

BRITISH JOURNAL OF PHARMACOLOGY

The *British Journal of Pharmacology* welcomes contributions in all fields of experimental pharmacology including neuroscience, biochemical, cellular and molecular pharmacology. The Board of Editors represents a wide range of expertise and ensures that well-presented work is published as promptly as possible, consistent with maintaining the overall quality of the journal.

Edited for the British Pharmacological Society by

A.T. Birmingham
(Chairman)

R.W. Horton **W.A. Large**
(Secretaries)

Editorial Board

P.I. Aaronson *London*
J.A. Angus *Melbourne, Australia*
G.W. Bennett *Nottingham*
T.P. Blackburn *Harlow*
N.G. Bowery *London*
W.C. Bowman *Glasgow*
S.D. Brain *London*
K.D. Butler *Horsham*
M. Caulfield *London*
R. Chess-Williams *Sheffield*
T. Cocks *Melbourne, Australia*
S.J. Coker *Liverpool*
R.A. Coleman *Ware*
Helen M. Cox *London*
A.J. Cross *London*
V. Crunelli *Cardiff*
T.C. Cunnane *Oxford*
F. Cunningham *London*
A. Dickenson *London*
J.R. Docherty *Dublin*
A. Dray *London*
L. Edvinsson *Lund, Sweden*
G. Edwards *Manchester*
J.M. Edwardson *Cambridge*
R.M. Eglen *Palo Alto, USA*
P.C. Emson *Cambridge*
A.C. Foster *San Diego, USA*
J.R. Fozard *Basle, Switzerland*
Allison D. Fryer *Baltimore, USA*

J.P. Gallagher *Galveston, USA*
Sheila M. Gardiner *Nottingham*
C.J. Garland *Bristol*
A. Gibson *London*
M.A. Giembycz *London*
W.R. Giles *Calgary, Canada*
R.G. Goldie *Perth, Australia*
R.J. Griffiths *Connecticut, USA*
R.W. Gristwood *Cambridge*
Judith M. Hall *London*
D.W.P. Hay *Philadelphia, USA*
P.G. Hellewell *London*
P.E. Hicks *Edinburgh*
S.J. Hill *Nottingham*
K. Hillier *Southampton*
S.M.O. Hourani *Guildford*
J.C. Hunter *Palo Alto, USA*
E.J. Johns *Birmingham*
R.S.G. Jones *Oxford*
C.C. Jordan *Ware*
P.A.T. Kelly *Edinburgh*
D.A. Kendall *Nottingham*
C. Kennedy *Glasgow*
P. Leff *Loughborough*
A.T. McKnight *Cambridge*
C.A. Maggi *Florence, Italy*
Janice M. Marshall *Birmingham*
G. Martin *Beckenham*
W. Martin *Glasgow*

A. Mathie *London*
D.N. Middlemiss *Harlow*
P.K. Moore *London*
C.D. Nicholson *Oss, The Netherlands*
H. Osswald *Tübingen, Germany*
F.L. Pearce *London*
J.D. Pearson *London*
A.G. Renwick *Southampton*
P.J. Roberts *Bristol*
G.J. Sanger *Harlow*
W.C. Sessa *Connecticut, USA*
P. Sneddon *Glasgow*
K. Starke *Freiburg, Germany*
R.J. Summers *Melbourne, Australia*
P.V. Taberner *Bristol*
J. Tamargo *Madrid, Spain*
C. Thiemeermann *London*
M.D. Tricklebank *Basle, Switzerland*
T.J. Verbeuren *Suresnes, France*
R.R. Vollmer *Pittsburgh, USA*
K.J. Watling *Boston, USA*
A.H. Weston *Manchester*
J. Westwick *Bath*
Eileen Winslow *Riom, France*
E.H.F. Wong *Milan, Italy*
B. Woodward *Bath*

Corresponding Editors

P.R. Adams *Stony Brook, U.S.A.*
C. Bell *Dublin*
F.E. Bloom *La Jolla, U.S.A.*
A.L.A. Boura *Newcastle, Australia*
N.J. Dun *Toledo, U.S.A.*
R.F. Furchgott *New York, U.S.A.*
T. Godfraind *Brussels, Belgium*
S.Z. Langer *Paris, France*

R.J. Miller *Chicago, U.S.A.*
R.C. Murphy *Denver, U.S.A.*
E. Muscholl *Mainz, Germany*
R.A. North *Geneva, Switzerland*
M. Otsuka *Tokyo, Japan*
M.J. Rand *Melbourne, Australia*
S. Rosell *Södertälje, Sweden*
P. Seeman *Toronto, Canada*

L. Szekeres *Szeged, Hungary*
B. Uvnäs *Stockholm, Sweden*
P.A. Van Zwieten *Amsterdam, Netherlands*
V.M. Varagić *Belgrade, Yugoslavia*
G. Velo *Verona, Italy*
Wang Zhen Gang *Beijing, China*
M.B.H. Youdim *Haifa, Israel*

Submission of manuscripts: Manuscripts (two copies) should be sent to The Editorial Office, British Journal of Pharmacology, St. George's Hospital Medical School, Cranmer Terrace, London SW17 0RE.

Authors should consult the Instructions to Authors and the Nomenclature Guidelines for Authors in Vol. **114**, 245–255. These Instructions and Guidelines also appear with the journal Index for Volumes **111–113**, 1994. A checklist of the essential requirements is summarised in each issue of the journal, or as the last page of the issue.

Whilst every effort is made by the publishers and editorial committee to see that no inaccurate or misleading data, opinion or statement appears in this Journal, they and the *British Pharmacological Society* wish to make it clear that the data and opinions appearing in the articles and advertisements herein are the responsibility of the contributor or advertiser concerned. Accordingly, the *British Pharmacological Society*, the publishers and the editorial committee and their respective employees, officers and agents accept no liability whatsoever for the consequences of any such inaccurate or misleading data, opinion or statement.

The *British Journal of Pharmacology* is published by Stockton Press, a division of Macmillan Press Ltd. It is the official publication of the British Pharmacological Society.

Scope The *British Journal of Pharmacology* is published twice a month. It welcomes contribution in all fields of experimental pharmacology including neuroscience, biochemical, cellular and molecular pharmacology. The Board of Editors represents a wide range of expertise and ensures that well-presented work is published as promptly as possible, consistent with maintaining the overall quality of the journal.

This journal is covered by Current Contents, Excerpta Medica, BIOSIS, CABS, CINAHL and Index Medicus.

Editorial Manuscripts (plus two copies) and all editorial correspondence should be sent to: The Editorial Office, British Journal of Pharmacology, St George's Hospital Medical School, Cranmer Terrace, London SW17 0RE, UK. Tel: +44 (0)181 767 6765; Fax: +44 (0)181 767 5645.

Advertisements Enquiries concerning advertisements should be addressed to: Michael Rowley, Hasler House, High Street, Great Dunmow, Essex CM6 1AP, UK. Tel: +44 (0)1371 874613; Fax: +44 (0)1371 872273.

