CR achievement
Univariate analysis identified nine variables that were significantly related to the realisation of a chest radiography: age ≤ 3 months (p = 0.03), feeding difficulties (p < 0.001), fever (p < 0.001), tachypnea (p = 0.002), global score of respiratory distress (p < 0.001), hypoxia (SpO2 < 95%, p < 0.001), crackles (p < 0.001), bronchitis rales (p = 0.04) and toxic appearing (p = 0.04). All these variables were included in the multivariate model (Table 5) which found a significant association between the realisation of CR and age ≤ 3 months (p = 0.003), feeding difficulties (p = 0.01), presence of fever (p < 0.001), hypoxia (p = 0.04), global score of respiratory distress (p = 0.03), crackles (p = 0.003) or bronchitis rales (p = 0.004) (Table 4).

Radiographic abnormalities
Univariate analysis only identified fever as significantly related to the presence of radiographic abnormality (p = 0.02). Finally, multivariate analysis only identified fever as an independent clinical predictor of radiographic abnormalities (p = 0.04) (Table 6). Presence of crackles was not an independent predictor in our population (p = 0.1).

Discussion
To our knowledge, this study is one of the largest prospective cohorts of infants under two years of age who have been admitted to a PED for bronchiolitis. We have been able to determine clinical predictors of CR prescription and radiographic abnormalities. We observed a high rate of CR (52.7%) with a rate of abnormalities of

Table 2 Clinical score of weighted retraction signs (total /8)

|                      | None | Moderate | Important |
|----------------------|------|----------|-----------|
| Intercostal indrawing| 0    | 1        | 2         |
| Sub-costal indrawing | 0    | 1        | 2         |
| Nasal flaring        | 0    | 1        | 2         |
| Thoraco-abdominal movement| 0 | 1        | 2         |

Table 3 Global score of respiratory distress

| Global score of respiratory distress | 0 point | 1 point | 2 points | 3 points |
|--------------------------------------|---------|---------|----------|----------|
| Clinical score of retraction signs   | 0       | 1 or 2  | 3 or 4   | 5 to 8   |

Table 4 Description of the study population (n = 821)

|                                | N      | n (%)  |
|--------------------------------|--------|--------|
| First wheezing episode         | 801    | 492 (59.9) |
| Sex (male/female)              | 821    | 476/345 |
| Age under 3 months            | 821    | 170 (20.7) |
| Prematurity <37 weeks          | 780    | 108 (13.1) |
| Respiratory symptom length (j) | 806    | 4.1 (5.3) |
| Feeding difficulties           | 799    | 309 (37.6) |
| Temperature                    |        |        |
| < 38°C                         | 738    | 373 (45.4) |
| [38-39°C]                      | 211    | 25.7 |
| ≥ 39°C                         | 154    | 18.8 |
| Previous antibiotherapy        | 816    | 124 (15.1) |
| Age-related respiratory rate   | 747    |        |
| 1 point                        | 427    | 52.0 |
| 2 points                       | 101    | 12.3 |
| 3 points                       | 219    | 26.7 |
| Global score for retraction signs | 798 |        |
| 0 point                        | 233    | 29.2 |
| 1 point                        | 454    | 56.9 |
| 2 points                       | 83     | 10.4 |
| 3 points                       | 28     | 3.5 |
| Oxygen saturation              | 747    |        |
| ≥95%                           | 642    | 85.9 |
| <95%                           | 105    | 14.1 |
| Crackles                       | 814    |        |
| 81                             | 9.9 |
| Bronchitis rales               | 814    | 272 (33.1) |
| Otitis                         | 789    | 104 (12.7) |
| Conjunctivitis                 | 802    | 27 (3.3) |
| Toxic appearing                | 817    | 35 (4.3) |
| Chest radiography              | 821    | 427 (52.0) |
| Hospitalisation                | 821    | 350 (42.6) |

Discussion
To our knowledge, this study is one of the largest prospective cohorts of infants under two years of age who have been admitted to a PED for bronchiolitis. We have been able to determine clinical predictors of CR prescription and radiographic abnormalities. We observed a high rate of CR (52.7%) with a rate of abnormalities of

Figure 1 Description of chest radiographs in the study.
Table 5 Factors associated with CR achievement: multivariate analysis

| Factor                                | RR  | IC.95          | p   |
|---------------------------------------|-----|----------------|-----|
| Age ≤ 3 months                        | 1.4 | [1.1 – 1.8]    | 0.003 |
| Prematurity                           | 1.3 | [1.0 – 1.6]    | 0.08 |
| Feeding difficulties                  | 1.3 | [1.1 – 1.6]    | 0.01 |
| Fever                                 |     |                |     |
| [38-39°C]                             | 1.4 | [1.1 – 1.8]    | <0.001 |
| ≥ 39°C                                | 1.9 | [1.5 – 2.5]    |     |
| SpO2 < 95%                            | 1.3 | [1.0 – 1.7]    | 0.04 |
| Age related respiratory rate*         |     |                |     |
| score = 2                             | 1.2 | [0.9 – 1.7]    | 0.2  |
| score = 3                             | 1.2 | [1.0 – 1.5]    |     |
| Global score of respiratory distress  |     |                |     |
| score = 1                             | 1.4 | [1.1 – 1.8]    | 0.03 |
| score = 2                             | 1.7 | [1.2 – 2.4]    |     |
| score = 3                             | 1.7 | [1.1 – 2.8]    |     |
| Toxic-appearing                       | 1.0 | [0.6 – 1.5]    | 0.9  |
| Crackles                              | 1.5 | [1.2 – 2.0]    | 0.003 |
| Bronchitis rales                      | 1.4 | [1.1 – 1.7]    | 0.004 |

*According to Liu et al. [6].

