

Thorax

The Journal of the British Thoracic Society
A Registered Charity

President: J E STARK

Executive Editors: J R Britton and A J Knox

Associate Editors: N C Barnes, M W Elliott, J A Fleetham, N M Foley, B G Higgins, N Høiby,
S A Lewis, M R Miller, M F Muers, D F Rogers, D P Strachan, W S Walker,
J O Warner, R J D Winter

Technical Editor: Elizabeth Stockman

Editorial Assistant: Hilary Hughes

Advisory Board

S H Abman USA

D M Geddes UK

F D Martinez USA

P D Sly Australia

J M Anto Spain

P Goldstraw UK

D M Mitchell UK

M J Tobin USA

P J Barnes UK

C Haslett UK

A J Peacock UK

M Woodhead UK

E D Bateman South Africa

P J Helms UK

R M Rudd UK

Editor, *British Medical Journal*

P S Burge UK

G J Laurent UK

N A Saunders Australia

Notice to contributors

SUBMISSION AND PRESENTATION The original typescript and three copies of all papers should be sent to the Executive Editors, *Thorax* Editorial Office, Division of Respiratory Medicine, City Hospital, Nottingham NG5 1PB, UK. Editorial and historical articles are normally commissioned but the Editors may accept uncommissioned articles of this type. Manuscripts must be accompanied by a declaration, signed by all authors, that the paper is not under consideration by any other journal at the same time and that it has not been accepted for publication elsewhere. The typescript should bear the name and address of the author who will deal with editorial correspondence, and also a fax number if possible. Authors may be asked to supply copies of similar material they have published previously. If requested, authors shall produce the data upon which the manuscript is based for examination by the Editors. Papers are accepted on the understanding that they may undergo editorial revision. In the event of rejection one copy of the text may be retained for future reference. **Authors are asked to supply the name and address of a possible referee for their work.**

Papers must be typed in double spacing with wide margins for correction and on one side of the paper only. They should include a structured abstract on a separate sheet (see below). Full papers should follow the basic structure of abstract, introduction, methods, results, discussion, references, and tables and figures as appropriate. They should not normally exceed 3000 words or include more than 30 references; priority will be given to papers that are concise. In each issue of the journal we will publish a small number of Rapid Communications, intended for reports of work of major importance in any areas of research, which will undergo an accelerated reviewing and publication process. Rapid Communications must not exceed 2000 words, 15 references, and two figures or tables. Short reports of experimental work, new methods, or a preliminary report can be accepted as two page papers and should comprise no more than 1300 words including a structured abstract, one table or illustration, and a maximum of 10 references. Case reports should not exceed 850 words with one table or illustration, a short unstructured abstract, and 10 references.

ABSTRACT Abstracts, which should be of no more than 250 words, should state clearly why the study was done, how it was carried out (including number and brief details of subjects, drug doses, and experimental design), results, and main conclusions. They should be structured to go under the headings "Background", "Methods", "Results", and "Conclusions".

KEYWORDS Authors should include on the manuscript up to three key words or phrases suitable for use in an index.

STATISTICAL METHODS The Editors recommend that authors refer to Altman DG, Gore SM, Gardner MJ, Pocock SJ. Statistical guidelines for contributors to medical journals. *BMJ* 1983;286:1489-93. Authors should name any statistical methods used and give details of randomisation procedures. 95% confidence intervals should be quoted for main results given as means or medians. The power of the study to detect a significant difference should be given when appropriate and may be requested by referees. Standard deviation (SD) and standard error (SE) should be given in parenthesis (not preceded by \pm) and identified by SD or SE at the first mention.

SI UNITS The units in which measurements were made should be cited. If they are not SI units the factors for conversion to SI units should be given as a footnote. This is the responsibility of the author.

ILLUSTRATIONS Line drawings, graphs, and diagrams should be prepared to professional standards and submitted as originals or as unmounted glossy photographic prints. Particular care is needed with photomicrographs, where detail is easily lost—it is often more informative to show a small area at a high magnification than a large area. Scale bars should be used to indicate magnification. The size of the symbols and lettering (upper and lower case rather than all capitals) and thickness of lines should take account of the likely reduction of the figure—usually to a width of 65 mm. Four copies of each illustration should be submitted. Each should bear a label on the back marked in pencil with the names of the authors and the number of the figure, and the top should be indicated. Legends should be typed on a separate sheet. Authors must pay for colour illustrations.