Publisher All business correspondence, supplement enquiries and reprint requests should be addressed to British Journal of Pharmacology, Stockton Press, Houndmills, Basingstoke, Hampshire RG21 2XS, UK. Tel: +44 (0)1256 29242; Fax: +44 (0)1256 810526. Publisher: Marija Vukovojac. Production Controller: Nicci Crawley.

Subscriptions – EU/Rest of World Subscription price per annum (3 volumes, 24 issues) £665, rest of world £880 (Airmail), £735 (Surface mail) or equivalent in any other currency. Orders must be accompanied by remittance. Cheques should be made payable to Macmillan Magazines and sent to: The Subscription Department Macmillan Press Ltd, Houndmills, Basingstoke, Hampshire RG21 2XS, UK. Where appropriate, subscribers may make payments into UK Post Office Giro Account No. 519 2455. Full details must accompany the payment. Subscribers from EC territories should add sales tax at the local rate.

Subscriptions – USA USA subscribers call toll free 1-800-221-2123 or send check/money order/credit card details to: Stockton Press, 49, West 24th Street, New York, NY 10010; Tel: 212 627 5757, Fax: 212 627 9256. USA annual subscription rates: \$1400 Airmail; \$1170 Surface (Institutional/Corporate); \$240 (Individual making personal payment).

British Journal of Pharmacology (ISSN 0007-1188) is published twice a month by Macmillan Press Ltd, c/o Mercury Airfreight International Ltd, 2323 Randolph Avenue, Avenel, NJ 07001, USA. Subscription price for institutions is \$1400 per annum (surface). 2nd class postage is paid at Rahway NJ. Postmaster: send address corrections to Macmillan Press Ltd, c/o Mercury Airfreight International Ltd, 2323 Randolph Avenue, Avenel NJ 07001.

Reprints of any article in this journal are available from Stockton Press, Houndmills, Basingstoke, Hampshire RG21 2XS, UK. Tel: +44 (0)1256 29242; Fax: +44 (0)1256 810526.

Copyright © 1995 Stockton Press
ISSN 0007-1188

All rights of reproduction are reserved in respect of all papers, articles, illustrations, etc., published in this journal in all countries of the world.

All material published in this journal is protected by copyright, which covers exclusive rights to reproduce and distribute the material. No material published in this journal may be reproduced or stored on microfilm or in electronic, optical or magnetic form without the written authorisation of the Publisher.

Authorisation to photocopy items for internal or personal use of specific clients, is granted by Stockton Press for libraries and other users registered with the Copyright Clearance Center (CCC) Transaction Reporting Service, provided that the base fee of \$12.00 per copy is paid directly to CCC, 21 Congress St., Salem, MA 01970, USA. 0007-1188/95 \$12.00 + \$0.00.

Apart from any fair dealing for the purposes of research or private study, or criticism or review, as permitted under the Copyright, Designs and Patent Act 1988, this publication may be reproduced, stored or transmitted, in any form or by any means, only with the prior permission in writing of the publishers, or in the case of reprographic reproduction, in accordance with the terms of licences issued by the Copyright Licensing Agency.



INSTRUCTIONS TO AUTHORS

With effect from 1 January 1996

The *British Journal of Pharmacology* welcomes contributions in all fields of experimental pharmacology, including neuroscience, biochemical, cellular and molecular pharmacology, for publication as full papers or as high priority Special Reports.

Papers should normally be based on new results obtained experimentally and should constitute a significant contribution to pharmacological knowledge. Papers which reassess pharmacological concepts based on earlier results will also be considered as will purely theoretical papers. Papers dealing only with descriptions of methods are acceptable if new principles are involved.

Contributions that have already been published, or accepted or are under consideration for publication, with essentially the same content will not be considered. This restriction does not apply to results published as abstracts of communications, letters to editors, or as contributions to symposia, provided that the submission adds significantly to the information available in the previously published contribution.

Papers are only accepted if accompanied by a Declaration which must be signed by all Authors. This Declaration concerns the originality of the submitted paper and assigns the copyright of all papers accepted for publication to Stockton Press Ltd. on behalf of the British Pharmacological Society. See pages 7 and 8 for details.

The Journal will not consider papers which describe experiments on animals which do not fall clearly within the current laws governing animal experimentation in the United Kingdom. Authors must make it clear that the procedures they use were as humane as possible and the doses (initial and subsequent) of anaesthetics and analgesics should be clearly stated; the method of assessing anaesthesia, particularly after the administration of skeletal muscle relaxants (neuromuscular blocking drugs), must be well defined. The Society has an Ethics Committee which can be consulted by authors through the Secretaries to the Editorial Board.

When investigations on normal human subjects are reported, evidence of approval by a local Ethics Committee must be given. Papers concerned with clinical trials or investigations of the effects of drugs on patients are not appropriate for this Journal.

Authors are strongly urged to keep their manuscripts as short as they reasonably can. An effective way is to reduce the Discussion and the number of figures to a minimum and to avoid repetition of information that has already been published. Authors should remember that a reader may be influenced by literary style and will appreciate simple but accurate prose.

It is important to note that failure to comply with 'Instructions to Authors' may lead to considerable editorial delays.

FULL PAPERS

Manuscripts must be typed on one side of A4 paper. Words at the end of lines should not be divided because they may become incorrectly hyphenated. Handwritten characters or symbols (e.g. Greek letters) should be spelled out in full in the margin. Papers in recent issues of the *British Journal of Pharmacology* should be consulted for the general layout of the paper and also for details. The following subsections are used:

1. Title page
2. Summary
3. Introduction
4. Methods

5. Results
6. Discussion and conclusions
7. Acknowledgements
8. List of references
9. Tables
10. Figures and captions

The type must not be smaller than 12 pitch or 10 point. Each section must be typed in **double spacing** with margins of not less than 2.5 cm all round and each page should be numbered. **The original and one copy of the typescript should be supplied.**

Title page

The title should normally contain **no more than 150 characters** and should not consist of a sentence (statement or conclusion) or be interrogative. A **short running title** containing not more than 50 characters and spaces **is also required**. The title page should include the names of authors and their appropriate addresses. It should be made clear which address relates to which author. Authors' present addresses differing from those at which the work was carried out should be given as footnotes on the title page and references at the appropriate place in the author list by superscript numbers. A footnote may also be used to indicate the author to whom correspondence should be sent. The use of footnotes for any other reason is not allowed. If the address to which proofs should be sent is not that of the first mentioned author, clear instructions should be given in a covering note and not on the title page. The title page should be paginated as page 1 of the paper.

Summary

The summary will be printed at the beginning of the paper. It should not exceed 5% of the length of the paper and should contain a brief account of the problem, the methods, results and the conclusions. It should be arranged in **numbered and concise paragraphs**. Up to ten **keywords** or phrases of two to three words (including names and terms used in the title) should be displayed at the end of the summary. Keywords will be used to compile the annual index. The quality of the index will thus be determined by the appropriateness of the keywords. These may be selected by reference to the most recent Index of the Journal. Avoid unhelpful or unqualified terms such as 'rat', 'drug' etc.

Introduction

The introduction should give a short and clear account of the background of the problem and the rationale of the investigation. Only previous work that has a direct bearing on the present problem should be cited.

