airway obstruction and not against its causes will not be beneficial.

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Drs Skoner and Fireman comment:
In response to Dr Amirav: (1) Persistence of post treatment rises of the measured prosta-
glandin F2α metabolite were not unexpected, as this metabolite is stable in plasma and may be detectable for hours after the release of prostaglandin F2α, which has a relatively short half life. (2) All of the infants in group II (infants after treatment) had residual signs of airway obstruction. (3) Pulmonary function testing would certainly be superior to clinical parameters in assessing the degree of airway obstruction, but this methodology was unavailable and may be unreliable in infants. (4) Inflammatory mediators are released either as a primary event causative of a disease process or as a secondary event, which may not be related to pathophysiology. In either case, the relationship of the mediator to the pathophysiological process is very complex, as inflammation is likely an orchestrated response, with release of multiple mediators simultaneously and synergism or antagonism between various mediators in provoking pathophysiologies. Additionally, the desired human pathophysi-
ological response may be inaccessible to direct measurement. As such, absolute proof of a causal relationship between a mediator and a pathophysiological process is frequently lacking and rests on a body of indirect rather than direct evidence.

Three criteria for implicating a mediator in disease pathophysiology are generally accepted: (1) recovery at or near the site of active disease; (2) reproduction by exposure to the mediator; and (3) attenuation or preven-
tion by pretreatment with a mediator antag-
osta. The results of our study indicate that a metabolite of prostaglandin F2α was detect-
able in plasma during acute airway obstruction and satisfy criteria (1). Other investigators have documented that inhaled prostaglandin F2α can cause bronchoconstriction (criteria 2).

1 This evidence, although suggestive, is insufficient to prove a causal relationship. In situations such as this, clinical trials with specific antagonists (criteria 3) can provide confirmation that a given mediator participates in a pathophysiological process. For this reason, we believe that therapeutic trials using specific anti-inflammatory agents for the treat-
ment of airway obstruction are warranted.


Diagnostic accuracy of pH monitoring in gastro-oesophageal reflux

Str—Da Dalt and colleagues believe their data show that a pH study has 100% sensi-
tivity and 94% specificity for the diagnosis of gastro-oesophageal reflux.1 In particular, in their discussion they make comment of the fact that they did not find any false negatives and conclude that a negative result must be thought of as being sufficient to rule out gastro-oesophageal reflux.

In order to comment about the sensitivity or specificity, a comparison must be made with a gold standard. In the study by Da Dalt and colleagues, the indication for 24 hour hoes-
egophageal pH study was a clinical suspicion of gastro-oesophageal reflux. If the pH study was negative, the children were followed up clini-
cally for a period ranging from eight months to two years. On the basis of the observations made over this period of time, the authors decided none of these children had gastro-
oesophageal reflux. I would be very sceptical that clinical follow up was sufficient to use as a gold standard and would be reluctant to accept that clinical suspicion would be used as an indication for a test on the one hand, and then subsequently as the confirmation that the test was accurate.

By contrast, the gold standard used for con-
firmation that abnormal tests were indeed abnormal was the combination of a barium swallow or oesophagoscopy, or both. Although no indication is made of what constituted a positive barium swallow examination or an oesophagoscopy, it would be more confi-
 dent with the specificity of the test. It would be useful to know whether the oesophago-
sopy included a biopsy.

Finally, I question the conclusion that pH monitoring should be used as the first line of investigation in the diagnosis of gastro-
oesophageal reflux. A case may be able to be made for this suggestion in children admitted to hospital, but the extrapolation that the diagnosis of gastro-oesophageal reflux in a child who is an outpatient should involve his or her admission to hospital, is premature.

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Da Dalt L, Zazzoleni S, Montini G, Donzelli F, Zachelli F. Diagnostic accuracy of pH moni-

Str—Drs Skoner and Fireman comment:
We are grateful to Dr Henry for the comments on our paper and for the opportunity to clarify some aspects of our study.

We agree with the necessity of a gold standard but we would like to point out that neither barium swallow nor oesophagoscopy has got the characteristics to be a gold standard in the diagnosis of gastro-
oesophageal reflux.

For that reason and for ethical consider-
ations we decided not to perform these exami-
nations on children with normal pH study and we thought that a clinical follow up would be sufficient. In fact in most of these children symptoms occurred only because they were due, we think, to a physiological gastro-oesophageal reflux, which is not detected as abnormal by pH study and for which no treatment is necessary. Only in a few children, in whom symptoms were persistent, did we reach a different final diagnosis such as colicad disease, anorexia, etc.

We also like to point out that in patients with an abnormal pH study we performed a barium swallow and/or oesopha-
goscopy especially in order to rule out structural anomalies; however, a radiological gastro-
oesophageal reflux was diagnosed, that at least two episodes of spontaneous reflux were present in five minutes (according to Meyers et al). A biopsy was performed in all our oesophagoscopies and signs of oesophagitis were always present.

Finally we agree with Dr Henry that not all children who vomit need a pH study or other investigations, but we think that when a major clinical problem is present the important gastro-oesophageal reflux is suspected, admis-
sion is to be considered. An alternative could be to perform an ambulatory 24 hour pH test.

We have recently been attempted even in children.


A catabolic state in dexamethasone treatment of bronchopulmonary dysplasia

Str—Dexamethasone treatment is being increasingly used in babies with broncho-
pulmonary dysplasia because of its benefit.

We have reviewed the effect of treatment in the first nine infants we have treated. All were infants with bronchopulmonary dysplasia who had been ventilated from birth for respiratory distress syndrome. Gestation ranged from 25 to 30 weeks (mean 27.5) and birth weight ranged from 740 to 1510 g (mean 1050). Four babies had required surgical ligation of a patent ductus arteriosus. Dexamethasone was commenced between days 32 and 67, and the starting dose was varied between 0.46 and 2.13 mg/kg/ day (mean 1-07). The infants’ ventilatory requirement had been either static or deteriorating over the week before treatment. We found the treatment effective in the short term managing successfully to exhaust all the infants within a week and seven of nine by the third day of treatment.

There has been concern over the incidence of side effects including hypertension, septi-
caemia, necrotising enterocolitis, hyper-
glycaemia, and the possibility of longer term adrenal suppression. In our patients infection, hyperglycaemia, and hypercaemia did not cause any real problem. Three of our nine patients, however, required prolonged courses in excess of five weeks to maintain the effect and appeared to exchange dexamethasone for ventilator dependence, this is of particular concern with regard to possible adrenal sup-
pression.

We would highlight the fact that a pronounced catabolic state developed on commencing treatment, to our knowledge this has not been commented on in previous reports. Mean weight gain fell to -57 g/week from +190 g/week. There was no weight in the first week of treatment and all had previously been gaining weight. Blood urea concentrations averaged over the first week on treatment were 5.43 times those of

1 Da Dalt L, Zazzoleni S, Montini G, Donzelli F, Zachelli F. Diagnostic accuracy of pH moni-

Diagnostic accuracy of pH monitoring in gastro-oesophageal reflux.

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