9.7% (4.9% of the whole study population). Our study found a great difference between clinical predictors of CR prescription (age ≤ 3 months, feeding difficulties, fever, presence of retraction signs, hypoxia, crackles, bronchial rales and toxic appearing) and the only clinical factor predictive of radiographic abnormalities (fever). Others clinical factors such hypoxia and conjunctivitis were not predictive of radiographic abnormalities.

The prevalence of pneumonia in previous investigations varied widely, ranging from 0.75 to 63% [3,4,12-19]. These variations may be due to several factors (age of patients, inclusion of patients with or without history of wheezing, percentage of children who underwent CR, considered abnormalities). Wide variations in prescription rates of CR (ranging from 42 to 72 [2,15-17]) reflect lack of consensus, even if recent guidelines recommend to limit their routine use [17]. Limiting the number of CR is important for several reasons: although the radiation associated with achieving a CR is negligible (0.02 mSv; whereas natural exposure is estimated at about 0.05 mSv per week), a recent report on the French population exposure to ionizing radiation related to acts of medical diagnosis, reports a number of CR equal to 0.2 acts per year per child under one year (approximately 160,000 procedures per year for a country with 800,000 births per year) [20]. Furthermore, CR achievement appears to be associated with an increased likelihood of prescribing unnecessary antibiotic [2,18,21]. To reduce the number of unnecessary CR, it is important to understand which clinical variables are associated with the realisation of this diagnostic test, and to compare these clinical variables with those associated with radiographic abnormalities. In our study, clinical features such as age (less than 3 months), hypoxia, conjunctivitis, feeding difficulties, retraction signs, and presence of bronchitis rales and crackles were independent clinical predictors of CR, while these variables did not appear to predict a greater risk of radiographic abnormalities. Thus, our results encourage to restrict CR prescription to infants with fever. Those clinical predictors of radiographic abnormalities are consistent with previous studies [3,4,12,13,16-18]. Neuman et al. conducted a large prospective cohort study to assess the relation between historical features and physical examination findings and radiographic pneumonia [4]. Fever, duration of fever, hypoxia and focal rales emerged as significant predictors of pneumonia on the subgroup of patients younger than five years. Furthermore they attempted to stratify children in low and high risk for pneumonia by recursive partitioning analysis but they were unable to characterize a low risk population among children of this subgroup.

As recommended, no microbiological testing has been performed in these patients. Indeed, the question of the benefit of their implementation for limiting antibiotics prescriptions or chest radiographs arises (even if a secondary bacterial infection may be a complication of an authentic viral bronchiolitis and that the presence of RSV does not exclude a bacterial superinfection).

Our study had several limitations. First it was a monocentric study and our results could be considered representative of all PED. As CR has only been performed in 52% of infants, the scope of our assessment of factors associated with radiographic abnormalities was limited. However, the number of CR studied (410) represented a large population compared with previous studies. Moreover, given the current data in the literature and recommendations, it would be unethical to practice systematic CR in bronchiolitis. Finally, our population was heterogeneous regarding the episode of bronchiolitis considered for our study: 40% of the patients already had a wheezing episode. Our work reflected current practices in emergency room.

Table 6 Clinical predictors of radiographic abnormalities: multivariate analysis

| Factor                  | RR  | IC.95          | p   |
|-------------------------|-----|----------------|-----|
| Fever                   |     |                |     |
| [38-39°C]               | 1.1 | [0.4 – 2.7]    | 0.04 |
| ≥ 39°C                  | 2.4 | [1.1 – 5.1]    |     |
| SpO2 < 95%              | 1.8 | [0.9 – 3.5]    | 0.08 |
| Conjunctivitis          | 2.7 | [0.9 – 7.9]    | 0.07 |
| History of wheezing     | 1.5 | [0.8 – 2.8]    | 0.3  |
| Crackles                | 1.7 | [0.8 – 3.4]    | 0.1  |
Conclusion
Our study provides arguments for reducing CR achievement in infants admitted to PED for bronchiolitis. Especially, clinical factors such as age less than three months, feeding difficulties, presence of retraction signs without hypoxia, or isolated rales and crackles are not considered as predictive of radiographic abnormalities and should not lead to the prescription of CR. For those patients, careful clinical follow-up is recommended as an alternative. Moreover further studies are required to evaluate the real link between the achievement of CR and antibiotics prescription.