Methods

The methods must be described in sufficient detail to allow the experiment to be interpreted and repeated by the reader. However, detailed repetition of methods which have been adequately described previously should be avoided and references given, although a brief outline is often helpful.

Drugs should be listed in a separate paragraph. Their

names should be 'approved names' as published previously in British Approved Names, 1990 (HMSO). If a drug has no 'approved name' its chemical name must be used and the rules set out in the current *Handbook for Chemical Society Authors* (London, Chemical Society) observed, or its structural formula given. Cumbersome chemical names should be suitably abbreviated for later reference in the paper.

The doses of drugs should be given as unit weight per body weight, e.g. mmol kg^{-1} or mg kg^{-1} ; concentrations should be given in terms of molarity, e.g. nM or μM .

Reference should be made to any statistical analyses that have been performed on the results in order, for example, to determine the significance of differences between results obtained under different conditions.

Results

The description of the experimental results should be succinct but, nevertheless, in sufficient detail to allow the experiments to be repeated by others. Typical single experiments may be presented with a clear statement that n number of similar experiments had similar results. Where appropriate, however, the mean results with confidence limits or with standard errors of the means and the number of observations should be given. Statistical tests of significance should be performed where appropriate. The results of such tests should be stated as the numerical value of the probability (P) that is calculated, with any necessary clarification (e.g. one-tail or two-tail test).

Every effort should be made to avoid unnecessary repetition of data in the text, tables and figures. Conclusions and theoretical considerations should not be elaborated in this section.

Discussion

The purpose of the discussion is to present a brief and pertinent interpretation of the results against the background of existing knowledge. Any assumptions on which conclusions are based must be stated clearly. A mere recapitulation of the results is not acceptable. A review-like treatment, which reduces the impact on the reader, should also be avoided. The main conclusion should be conveyed in a final paragraph.

Acknowledgements

Acknowledgements should be brief but should include reference to sources of support. Sources of drugs not widely available commercially should be acknowledged.

References

In the text, references to other work should take the form: (Bolton & Kitamura, 1983) or, 'Bolton & Kitamura (1983) showed that . . .'. If there are more than two authors, the first author's name should be given followed by *et al.* (Bülbring *et al.*, 1981).

References to 'unpublished observations' or 'personal communications' should be mentioned in the text only, and not included in the list of references. Papers which have been submitted and accepted for publication, should be included in the list of references with the names of the periodicals and 'in press'. A photocopy should normally be submitted with the manuscript. If this is not possible, authors should indicate whether the work cited is an abstract or a full paper. Papers in preparation or which have been submitted but not yet finally accepted for publication must not be included in the list of references.

The reference list at the end of the manuscript must be arranged alphabetically according to the surname of the first author. When the surnames of authors are identical, the alphabetical order of their initials takes precedence over the year of publication. The AUTHORS' names are followed by the year of publication in brackets. If more than one paper by the same authors in one year are cited, a, b, c, etc. are placed after the year of publication, both in the text and in the list of references. The title of the article is given in full, followed by the *abbreviated title of the periodical*, volume number and first and last page numbers. The abbreviations used for periodicals are those of the most recent edition of the International List of Periodical Title Word Abbreviations. References to articles in books should consist of names of authors, year of publication, title of article followed by the *title of the book*, the editors, volume numbers, if any, and page numbers, the place of publication and the names of the publishers. For example:

BOLTON, T.B. & KITAMURA, K. (1983). Evidence that ionic channels associated with the muscarinic receptor of smooth muscle may admit calcium. *Br. J. Pharmacol.*, **78**, 405–416.
BRADING, A.F. (1981). Ionic distribution and mechanisms of transmembrane ion movements in smooth muscle. In *Smooth Muscle: An Assessment of Current Knowledge*. ed. Bülbring, E., Brading, A.F., Jones, A.W. & Tomita, T. pp. 65–92. London: Edward Arnold.

Tables

Each table should be given on a separate page, paginated as part of the paper. Tables should be numbered consecutively with arabic numerals and the number should be followed by a brief descriptive caption, occupying not more than two lines, at the head of the table. The proportions of the text area should be borne in mind when designing the layout of tables. For the sake of clarity, tables should not have more than 120 characters to a line, with spaces between columns counted as four characters. The absolute maximum is 180 characters to a line. Each column should have a heading and the units of measurement should be given in parentheses in the heading. Except in special circumstances, tables should be self-explanatory; the necessary descriptions should be at the bottom of the table.































Figures

To avoid unnecessary Figures, particularly those requiring half-tone reproduction, only critical points of the text should be illustrated. If coloured Figures are desired, the Authors should discuss their requirements with the Secretaries, preferably before submission.

Please note that unsatisfactory Figures will be returned to the Author for revision. The Journal reserves the right to reject a manuscript if the Figures are unacceptable.

Submission Requirements

- The Authors' names and the Figure number must be indicated lightly *in pencil* on the back of each Figure; if necessary, use an adhesive label to avoid damage to the Figure.
- Each copy of the manuscript must be accompanied by one set of labelled Figures (i.e. complete with lettering and numbering, arrows, etc.). An original set and one high quality photocopy will suffice.
- Another original set of Figures identical in size but without any letters or numbers must also be supplied for the use of the Publisher. Arrows and event marks on experimental

Line width (axes)	Line width (graphs)	Symbol size	Figure will reduce to this percentage of the original size
		  	100 (No reduction)
		  	80
		  	70
		  	60
		  	50
		  	40

- records may be retained, provided they are larger than 3 mm wide. The Publisher will choose the correct style of typeface of an appropriate size to suit the final size of the Figure on the printed page.
- (d) No submitted Figure should exceed 210 × 297 mm (A4).
- (e) Each Figure must be accompanied by a legend; each legend should be typed on a *separate* sheet of paper and paginated as part of the manuscript. Legends should explain the Figures in sufficient detail that, whenever possible, they can be understood without reference to the text.

Illustrations

All illustrations will be scanned electronically for inclusion in the journal therefore care should be taken when preparing artwork for publication.

Any illustrations that have been prepared on a computer may be submitted on disk, provided the material is supplied as:

- 1 Macintosh TIFF (or EPS with bounding box directly around the graphic to allow for reduction without excessive loss of resolution).
- 2 IBM/PC TIFF (file must be in graphic format NOT document).

Minimum resolution for TIFF files to be 300 dpi for tone work and 800 dpi for line work. Preferred packages: Adobe illustrator, Aldus Freehand and Mac Photoshop.

The disk **must** be clearly marked with journal/author/manuscript number, file name and programme used and accompany the manuscript to which it refers, and **include** printed copies of the illustration for identification. All illustrations will be reduced to fit single column width wherever possible. Amendments to the illustration may be made to conform to journal style.

Any illustrations containing blots from gels, histochemical stains or pen traces, that have been prepared via a computer programme cannot be reproduced from a laser printout, as this creates a cross-hatched pattern. Such material must be submitted on disk as above, unless unscanned continuous tone originals are supplied. Amendments to the illustrations may be made to conform to the journal style (ie: labelling).