REFERENCES Responsibility for the accuracy and completeness of references rests entirely with the authors. References will not be checked in detail by the Editors but papers in which errors are detected are unlikely to be accepted. Reference to work published in abstract form is allowed only in exceptional circumstances—for example, to acknowledge priority or indebtedness for ideas. References should be numbered in the order in which they are first mentioned and identified in text, tables, and legends to figures by arabic numerals above the line. References cited only (or first) in tables or legends should be numbered according to where the particular table or figure is first mentioned in the text. The list of references should be typed in double spacing and in numerical order on separate sheets. The information should include reference number, authors' names and initials (all authors unless more than six, in which case the first six names are followed by *et al*), title of article, and in the case of journal articles name of journal (abbreviated according to the style of *Index Medicus*), year of publication, volume, and first and last page numbers. The order and the punctuation are important and should conform to the following examples:

- 1 Anderson HR. Chronic lung disease in the Papua New Guinea Highlands. *Thorax* 1979;34:647-53.
- 2 Green AB, Brown CD. *Textbook of pulmonary disease*. 2nd ed. London: Silver Books, 1982:49.
- 3 Grey EF. Cystic fibrosis. In: Green AB, Brown CD, eds. *Textbook of pulmonary disease*. London: Silver Books, 1982:349-62.

REVIEWING PROCESS Papers submitted to *Thorax* will be assessed by the Executive Editors and those considered unsuitable for publication will be returned directly to the authors. All other papers will be peer reviewed by an associate editor and at least one other reviewer. Rapid Communications will be reviewed and returned to the authors within 4 weeks, and published 2 or 3 months after acceptance.

CORRESPONDENCE The Editors welcome letters related to articles published in *Thorax*. These should not exceed 300 words or contain more than three references, which should be listed at the end of the letter. Letters should be typed in double spacing with wide margins and must be signed by all authors.

REPRINTS Reprints are available at cost if they are ordered when the proof is returned.

NOTICE TO ADVERTISERS Applications for advertisement space and for rates should be addressed to the Advertisement Manager, *Thorax*, BMJ Publishing Group, BMA House, Tavistock Square, London WC1H 9RJ.

NOTICE TO SUBSCRIBERS *Thorax* is published monthly. The annual subscription rate is £203.00 (\$319.00) worldwide. Orders should be sent to the Subscription Manager, *Thorax*, BMJ Publishing Group, BMA House, Tavistock Square, London WC1H 9RJ. Orders may also be placed with any leading subscription agent or bookseller. Subscribers may pay for their subscriptions by Access, Visa, or American Express by quoting on their order the credit or charge card preferred together with the appropriate personal account number and the expiry date of the card. For the convenience of readers in the USA subscription orders with or without payment may also be sent to *British Medical Journal*, PO Box 408, Franklin, MA 02038, USA. All inquiries, however, must be addressed to the publisher in London. All inquiries about air mail rates and single copies already published should also be addressed to the publisher in London. Second class postage paid at Rahway New Jersey. Postmaster: send address changes to *Thorax* c/o Mercury Airfreight International Ltd Inc, 2323 Randolph Avenue, Avenel, NJ 07001, USA.

COPYRIGHT © 1996 THORAX This publication is copyright under the Berne Convention and the International Copyright Convention. All rights reserved. Apart from any relaxations permitted under national copyright laws, no part of this publication may be reproduced, stored in a retrieval system, or transmitted in any form or by any means without the prior permission of the copyright owners. Permission is not, however, required for copying abstracts of papers or of articles on condition that a full reference to the source is shown. Multiple copying of the contents of the publication without permission is always illegal.

ISSN 0040-6376

CLINICAL SYMPOSIUM

CURRENT ASPECTS OF LUNG CANCER An International Perspective

University of Liverpool
Friday, 28th June 1996

An international panel of speakers will address topical issues related to lung cancer with particular reference to early diagnosis and intervention.

Information from:
R. J. Donnelly FRCSE
The Cardiothoracic Centre
Thomas Drive
Liverpool L14 3PE.
Tel: 0151-228-1616
Fax: 0151-230-0328



Imperial College
OF SCIENCE, TECHNOLOGY AND MEDICINE
University of London

National Heart & Lung Institute and Royal Brompton Hospital Advances In Respiratory Medicine

22 - 26 April, 1996

Course Organiser: Dr Margaret Hodson

This course will update physicians in important clinical developments in respiratory medicine. It is intended for respiratory physicians, general physicians and senior registrars (or equivalent) with special responsibilities for respiratory medicine. Contributors will come from many European countries.