Abbreviations
CR: Chest radiography; PED: Pediatric emergency department.

Competing interests
The authors declare that they have no competing interests.

Authors’ contributions
EDE, JB, RE, PL, and VG conceived the study and its design. MB and FP coordinated inclusions in the study. JDL and HDL reviewed all chest X-rays. EDE, JB, PL, and VG helped drafting the manuscript. All authors read and approved the final manuscript.

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References
1. Alpern ER, Stanley RM, Gorelick MH, Donaldson A, Knight S, Teach SJ, Singh T, Mahajan P, Goepp JG, Kuppermann N, Dean JM, Chamberlain JM: Pediatric Emergency Care Applied Research Network: Epidemiology of a pediatric emergency medicine research network: the PECARN core data project. Pediatr Emerg Care 2006, 22(10):689–699.
2. Christakis DA, Cowan CA, Garrison MM, Molteni R, Marcuse E, Zerr DM: Variation in inpatient diagnostic testing and management of bronchiolitis. Pediatrics 2005, 115(4):878–884.
3. Mathews B, Shah S, Cleveland RH, Lee EY, Bachur RG, Neuman MI: Clinical predictors of pneumonia among children with wheezing. Pediatrics 2009, 124(1):e29–e36.
4. Neuman MI, Monuteaux MC, Scully KJ, Bachur RG: Prediction of pneumonia in a pediatric emergency department. Pediatrics 2011, 128(2):246–253.
5. Zorc JJ, Hall CB: Bronchiolitis: recent evidence on diagnosis and management. Pediatrics 2010, 125(2):342–9.
6. Liu L, Gallaher MA, Davis RL, Rutter CM, Lewis TC, Marcuse EK: Use of a respiratory clinical score among different providers. Pediatr Pulmonol 2004, 37(3):243–248.
7. Gajdos V, Beydon N, Bommenel L, Pellegrino B, de Pontual L, Bailleux S, Labrune P, Bouyer J: Inter-observer agreement between physicians, nurses, and respiratory therapists for respiratory clinical evaluation in bronchiolitis. Pediatr Pulmonol 2009, 44(8):754–762.
8. McNutt LA, Wu C, Xue X, Hafner JP: Estimating the relative risk in cohort studies and clinical trials of common outcomes. Am J Epidemiol 2003, 157(10):940–943.
9. Royston P: Multiple imputation of missing values: update. Stata J 2005, 5(2):188–201.
10. Rubin D: Multiple Imputation for Nonresponse in Surveys. New York: Wiley; 1987.
11. Van Bauren S, Boshuizen H, Knook D: Multiple imputation of missing blood pressure covariates in survival analysis. Stat Med 1999, 18(6):681–694.
12. Farah MM, Padgett LB, McLaren DJ, Sullivan KM, Simon HK: First-time wheezing in infants during respiratory syncytial virus season: chest radiograph findings. Pediatr Emerg Care 2002, 18(6):333–336.
13. Garcia Garcia ML, Calvo Rey C, Quevedo Teruel S, Martinez Perez M, Sanchez Ortega F, Martin del Valle F, Verjanco Sanchez F, Perez-Brena P: Chest radiograph in bronchiolitis: is it always necessary? An Pediatr (Barc) 2004, 61(3):219–225.
14. Genthell JC, Goldmann HS, Stein RE, Shelov SP, Zipkowsk M: The usefulness of chest radiographs in first asthma attacks. N Engl J Med 1983, 309(6):336–339.
15. Kneyber MC, Moons KG, Van HA: Predictors of a normal chest x-ray in respiratory syncytial virus infection. Pediatr Pulmonol 2001, 31(4):277–283.
16. Mahabee-Gittens EM, Bachman DT, Shapiro ED, Dowd MD: Chest radiographs in the pediatric emergency department for children < or = 18 months of age with wheezing. Clin Pediatr 1999, 38(7):395–399.
17. Roback MG, Dreitlein DA: Chest radiograph in the evaluation of first time wheezing episodes: review of current clinical practice and efficacy. Pediatr Emerg Care 1998, 14(3):181–184.
18. Schuh S, Latali A, Allen U, Manson D, Babyn P, Stephens D, MacPhee S, Mokansi M, Kahanin S, Dick P: Evaluation of the utility of radiography in acute bronchiolitis. J Pediatr 2007, 150(4):429–433.
19. Walsh-Kelly CM, Kim MK, Hennes HM: Chest radiography in the initial episode of bronchopasm in children: can clinical variables predict pathologic findings? Ann Emerg Med 1996, 28(4):391–395.
20. Etard C, Sinno-Tellier S, Aubert B: Exposition de la population française aux rayonnements ionisants liés aux actes de diagnostic médical en 2007. Institut de veille sanitaire 2010, http://www.inv.sante.fr/publications/2010/rayonnements_ionisants_diagnostic_medical/rapport_expiri.pdf.
21. Swingler GH, Hussey GD, Zwarenstein M: Randomised controlled trial of clinical outcome after chest radiograph in ambulatory acute lower-respiratory infection in children. Lancet 1998, 351(9090):404–408.