Line Figures

It is best to submit an original drawing (black ink on heavy white paper or faint blue graph paper) which has been

prepared to conform with the style and convention of the Journal, because redrawing is expensive. The original drawing should be lettered in pencil and should be larger (up to two times as large) than the intended size in the Journal.

It is important that the printed symbols and lines should retain their clarity. To achieve this the symbols and lines in original drawings should be sharply defined and of an even density and breadth. When graphs are generated by computer, lines must not show noticeable stepping. Heavier (broader) lines should be used for curves than for the axes of graphs. The table above illustrates line widths and symbol sizes to be used on a figure and the appropriate reductions in the final printed form.

Symbols should be chosen from the following set

○ ● □ ■ △ ▲ ▽ ▼ ◇ ◆ + ×

The preferred order to shading of histogram columns is: open (clear), closed (solid), cross-hatched, heavily stippled and other (if required).

The explanation of the symbols and column headings should be given in the Figure legend and not as a key in the Figure itself.

Line Figures should normally have only left and bottom axes; box-style Figures and those using 3-dimensions are not acceptable.

Where the Figure is a composite of more than one graph, experimental record, etc., particular care is needed to minimise the spaces between each part, without overcrowding the entire Figure.

Figure 1 illustrates a simple properly-drawn graph in its original form (a) and in its reduced form (b) as it would appear in the Journal (single column width).

Photographs and photomicrographs

These should be submitted, twice as large as their intended published size, as good quality prints of high contrast especially where traces and records are illustrated. The originals must not contain arrows, lettering or numbering; these must be accurately located on a duplicate print (or photocopy). When submitting half-tone illustrations for publication authors should remember that it is not possible to reproduce Figures to a finer quality than the original photographs/photomicrographs provided. Critical areas should be marked on a second copy or on an overlay, so that the Printer can choose the correct exposure. Maximum trim areas should be marked on a second copy of the photograph/

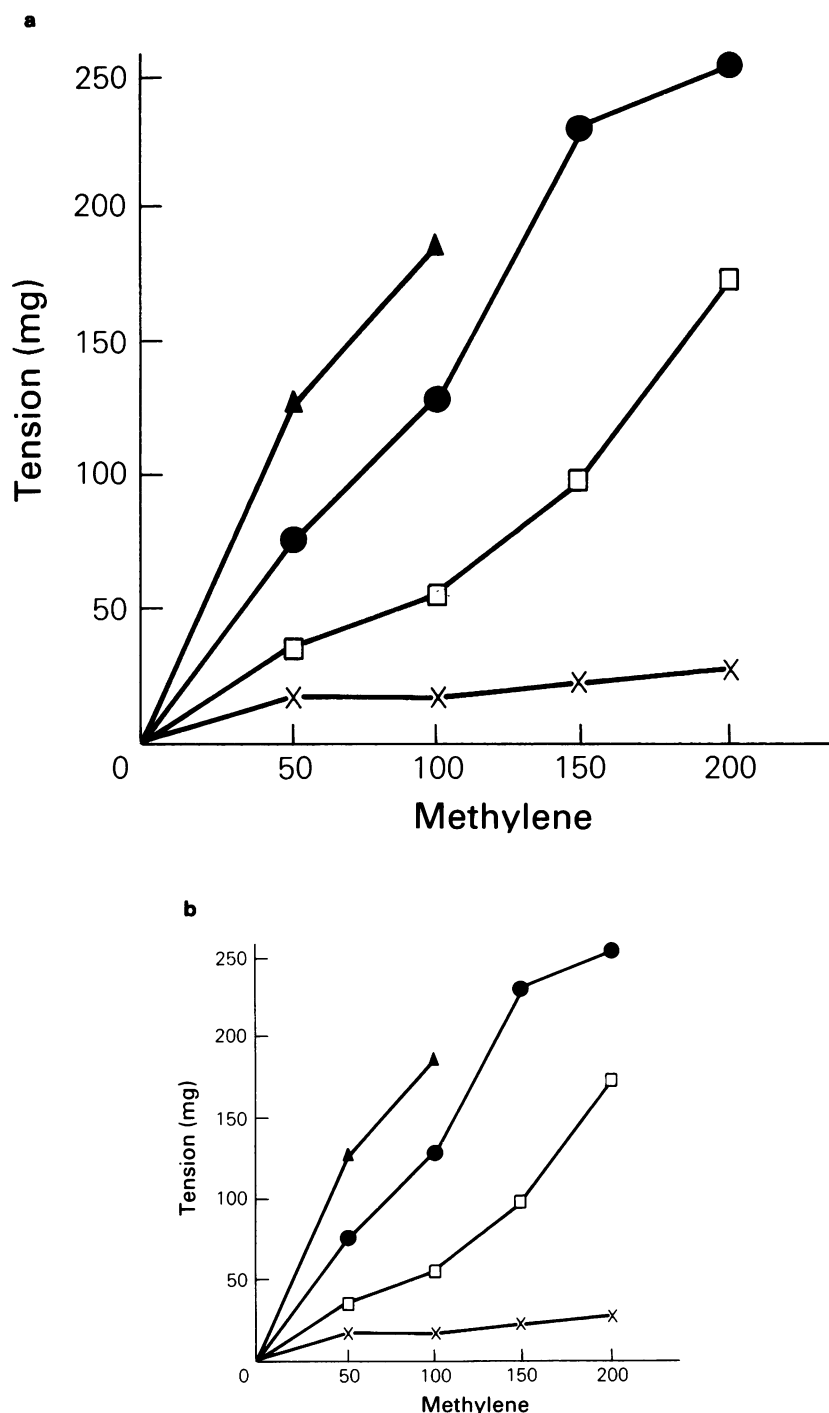


Figure 1 (a) Artwork as drawn. (b) Artwork reduced to 60 per cent of its original size for publication in the Journal to fit in single column width.

photomicrograph or on a tracing overlay, i.e. authors should show any parts of the photographs that could be excluded from the finished half-tone illustration. **A calibration bar must be provided on the photomicrograph** to ensure that, if the Printer reduces the plate, the scale is reduced in the correct proportion.

Submitting manuscripts on disk

Manuscripts, once accepted for publication, may be presented on disk as long as they meet the following criteria:

- The manuscript on disk **must** be the final version
- A double spaced hard-copy accompanies the disk and matches the disk version **exactly**.
- The disk is text only (i.e. no figures)
- The preferred program is WordPerfect 5.1 (DOS). (Acceptable alternatives are WordPerfect for Windows saved as 5.1 or ASCII and Microsoft Word for Mac)
- Diskettes should be 3.5" (90 mm).

Please ask the Publisher for full details if you wish to proceed with submitting your manuscript on disk.

We do not guarantee that the disk will be used or that it will expedite publication.

Proofs

One set of page proofs will be supplied, which should be corrected immediately and returned to the Publisher and a photocopy retained by the author. Corrections should be kept to a minimum.

SPECIAL REPORTS

The purpose of *Special Reports* is to provide rapid publication for new and important results which the Editorial Board considers are likely to be of special pharmacological significance. *Special Reports* will have publication priority over all other material and so authors are asked to consider carefully the status of their work before submission.