Topics Will Include

Pharmacology and management of Asthma, critical care, sleep related disorders, lung transplantation and updates on the management of fibrosing lung disease, cystic fibrosis and 'COPD' together with case presentations.

Speakers

Experts from the UK and guest speakers from mainland Europe

Course Fee: £450.00

CME accreditation sought

Enquiries and payments should be made to: Imperial College School of Medicine at the Conference Centre, National Heart & Lung Institute, Dovehouse Street, London SW3 6LY. Telephone: 0171 351-8172 (24 hr answering service), Facsimile: 0171 376-3442

The National Heart and Lung Institute operates a NO SMOKING policy.
The College is striving towards Equal Opportunities.

At the leading edge of research, innovation and learning

Beclazone Easi-Breathe Inhaler

Beclomethasone Dipropionate BP

(Please refer to full data sheet before prescribing)

■ **Presentation** Metered-Dose Aerosol supplied in a Breath-Operated Inhaler containing 200 doses. **Beclazone 50 Easi-Breathe, Beclazone 100 Easi-Breathe and Beclazone 250 Easi-Breathe Inhalers** deliver 50, 100 and 250 microgram Beclomethasone Dipropionate BP per actuation of the valve.

■ **Uses** Provides automatic actuation of inhaler with inspiration. For the management of bronchial asthma especially in patients inadequately controlled by bronchodilators and sodium cromoglycate.

■ **Dosage and Administration** Use regularly. **Adults, Beclazone 50 and 100 Easi-Breathe Inhalers**; 100 microgram three or four times daily. **Beclazone 250 Easi-Breathe Inhaler**; 500 microgram twice a day or 250 microgram four times a day. **Elderly**, no dose adjustment necessary, including patients with renal or hepatic impairment. **Children, Beclazone 50 and 100 Easi-Breathe Inhalers**; 50 to 100 microgram two to four times daily. **Beclazone 250 Easi-Breathe Inhaler** is not indicated for use in children.

■ **Contra-indications** Hypersensitivity to the ingredients.

■ **Precautions** Patients should be instructed in the correct use of inhalers. May induce systemic cortico-steroid effects (with reduction in plasma cortisol levels) and adrenal suppression (above 2000 microgram daily) - monitor adrenal function and provide systemic steroids in appropriate cases of stress. Caution in patients with history of, or active pulmonary tuberculosis. Avoid sudden cessation of treatment.

■ **Pregnancy/Lactation** Use inhalers only if the potential benefit outweighs the risk.

■ **Side Effects** Paradoxical bronchospasm - discontinue use immediately and seek medical advice. Candidiasis, hoarseness or throat irritation - relieve by rinsing throat with water.

■ **Product Licence Numbers and Basic NHS Cost**

Beclazone 50 Easi-Breathe Inhaler PL 0530/0451 (£4.34)

Beclazone 100 Easi-Breathe Inhaler PL 0530/0452 (£8.24)

Beclazone 250 Easi-Breathe Inhaler PL 0530/0453 (£18.02)

■ **Legal Category** POM.

■ **Further Information** is available on request from: Baker Norton Gemini House, Flex Meadow, Harlow, Essex CM19 5TJ

■ **Date of Issue** July 1995

Salamol Easi-Breathe Inhaler

Salbutamol BP

(Please refer to full data sheet before prescribing)

■ **Presentation** Metered-Dose Aerosol supplied in a Breath-Operated Inhaler containing 200 doses.

Salamol Easi-Breathe Inhaler metered-dose aerosol delivering 100 microgram of Salbutamol BP per actuation.

■ **Uses** Provides automatic actuation of inhaler with inspiration. For the treatment and prophylaxis of bronchial asthma.

■ **Dosage and Administration** For optimum results use as required. Each administration has a bronchodilator effect which should last about 4 hours. **Adults** (i) Acute bronchospasm and intermittent episodes of asthma, including relief of symptoms such as wheezing, breathlessness and tightness of the chest - one or two inhalations as a single dose. (ii) Chronic maintenance or prophylactic therapy - two inhalations three or four times daily. (iii) To prevent exercise induced bronchospasm - two inhalations should be taken before exertion. **Children** (i) Acute bronchospasm and episodic asthma, including relief of symptoms such as wheezing, breathlessness and tightness of the chest, or before exercise - one inhalation. (ii) Routine maintenance or prophylactic therapy - one inhalation three or four times daily. The doses in children may be increased to two inhalations if necessary. Children should be supervised.

■ **Contra-indications** In spite of the fact that salbutamol has been used intravenously and orally in the management of uncomplicated premature labour, **Salamol Easi-Breathe Inhaler** should not be used for managing premature labour or for threatened abortion. **Salamol Easi-Breathe Inhaler** is contra-indicated in patients with a history of hypersensitivity to any of its components.

