Assessment of hyperinflation in children with cystic fibrosis

J Marchant, D M Hansell, A Bush

Abstract

Background – Radiological estimates of hyperinflation are used in several clinical and radiographic scoring systems for cystic fibrosis, but it is not known if these estimates of hyperinflation are related to measured total lung capacity.

Methods – Comparison was made of independent clinical estimates of hyperinflation from chest radiographs with objective plethysmographic and radiographic measurements of total lung capacity in 25 children with cystic fibrosis.

Results – There was good agreement between plethysmographic and radiographic measurements. Clinical estimation correctly predicted the extremes of hyperinflation, but grading was no more than 50% accurate in all other groups.

Conclusion – The degree of hyperinflation cannot be estimated by inspecting chest radiographs in many children with cystic fibrosis. This does not invalidate the scoring systems, but suggests that a better term than “hyperinflation” should be sought.

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The commonly used quantitative scoring systems for the chest radiograph in cystic fibrosis require scoring of the degree of hyperinflation.¹² For example, the Brasfield score² grades air trapping from 0 (absent) to 4 (most severe) on the basis of overdistension manifest by sternal

bowing, depression of diaphragms, and/or thoracic kyphosis. However, no study has compared estimates of hyperinflation with simultaneous objective lung function data to see if such estimates are valid.

Methods

Twenty five children (16 boys) of mean age 11.4 (range 5.9-16.6) years from the paediatric cystic fibrosis clinic were studied. The diagnosis had been established on the basis of a compatible history and duplicate sweat sodium levels of more than 70 mmol/l on at least 100 mg sweat. As part of their routine annual assessment they underwent, in random order on the same day, posteroanterior and lateral chest radiography at full inspiration taken under known conditions of magnification, and whole body plethysmography.

Total lung capacity was estimated by digitising the chest radiographs³⁻⁵ by the same observer (JM); this involves using a digitising tablet linked to a personal computer to measure the dimensions of the chest wall, heart, mediastinal, and subdiaphragmatic structures from posteroanterior and lateral chest radiographs. Allowance is made for magnification. The algorithms use shape assumptions to measure lung volume to a good degree of accuracy. Care is necessary to ensure that the patient has taken a full inspiration. Total lung capacity was also recorded using a Fenyves and Gut whole body plethysmograph in the routine lung function laboratory (technical staff) taking the mean of three measurements required to be within 5% of each other. Hyperinflation was scored as none, minimal, moderate, marked, severe¹ and graded 0-4² independently by a radiologist (DH) and a paediatric respiratory physician (AB) using the standard criteria.¹² All measurements and assessments were made without knowledge of the other results.

Results

The results of plethysmographic and radiographic measurement of total lung capacity were very similar (radiographic lung volume (ml) = 1.23 × plethysmographic lung volume - 224, $r^2 = 0.88$; table). Radiographic estimation of lung volume was a mean of 563 ml greater than plethysmographic lung volume. Hyperinflation was defined as total lung capacity above the normal range for our laboratory - that is, >115% predicted. Mild, medium, marked, and severe hyperinflation were arbitrarily defined as 115-125%, 125-135%, 135-145%, and >145% of predicted total lung capacity. Although the two observers

Department of
Paediatric Respiratory
Medicine
J Marchant
A Bush

Department of
Radiology
D M Hansell

Royal Brompton
National Heart and
Lung Hospitals,
Sydney Street, London
SW3 6NP, UK

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Individual patient data

Patient no.	TLC pred (ml)	TLC chest radiography	TLC plethysmography	Observer 1	Observer 2
		Actual (% pred)	Actual (% pred)		
1	4526	4690 (104)	4390 (97)	0	1
2	6065	7951 (131)	6450 (108)	2	2
3	5052	5980 (119)	4850 (96)	1	2
4	6196	7249 (117)	6630 (107)	1	0
5	1792	2628 (147)	2150 (120)	2	2
6	2860	3560 (124)	3260 (114)	3	3
7	4329	4746 (110)	3420 (79)	3	3
8	4614	4663 (101)	3830 (83)	1	3
9	2186	2684 (122)	2230 (102)	1	2
10	2402	1727 (72)	2330 (97)	1	2
11	1816	2813 (155)	2670 (147)	4	4
12	3176	3113 (98)	3350 (105)	1	3
13	2485	3587 (144)	3280 (132)	2	3
14	2896	3954 (137)	3620 (125)	3	3
15	3804	5423 (123)	3500 (92)	2	3
16	1958	2508 (128)	2350 (120)	0	2
17	3765	3994 (106)	3690 (98)	2	3
18	3151	3547 (113)	2710 (86)	2	3
19	3418	4486 (132)	3350 (98)	2	3
20	2558	4358 (171)	3300 (129)	4	4
21	5056	5935 (118)	4500 (89)	1	1
22	4058	4470 (110)	4180 (103)	3	3
23	1805	2350 (130)	2040 (113)	3	2
24	3186	3231 (101)	3250 (102)	1	2
25	1906	2200 (115)	2440 (128)	0	2

TLC = total lung capacity; observer = clinical score (0-4).

Figure 1 Comparison between the mean of the two clinical scores and total lung capacity (TLC) measured from the digitiser. Although the extremes are correctly graded, there is poor agreement between clinical score and plethysmographic measurement.

agreed to within a single grade in 22 of 25 children (table), overall concordance was poor ($\kappa=0.19$). Both observers assigned the patients with mild and severe hyperinflation to the correct grades defined by plethysmographic and radiographic measurements. No patient with severe hyperinflation was incorrectly graded. In all other groups, however, neither observer scored with more than 50% diagnostic accuracy. The mean score of both observers was no better than either alone (fig 1). The results were the same whether the observers were compared with plethysmographic or radiographic measurements of total lung capacity.