In order to speed publication there is normally no revision allowed beyond very minor typographical or grammatical corrections. If significant revision is required, the Board may either invite rapid re-submission or, more probably, propose that it be re-written as a Full Paper and be re-submitted for consideration. In order to reduce delays, proofs of *Special Reports* will be sent to authors but **essential corrections must reach the Publisher within 48 hours of receipt**. Authors should ensure that their submitted material conforms exactly to the following requirements.

Special Reports should normally occupy no more than two printed pages of the Journal; two illustrations (Figures or Tables, with legends) are permitted. As a guideline, with type face of 12 pitch and double-line spacing, a page of A4 paper could contain about 400 words. The absolute maximum length of the *Special Report* is 1700 words. For each Figure or Table, please deduct 200 words. The manuscript should comprise a Title page, a Summary consisting of a single short paragraph, followed by keywords (maximum of 10), Introduction, Methods, Results, Discussion and References (maximum of 10). In all other respects, the requirements are the same as for Full Papers.

STATEMENT AND COPYRIGHT AGREEMENT

The following Statement, Declaration and Copyright Agreement should be read carefully by Authors who should then send a copy of their Declaration with their manuscript following the example given in this section.

Statement

- 1. Submission of a manuscript will be taken to imply that the Authors have obtained permission to publish from (a) their employers or institution, if they have a contractual or moral obligation to do so, and (b) those whose unpublished work, including papers accepted for publication (i.e. in press), has been cited or those whom the Authors wish to acknowledge as having improved the content or presentation of the manuscript.
- 2. The Authors must declare that the manuscript contents are original and that they have not already been published or accepted for publication, either in whole or in part, in any form other than as an abstract or other preliminary publication in an unrefereed article. Furthermore, the Authors must verify that no part of the manuscript is under consideration for publication elsewhere and it will not be submitted elsewhere if accepted by the *British Journal of Pharmacology* and not before a decision has been reached by the Editorial Board.

Declaration

I/We assign to Stockton Press, on behalf of the British Pharmacological Society, the copyright of my/our manuscript, currently entitled

.....,
for publication in the *British Journal of Pharmacology*

Furthermore I/We have read, understood and accepted the terms and conditions as set out in Statement and Copyright, Instructions to Authors *Br. J. Pharmacol.* 1996, 117, 389–396.

Name

Signed

Name

Signed

Name

Signed

Date

Copyright

- 1. The Authors must agree that, when the above manuscript has been accepted for publication in this Journal, the worldwide copyright shall pass to the Stockton Press Ltd. on behalf of the British Pharmacological Society, on the understanding that the assignment of copyright will not affect subsisting Patent Rights arrangements pertaining to it. The Authors also accept that, when accepted, the contents will not be published subsequently in the same or similar form in any language without the consent of the Publisher or Editorial Board of the Journal.
This Agreement shall not compromise the Authors' rights to reproduce their own work (see 3 below). For its part, the British Pharmacological Society will protect the interest of Authors in the matter of copyright.
- 2. The Authors must declare that, where excerpts from copyrighted works have been included, the Authors have obtained written permission from the Copyright owners and have credited the sources in the manuscript. They must also warrant that the article contains no libellous or unlawful statements and does not infringe the rights of others.
- 3. The Authors will be entitled to publish any part of the paper in connection with any other work by them, provided adequate acknowledgement is given.
- 4. If it is appropriate, the Authors' employer may sign this Declaration. It is understood that proprietary rights, with the exceptions of Copyright and Patent Rights are reserved by the signee.
- 5. If an Author is a U.S. Government employee and the work was done in that capacity, the assignment applies only to the extent allowed by U.S. law. If an Author is an employee of the British Government, HMSO will grant a non-exclusive licence to publish the paper in the Journal, provided British Crown Copyright and user rights (including Patent Rights) are reserved.
- 6. If for good reason a co-author is unable to sign this Declaration, the other Author or co-authors may sign on his or her behalf, provided that this is clearly stated and on the understanding that they will make every effort to inform the person concerned of the terms of the agreement.

When submitting a manuscript for editorial consideration, Authors should confirm their acceptance of these terms by signing a Declaration to that effect. The recommended wording is given in the example. No paper will be accepted for publication without such a Declaration being signed by each Author (see paragraph 6 above). If the manuscript is not accepted for publication, the assignment will be null and void.

ABBREVIATIONS AND SYMBOLS

Physico-chemical quantities

The *British Journal of Pharmacology* uses the SI symbols for units. The following prefixed for multiples of units should be used:

<i>Multiplier</i>	<i>Prefix</i>	<i>Symbol</i>
10 ⁻¹	deci	d
10 ⁻²	centi	c
10 ⁻³	milli	m
10 ⁻⁶	micro	μ
10 ⁻⁹	nano	n
10 ⁻¹²	pico	p
10 ⁻¹⁵	femto	f
10 ⁻¹⁸	atto	a
<i>Multiplier</i>	<i>Prefix</i>	<i>Symbol</i>
10 ³	kilo	k
10 ⁶	mega	M
10 ⁹	giga	G
10 ¹²	tera	T

Thus, micron = μm; ångstrom = 0.1 nm. Mixed prefixes are not permissible, thus mμg should be ng. The symbols d (10⁻¹) and c (10⁻²) should be restricted to those occasions on which there is a strongly felt need for them (e.g. cm).

Use of the solidus

The solidus should be avoided as far as possible and the negative index substituted, e.g. mg kg⁻¹ rather than mg/kg; pmol mm⁻² min⁻¹ rather than pmol/mm²/min.

SYMBOLS

Symbols denoting physical quantities are usually printed as italic capitals (indicated by single underline in typescript). A dash over the symbol indicates a mean value; a dot over the symbol indicates a time derivative. Suffixes may be used to indicate 'where' and 'what'. They are printed as inferiors on the line. Multiple suffixes should be avoided if a simpler symbol adequately defined is unambiguous, but if necessary should be separated by commas e.g. P_{A, CO₂} denotes partial pressure of CO₂ alveolar air.

CHEMICAL AND BIOLOGICAL ABBREVIATIONS

Authors should also consult *Nomenclature Guidelines for Authors* contained in this issue of the Journal. The abbreviations listed may be used without definition *except* those for chemicals, drugs and enzymes which must be written in full at first mention in the title, summary and again in the text. At first mention they should be followed by the abbreviation in brackets. Subsequently, the abbreviation alone may be used.

The list of abbreviations for chemical, drug and enzyme names is clearly not comprehensive and includes only a few commonly used examples.

Use abbreviations sparingly as extensive use can make the text hard to follow.

Physico-chemical quantities

<i>Quantity</i>	<i>Preferred unit</i>	<i>Symbol</i>
Amount (of substance)	mole	mol
Capacitance	farad	F
Concentration	moles per litre	M or mol l ⁻¹
Current	ampere	A
Electrical conductance	siemens	S
Electromotive force	volt	V
Flow (blood or other liquid)	litres per second (or min)	l s ⁻¹ or l min ⁻¹
Flow (air or other gas)	litres per second (or min)	l s ⁻¹ or l min ⁻¹
Force	newton	N
Frequency of regular event	hertz	Hz
Length	mètre	m
Mass	gram	g
Power	watt	W
Pressure (or partial pressure)	pascal*	Pa
Radioactivity	becquerel or curie	Bq (60 d.p.m.) or Ci (3.7 × 10 ¹⁰ Bq)
Resistance (electrical)	ohm	Ω
Temperature	degree celsius	°C
Time	second (preferred)	s
	minute	min
	hour	h
Volume (blood or other liquid)	litre	l
Volume (air or other gas)	litre	l
Work	joule	J

* mm of mercury (mmHg) are allowed if conventional, and if mercury manometer is used for calibration.