■ **Warnings** Potentially serious hypokalaemia may result from beta2-agonist therapy. It is recommended that serum potassium levels are monitored when the hypokalaemic effect may be potentiated by concomitant drugs or hypoxia. Propranolol and other non-cardioselective beta-adrenoceptor blocking agents antagonise the effect of salbutamol.

■ **Precautions** Patients with hyperthyroidism, who are hypersusceptible or who are suffering from diabetes mellitus, serious cardiovascular disorders or hypertension should use salbutamol containing products with caution. Asthmatic patients whose condition deteriorates despite salbutamol therapy or where a previously effective dose fails to give relief for at least three hours should seek medical advice. Alternative or additional therapy including corticosteroids should be instituted promptly although adverse metabolic effects of high doses of salbutamol may be exacerbated by concomitant administration of high doses of corticosteroids. Patients should not increase the dosage or frequency of administration without seeking medical advice.

■ **Side Effects** Potentially serious hypokalaemia may result from beta2-agonist therapy (see Warnings). Salbutamol may cause fine tremor of skeletal muscle (particularly the hands), palpitations and muscle cramps. Slight tachycardia, tenseness, headaches and peripheral vasodilatation have also been reported but these are less usually associated with the inhalation dosage form. Hypersensitivity reactions have been reported very rarely. Reports of hyperactivity in children are rare with beta2-agonists.

■ **Pregnancy/Lactation** **Salamol Easi-Breathe Inhaler** should be used during pregnancy or lactation only after careful consideration by the medical practitioner that the expected benefit outweighs the risk. **Salamol Easi-Breathe Inhaler** should not be used for managing premature labour or for threatened abortion (see Contra-indications).

■ **Product Licence Number and Basic NHS Cost**

Salamol Easi-Breathe Inhaler PL 0530/0399 (£6.30)

■ **Legal Category** POM.

■ **Further Information** is available on request from: Baker Norton Gemini House, Flex Meadow, Harlow, Essex CM19 5TJ

■ **Date of Issue** July 1995

Beclazone, Beclazone Easi-Breathe and Salamol, Salamol Easi-Breathe and Baker Norton are trademarks of Norton Healthcare Limited.

References:

- 1 Lindgren S, Bake B, Larsson S. Eur J Respir Dis 1987;70:93-98
- 2 Crompton G.K. Eur J Respir Dis 1982;63(Suppl. 119):101-104
- 3 Goodman D.E. et al. Am J Respir Crit Care Med 1994;150:1256-1261
- 4 Efficah M.J. et al. The Pharmaceutical Journal, 1994; 253:467-468
- 5 MIMS, July 1995
- 6 Data on file, Baker Norton
- 7 Based on IMS Data

**BAKER
NORTON**

Quality medicines at sensible prices

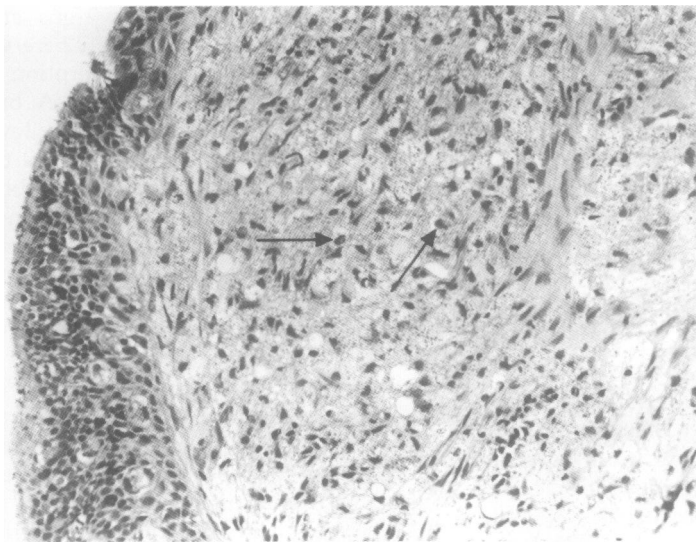


Figure 2 Histological features of an endobronchial lesion with PAS positive diastase resistant foamy macrophages (arrows). Magnification $\times 100$ reduced to 80% in origination.

and there is some doubt as to whether the granulomata were due to direct infection or resulted from a delayed hypersensitivity reaction. Differential diagnosis should include Wegener's granulomatosis, although this diagnosis was rejected in our patient in view of the negative anticytoplasmic antibody test and his benign clinical course.