Discussion

Radiographic and plethysmographic estimates of total lung capacity showed reasonable agreement in these patients with cystic fibrosis, similar to that found in other studies.³⁻⁵ The overestimate by radiography has been attributed to lung tissue volume, which is not measured by plethysmography. The main conclusion of this study, however, is that subjective assessment of hyperinflation from chest radiographs is unreliable, despite the good agreement between plethysmographic and radiographic values. Occasional radiographic estimates of total lung capacity are very different from plethysmography. We therefore checked our conclusions by repeating the comparisons with plethysmographic instead of radiological estimations. Comparison of radiographic estimates with plethysmography did not alter the conclusions. The allocation of degrees of hyperinflation to particular bands of lung volume was to some extent arbitrary; 115% predicted has been taken as the upper limit of normal,⁶ but different ways of dividing mild, moderate, and severe above this level did not significantly improve the results. Scoring systems for cystic fibrosis include an assessment of hyperinflation; from this study it is clear that, whatever is being assessed on the radiographs, it bears little relation to objective measurements of lung volume (fig 2). This does not necessarily invalidate the scoring systems, but does suggest that a better term should be sought and that clinicians should not delude themselves that they can objectively assess lung volumes from chest radiographs, at least in children with cystic fibrosis.

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Figure 2 (A) Posteroanterior and (B) lateral chest radiographs scored as severely hyperinflated by both observers. Total lung capacity measured radiographically and plethysmographically was normal.

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Effects of inhaled lignocaine and adrenaline on capsaicin-induced cough in humans

L Hansson, B Midgren, J-A Karlsson

Abstract

Background – The hypothesis that adrenaline can augment and/or prolong the antitussive effect of nebulised lignocaine was examined.

Methods – The effect of inhaled lignocaine alone (20 mg) and in combination with adrenaline (400 µg) was studied on capsaicin-induced cough in 10 healthy subjects.

Results – Cough was significantly reduced between five and 25 minutes by lignocaine. Adrenaline alone had no inhibitory effect and it neither augmented nor prolonged the antitussive effect of lignocaine. The subjective anaesthesia by lignocaine was short lasting (less than 15 minutes) and not altered by adrenaline, suggesting different sensory mechanisms for anaesthesia and cough suppression. Plasma concentrations of lignocaine were low (<30 ng/ml), not altered by adrenaline, and did not correlate with the local anaesthetic or the antitussive effect.

Conclusions – Lignocaine acts locally in the oropharynx and airways and adrenaline does not alter the effect or absorption of nebulised lignocaine on the human respiratory mucosa.

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levels of lignocaine were measured to determine the degree of systemic absorption.

Methods

Ten non-smoking healthy subjects (five women) of mean age 27 (range 18–33 years) took part in the study. They gave their written informed consent and the study was approved by the University Hospital medical ethics committee, Lund. Capsaicin (Sigma) was dissolved in ethanol and diluted with 0.9% NaCl to 0.4 µmol/l, 2 µmol/l, 10 µmol/l, and 50 µmol/l.

Capsaicin was inhaled by tidal breathing from a nebuliser (BIRD Asmastic, output 0.5 ml/min and mass median diameter 3 µm) filled with 2 ml of solution.¹ A microphone and tape recorder were used to register the sounds of breathing and cough. The number of coughs was counted from the tape recordings. Blood samples were drawn from an arm vein into heparinised tubes. All blood samples were centrifuged and plasma was then separated and stored at –25°C until analysed by gas chromatography (Astra Alab, Södertälje, Sweden).

The antitussive effects of nebulised lignocaine (20 mg), adrenaline (400 µg), lignocaine in combination with adrenaline (20 mg + 400 µg), and vehicle (saline) were studied on four separate study days. Treatments were administered in a randomised, double blind manner. Increasing concentrations of capsaicin (0.4–50 µmol/l) were inhaled until a response of at least 10 coughs per minute was reached, and the concentration was then repeated and the mean response of the two challenges used as a baseline value. On each study day the chosen concentration was repeated. Capsaicin challenges were repeated five, 15, 25, 45, and 60 minutes after treatment with the study drug. Subjects were asked to subjectively assess the level of oropharyngeal anaesthesia according to a five point scale immediately before each capsaicin challenge. Blood samples for determination of plasma levels of lignocaine were drawn before and 10, 20, 30, 45, and 60 minutes after lignocaine inhalation. Kruskal–Wallis test was used at each

Department of Lung Medicine, University Hospital, S-221 85 Lund, Sweden
L Hansson
B Midgren

Discovery Biology, Rhône-Poulenc Rorer Ltd, London, UK
J A Karlsson

Reprint requests to:
Dr L Hansson.

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Lignocaine inhibits experimental cough in humans in a dose-dependent manner¹ and inhalation of a large dose of lignocaine has been reported to be an effective treatment in patients with severe persistent cough.^{2,3} Unfortunately, lignocaine has a short duration of action. Phenylephrine reduces blood flow in the tracheobronchial mucosa and adrenaline increases vascular resistance in the nasal mucosa⁴ in the dog. We therefore determined whether α -adrenoreceptor stimulation would potentiate the effects of inhaled lignocaine. The aim was to examine the antitussive effect and the oropharyngeal numbness of inhaled lignocaine and adrenaline on capsaicin-induced cough in healthy human subjects. In addition, plasma