Chemical and biological abbreviations

acetylcholine	ACh	dextro-(absolute configuration)	D-
acetylcholinesterase	AChE	dextro-(optical rotation)	(+)-
adenosine 3':5'-cyclic monophosphate	cyclic AMP	diameter	diam.
adenosine 5'-phosphate	AMP	diameter, inside	i.d.
adenosine triphosphatase	ATPase	diameter, outside	o.d.
γ -aminobutyric acid	GABA	diffusion coefficient	<i>D</i>
analysis of variants	F	3,4-dihydroxyphenylalanine	DOPA
adrenaline	Ad	3,4-dihydroxyphenylethylamine	dopamine
analytical standard of reagent purity	A.R.	direct current	d.c.
anhydrous	anhyd.	disintegration per minute	d.p.m.
approximate(ly)	approx.	dissociation constant	K_D
approximately equals	\approx	dissociation constant, negative logarithm of	pK
aqueous	aq.	distilled	dist.
arg-vasopressin	AVP	dry ice	solid CO ₂
		edition	edn
		editor(s)	ed.
boiling point	b.p.	effective concentration	EC ₅₀
bovine serum albumin	BSA	effective dose, median	ED ₅₀
		electrocardiogram	ECG
		electrocorticogram	ECoG
cardiovascular system	CVS	electroconvulsive therapy	ECT
catechol- <i>O</i> -methyl transferase	COMT	electroencephalogram	EEG
central nervous system	CNS	electromyogram	EMG
cerebrospinal fluid	CSF	electron spin resonance	e.s.r.
chi-squared (statistics)	χ^2	endothelial-derived relaxing factor	EDRF
clearance	<i>c</i>	epithelial-derived relaxing factor	EpDRF
coenzyme A	CoA	equilibrium constant	<i>K</i>
concentrated	conc.	equivalent (general use)	equiv.
correlation coefficient	<i>r</i>	erythrocyte	r.b.c.
cubic	cu.	erythrocyte sedimentation rate	ESR
		ethylenediaminetetracetic acid	EDTA
degree of freedom (statistics)	d.f.	excitatory postsynaptic potential	e.p.s.p.
deoxyribonucleic acid	DNA	experiment	expt
deoxyribonuclease	DNase	experimental	exptl
fatty acids, nonesterified	NEFA	page/pages	p./pp.
figure(s) (with reference number)	Figure(s)	para-	<i>p</i> -
figure (diagram)	figure	paragraph	para. or ¶
		parts per million	p.p.m.
gas-liquid chromatography	g.l.c.	per cent	%
glomerular filtration rate	GFR	platelet activating factor	PAF
		posterior	post.
haemoglobin	Hb	probability (significance level in a statistical test)	<i>P</i>
half-life	<i>t</i> ₁		
high-frequency	h.f.		
high performance liquid chromatography	h.p.l.c.	radioimmunoassay	RIA
human serum albumin	HSA	rectus (configuration by the sequence rule)	R
hydrogen-ion concentration	[H ⁺]	red blood corpuscle	RBC
hydrogen-ion activity, negative logarithm of (hydrogen-ion exponent)	pH	relative band speed to front (chromatography)	<i>R_F</i>
6-hydroxydopamine	6-OHDA	relative molecular mass	<i>M_r</i>
N-[2-Hydroxyethyl]piperazine-N'-[2-ethanesulphonic acid]	HEPES	relative retention time (gas chromatography)	<i>t_r</i>
5-hydroxyindoleacetic acid	5-HIAA	renal plasma flow	RPF
5-hydroxytryptamine	5-HT	resistance (respiratory)	<i>R</i>
		respiratory conductance	Sgaw
		revolutions per minute	r.p.m.
immunoglobulins	IgA, IgD, IgE, IgG, IgM	ribonucleic acid	RNA
	<i>K_i</i>		
inhibitor constant	IC ₅₀	section	§
inhibitory concentration	i.p.s.p.	sedimentation coefficient (ultracentrifugation)	<i>s</i>
inhibitory postsynaptic potential	insol.	sinister (configuration by the sequence rule)	S
insoluble	iu		
international unit	i.a.	soluble	sol.
intra-arterial	ICF	solution	soln.
intracellular fluid	i.d.	Spearman rank coefficient	<i>r_s</i>
intradermal	i.m.	standard deviation	s.d.
intramuscular	i.p.	(of observed sample)	
intraperitoneal			

intracerebroventricular	i.c.v.	standard error (of estimate mean value)	s.e.mean
intravenous	i.v.	standard error (of sampling)	s.e.
isotope (atomic mass)	¹³¹ I	standard temperature and pressure	STP
e.g. iodine-131		subcutaneous	s.c.
isotopically substituted compounds e.g.	[¹⁴ C]-ethanol	sum (statistical):	
		of hypothetical population	Σ
laevo-(absolute configuration)	L-	of observed sample	S or Σ
laevo-(optical rotation)	(-)-		
lethal dose, median	LD ₅₀	temperature	temp.
leukotriene	LT	thin layer chromatography	t.l.c.
logarithm to base e	log _e or ln	time, clock – 24 h clock used	t
logarithm to base 10	log ₁₀	e.g. 18 h 30 min	
		time constant	τ
maximum	max.	2-amino-2-hydroxymethyl-	Tris
mean arterial pressure	MAP	propan-1,3-diol	
mean value of (statistics)	\bar{x}		
melting point	m.p.	ultraviolet	u.v.
meta	m-	unit	u
Michaelis constant	K _M		
minimum	min.		
mobility (electrophoresis)	m		
monoamine oxidase	MAO	vacuum	vac.
		valency	e.g. Fe ²⁺ ;
			Fe(II)
noradrenaline	NA		protoporphyrin
nuclear magnetic resonance	n.m.r.		
number	no. or No.		
number of observations	n	volume by volume	v/v
(statistics)			
		wavelength	λ
ortho	o-	weight	wt.
packed cell volume	PCV	weight by volume	w/v



NOMENCLATURE GUIDELINES FOR AUTHORS

With effect from 1 January 1996

The Nomenclature Working Party (NWP) of the Editorial Board of the *British Journal of Pharmacology* has consulted many acknowledged experts in an effort to clarify and standardize receptor and other nomenclature systems for use by Editors until a complete set of recommendations from the International Union of Pharmacology Committee on Receptor Nomenclature and Drug Classification (NC-IUPHAR) is published.

NWP is unanimous in its view that, with rare exceptions, the Journal should use spellings, names and abbreviations that have been chosen by international bodies convened for the purpose.