We feel that the concept that dissemination of infection to organs outside the gastrointestinal tract occurs late in the evolution of the disease should be challenged, as diarrhoea and biochemical evidence of malabsorption may be absent while involvement elsewhere may be significant, as our case suggests. Our patient

did not consent to having biopsy samples taken of the small bowel until several months of treatment had been completed, in spite of being persistently requested to do so. The absence of diagnostic changes at this stage is not surprising and is consistent with his excellent clinical and radiological response to antibiotics. We suggest that this case represents a unique presentation of Whipple's disease and that pulmonary manifestations of this disorder should be extended to include pulmonary nodules and endobronchial lesions which may be mistaken for features of malignancy. Pulmonary involvement may be the presenting feature of the condition and offers the opportunity to diagnose the disease at a stage where response to treatment is likely to be both more rapid and complete than if it is delayed until manifestations of gastrointestinal involvement appear.

- 1 Whipple GH. A hitherto undescribed disease characterized anatomically by deposits of fat and fatty acids in the intestinal and mesenteric lymphatic tissues. *Bull Johns Hopkins Hosp* 1907;18:382-91.
- 2 Russo FR. Whipple's disease. Review of the literature and report of two cases. *Arch Intern Med* 1952;8:600-14.
- 3 Clancy RL, Tomkins WAF, Muckle TJ, Richardson H, Rawls WE. Isolation and characterization of an aetiological agent in Whipple's disease. *BMJ* 1975;iii:568-70.
- 4 Kelly JJ, Weisiger BB. The arthritis of Whipple's disease. *Arthritis Rheum* 1963;6:615-32.
- 5 Cho C, Linscheer WG, Hirschhorn MA, Ashutosh K. Sarcoidlike granulomas as an early manifestation of Whipple's disease. *Gastroenterology* 1984;87:941-7.
- 6 Winberg CD, Rose ME, Rappaport H. Whipple's disease of the lung. *Am J Med* 1978;65:873-80.
- 7 Symmons DP, Shepherd A, Boardman P. Pulmonary manifestations of Whipple's disease. *J Med* 1985;56:497.
- 8 Enzinger FM, Helwig EB. Whipple's disease: a review of the literature and report of fifteen patients. *Virchow's Arch Pathol Anat* 1963;336:238-69.
- 9 Rodarte JR, Garrison CO, Holley KE, Fontana RS. Whipple's disease simulating sarcoidosis. A case with unique clinical and histologic features. *Arch Intern Med* 1972;129:479-82.
- 10 Otto HF, Siemssen S, Sill V. Zur Differentialdiagnose von Morbus Whipple und Sarkoidose. Klinische-biologische Untersuchungen. *Deutsch Med Wochenschr* 1972;97:1343-7.

Thorax 1996;51:344

BOOK REVIEW

A History of Breathing Physiology. Donald F Proctor. (Pp 416; \$150.00). New York: Marcel Dekker, 1995. 0 8247 9653 5.

I very much enjoyed reading this book. It is, like the proverbial football match, an affair of two halves. All but one of the first 10 chapters is written by the editor, and in these he traces up to the 17th century the story of the struggle to understand the nature and purpose of breathing. After the customary coverage of Babylonian, Egyptian and Greek physiology, there is a chapter on Galen which helps to explain how his ideas, although

wrong, became so authoritative. However, Dr Proctor's chief loves are clearly the English physiologists of the 17th century, particularly Mayow, and he devotes five chapters to them, providing a lot of detail on their lives and achievements, but unfortunately also with much repetition. He tackles the question of why they drifted into different fields of enquiry after coming so close to understanding the whole mystery, leaving it for Lavoisier, a century later, to provide the final essential missing piece in the jigsaw with the discovery of oxygen.

The second half of the book consists of eight chapters by various authors who bring the story up to date in their respective fields from pulmonary surfactant to regulation of breathing. These vary in quality; that by Permutt on the pulmonary circulation is outstanding. The book is lavishly illustrated and well produced, apart from the chapter on

da Vinci in which the 14 illustrations, their legends, and their text references have been shuffled into total non-correspondence which is disappointing in a book that is so exorbitantly priced. - SF

NOTICE

1st Congress of Surgery of Bosnia and Herzegovina

This Congress, which was originally planned for 8-11 October 1995, will now take place in Sarajevo on 12-15 May 1996. For further information please contact the Congress Committee. Telephone: 387 71 44 55 22. Fax: 387 71 47 19 76.