For receptor nomenclature, with few exceptions, the Journal generally follows the guidelines laid down in the current *Trends in Pharmacological Sciences* Receptor and Ion Channel Nomenclature Supplement and the reports of NC-IUPHAR published in *Pharmacological Reviews*.

1 Definition of receptors and subtypes

Receptors and their subtypes are defined in terms of the relative potencies of agonists and selectivities of antagonists in functional studies, by the binding of such ligands, and structural information, where available.

2 Format of receptor names

It was agreed that, until NC-IUPHAR provides full recommendations:

- (a) Editors will permit with reluctance new nomenclature systems in papers accepted for publication if, and only if, there are compelling reasons to introduce a new terminology (or modify an accepted one). The criteria upon which the new receptor type or subtype are defined must be given, together with adequate explanations of the relationship between the previous nomenclature (fully referenced) and the proposed one.

N.B. The new nomenclature should not appear in the Title, Short Title or Keywords, unless qualified by the adjective putative (e.g. . . . mediated by the putative imidazoline I₂ receptor).

- (b) Only well-established and universally accepted subtype names (e.g. muscarinic and nicotinic acetylcholine receptors; α - and β -adrenoceptors) will be acceptable without any reference to the originator of these terms. In cases of controversy concerning further subdivision of the subtype, full referencing must be given.
- (c) When receptors are expressed from DNA/RNA that has been introduced into cells and these receptors display a similar pharmacological profile to the native receptors, they should be denoted by use of lower case, e.g. m1 for expressed receptor and M₁ for native receptor. The stoichiometry of the expressed receptor should be indicated, where appropriate, e.g. for an immature muscle nicotinic acetylcholine receptor, it might be (α)₂ β 1 γ δ .
- (d) Receptor subtypes should be designated by means of a subscript numeral or capital letter. Some double subscripts (i.e. numeral plus letter) are acceptable.
- (e) Greek letters and Roman numerals should be avoided in any new nomenclature. The name should not include the letter "R" or "r" as an abbreviation for receptor.

- (f) Mammalian systems are the basis of receptor classifications. Evolutionary changes may be so great that receptors in non-mammalian species are difficult to classify within this nomenclature. Therefore non-mammalian species should be clearly indicated, e.g. torpedo nicotinic receptor, chick β -adrenoceptor, locust GABA receptor.

3 Types of receptor

The NWP accepts that there are additional receptors to those described below which can be considered to be well characterised. In many cases, however, their existence has been confirmed only in cloning studies and it is as yet unclear how they relate to similar subdivisions proposed on the grounds of differences in agonist and antagonist potencies in various tissues.

- (a) *Acetylcholine receptors* The two principal subfamilies are muscarinic and nicotinic acetylcholine receptors.
Muscarinic acetylcholine receptors The principal subtypes are M₁, M₂, M₃, M₄ and M₅.
Nicotinic acetylcholine receptors The principal subgroups are muscle and neuronal nicotinic acetylcholine receptors.
- (b) *Adenosine receptors* Known also as P₁ purinoceptors (see Purinoceptors, 3v). (See Fredholm, B.B., *et al.*, (1994) *Pharmacol. Rev.*, **46**, 145–156.)
- (c) *Adrenoceptors* The principal subtypes are α ₁-, α ₂-, β ₁-, β ₂- and β ₃-adrenoceptors. Additional subtypes must be fully referenced. (See Bylund, D.B., *et al.*, (1994) *Pharmacol. Rev.*, **46**, 121–136 and Hieble, J.P., *et al.*, (1995) *Pharmacol. Rev.*, **47**, 267–270.)
- (d) *Angiotensin receptors* The principal subtypes are AT₁ and AT₂.
- (e) *Bombesin receptors* Proposed subtypes such as BB₁, BB₂ may be used but must be fully referenced.
- (f) *Bradykinin receptors* The principal subtypes are B₁ and B₂ receptors.
- (g) *Calcitonin gene-related peptide (CGRP) receptors* Proposed CGRP receptor subtypes must be fully referenced.
- (h) *Cannabinoid receptors* The principal subtypes are CB₁ and CB₂.
- (i) *Chemokine receptors* The principal subgroups are CC and CXC receptors. Subtypes of these must be fully referenced.
- (j) *Cholecystokinin (CCK) receptors* The principal subtypes are CCK_A and CCK_B receptors.
- (k) *Dopamine receptors* The principal subtypes are D₁, D₂, D₃, D₄ and D₅.
- (l) *Endothelin receptors* The principle subtypes are ET_A and ET_B receptors. (See Masaki, T., *et al.*, (1994) *Pharmacol. Rev.*, **46**, 137–142.)
- (m) *Excitatory amino acid receptors* Three ionotropic subtypes are recognised and named: (1) NMDA receptors; (2) AMPA receptors, and (3) kainate receptors. A second class is the metabotropic (mGlu) receptor family. Further subtypes must be fully referenced.

- (n) *γ-Aminobutyric acid (GABA) receptors* The principal subtypes are GABA_A and GABA_B receptors. Modulatory sites on the GABA_A receptor should be referenced.
- (o) *Histamine receptors* The principal subtypes are H₁, H₂ and H₃ receptors.
- (p) *5-Hydroxytryptamine (5-HT) receptors* The principal subtypes are 5-HT₁, 5-HT₂, 5-HT₃ and 5-HT₄. Further subdivisions require full referencing. (See Hoyer, D., *et al.*, (1994) *Pharmacol. Rev.*, **46**, 157–203).
- (q) *Leukotriene (LT) receptors* Receptors should be designated according to the leukotriene that selectively or preferentially binds to them. All leukotriene receptor subtypes should be fully referenced.
- (r) *Neuropeptide Y (NPY) receptors* Proposed subtypes should be fully referenced.
- (s) *Opioid receptors* The principal subtypes are μ-, δ- and κ-opioid receptors. Other proposed subtypes should be fully referenced.
- (t) *Oxytocin receptors* (see Vasopressin and oxytocin receptors).
- (u) *Prostanoid receptors* The principal types are DP, EP, FP, IP and TP receptors. When first mentioned, the style prostanoid (XP) receptor should be used, thereafter XP receptor (where X denotes the type). Proposed subtypes should be referred to as XP_n, (e.g. EP₁, EP₂) and referenced. (See Coleman, R.A., *et al.*, (1994) *Pharmacol. Rev.*, **46**, 205–229).
- (v) *Purinoceptors* The principal subtypes are P₁ and P₂ receptors. Subdivision of P₁ into A₁, A₂ and A₃ subtypes and of P₂ into P_{2X} and P_{2Y} are permitted. Other subtypes should be fully referenced. (See Fredholm, B.B., *et al.*, (1994) *Pharmacol. Rev.*, **46**, 143–156).
- (w) *Somatostatin (SST) receptors* Proposed subtypes should be fully referenced.
- (x) *Tachykinin receptors* The term tachykinin is preferred to neurokinin. The principle subtypes are NK₁, NK₂ and NK₃ receptors.
- (y) *Vasoactive intestinal peptide (VIP) receptors* Proposed subtypes should be fully referenced.
- (z) *Vasopressin and oxytocin receptors* The principle subtypes are V_{1A}, V_{1B}, V₂ and OT receptors.

4 Naming of ion channels

The four main ion channels are named according to the abbreviation of the ion normally carrying the current (i.e. K⁺ channels, Na⁺ channels, Ca²⁺ channels and Cl⁻ channels). Further subtypes must be fully referenced. For Ca²⁺ channels, see Spedding, M. & Paoletti, P. (1992) *Pharmacol. Rev.*, **44**, 363–376.

5 Naming of nerve fibres

Many nerve fibres are now known to release more than one transmitter, and future work may show that this is in fact the general rule. In that case, the concept of the same transmitter being released either at different developmental stages or under various experimental conditions would no longer hold, and single adjectives that imply this (e.g. cholinergic, noradrenergic) would become inappropriate when applied to nerve fibres, as distinct from transmitter functions. For the present, those nerve fibres that are known to function by releasing more than one identified transmitter may be described accordingly; for example, noradrenergic-purinergetic, cholinergic-peptidergic (in alphabetical order, the order implying no priority of

function). N.B. The suffix 'ergic' should continue to be applied only to nerve fibres and to the transmission event, in accordance with Dale's intentions. For example, 'cholinergic' indicates that the nerve fibre, or the transmission, functions under particular conditions through the release of a choline-like substance. The suffix should not be used loosely to mean 'pertaining to'. Hence, for example, the expression 'cholinergic receptor' (rather than acetylcholine receptor) is an inappropriate use of the term. Transmission events involving nitric oxide may be referred to as *nitroergic*. However, *nitroergic* may be used to describe axons only when there is sufficient evidence that nitric oxide is released from them as a neurotransmitter.

- (a) *Catecholamine releasing nerve fibres* The adjective to be applied to nerve fibres that release dopamine as a transmitter is dopaminergic (not DAergic).

Nerve fibres that are known to function by releasing noradrenaline are to be described as noradrenergic. The term adrenergic should now be reserved for nerve fibres known to release adrenaline. Where the identity of the catecholamine is uncertain, catecholaminergic should be used.

- (b) *Some other adjectives describing nerve fibre function* NANC is an acceptable abbreviation of non-adrenergic, non-cholinergic for peripheral efferent nerve fibres when the identity of the transmitter(s) is unknown other than the fact that neither (nor)-adrenaline nor acetylcholine is involved. It should be defined when introduced. NANCergic, e-NANC (or NANC-e) and i-NANC (or NANC-i) are not acceptable terms.

Glutamatergic, not glutaminergic, should be used to describe nerve fibres releasing glutamate. In referring to peptide-releasing nerve fibres (e.g. those that may release substance P or vasoactive intestinal peptide) the nomenclature to be used is peptidergic (X), e.g. peptidergic (SP), peptidergic (VIP), not SPergic, VIPergic.

The terms 5-hydroxytryptamine (5-HT) and 5-hydroxytryptaminergic (i.e. nerves releasing 5-hydroxytryptamine) are preferred to those of serotonin and serotonergic. The term 5-HTergic is not acceptable.

Likewise, the terms purinergetic (ATP) and purinergetic (adenosine) are preferred.

Terms used to describe agonist and antagonist action

The following terms can be used without full definition. Where appropriate, other terms may be used but must be accompanied by a full definition.

Terms used to describe affinity and potency

- (a) *EC₅₀* The concentration of an agonist that produces 50% of the maximal response for that agonist *in vitro*. The agonist may be stimulatory or inhibitory. When EC₅₀ values are determined in the presence of other agonists or antagonists the concentration of the latter should be stated. Related terms, e.g. EC₂₅, are acceptable if accompanied by a full definition.
- (b) *IC₅₀* This term may be used in the following ways: (i) The concentration of antagonist that reduces the response to a sub-maximal concentration of agonist by 50%; the concentration of agonist should be stated. (ii) The concentration of competing agonist or antagonist that inhibits the binding of a radioligand by 50%; the concentration of radioligand should be stated.

- (c) ED_{50} This term may be used in the following ways: (i) The dose of an agonist or antagonist that produces 50% of the maximal possible effect of that agonist or antagonist *in vitro*. (ii) The dose of drug that produces the effect under investigation in 50% of the population.
- (d) K The dissociation equilibrium constant ($M l^{-1}$) for ligand-receptor interactions. The reciprocal is called the affinity constant or association equilibrium constant. When necessary for clarity, subscripts (letters or numerals, or a combination of both) may be added but these must be clearly explained when first used.
- (e) n_H The Hill coefficient.
- (f) pA_2 The negative logarithm to base 10 of the concentration of an antagonist that makes it necessary to double the concentration of agonist needed to elicit a given submaximal response. Note that the definition is empirical and does not pre-suppose the mechanism of antagonism. The pA_2 value can be determined from a Schild plot with unconstrained slopes, but only provides an estimate of the pK_B if the antagonism has been shown to meet all of the criteria of competition.
- (g) pD_2 The negative logarithm to base 10 of the EC_{50} .
- (h) pIC_{50} The negative logarithm to base 10 of the IC_{50} .
- (i) pK The negative logarithm to base 10 of K (with or without subscripts as appropriate: see 6(d)).

Terms used to describe the mode of antagonism

- (a) *Competitive antagonism* In competitive antagonism, the binding of agonist and antagonist is mutually exclusive. This may be because the agonist and antagonist compete for the same binding site or combine with adjacent sites that overlap. A third possibility is that different sites are involved but they influence the receptor macromolecule in such a way that agonist and antagonist molecules cannot be bound at the same time.
- (b) *Irreversible competitive antagonism* Used to describe antagonists that bind irreversibly.
- (c) *Non-competitive antagonism* Agonist and antagonist can be bound simultaneously; antagonist binding reduces or prevents the action of the agonist.
- (d) *Irreversible non-competitive antagonism* Used to describe non-competitive antagonists that bind irreversibly.

For a more detailed account of the terms used to describe agonist and antagonist action see Jenkinson, D.H., *et al.*, (1995) *Pharmacol. Rev.*, **47**, 225–266.

(Enzyme Nomenclature 1992, Academic Press, San Diego and London) must be quoted when first mentioned in text. Subsequently the accepted trivial name is used. Trivial names may be used in the title.

8 Other nomenclature requirements

- (a) *Racemates* Authors must state unambiguously in the Methods section of papers which isomers were used, e.g. (+)- or (–)-propranolol, and must bring to the attention of the reader the composite character of drugs that are mixtures of stereoisomers. Furthermore, the implications of the composite nature of such drugs studied for the interpretation of the data measured and the conclusions drawn must be made explicit. Note that the terms d- or l- for dextro- and laevo-rotatory are now obsolete, and the prefixes (+)- or (–)- respectively should be used. Capital D and L refer to the absolute configurations and of course remain acceptable when appropriate.
- (b) *Purines* This term should not be used as a synonym for purine nucleotides or nucleosides.
- (c) *Eicosanoids* The system of nomenclature to be used for eicosanoids is that published in *Methods in Enzymology* (1990) **187**, 1–9. This scheme incorporates recent changes in the style of abbreviation of hydroperoxy-, epoxy- and oxo-unsaturated fatty acids, e.g. 12(S)-hydroperoxyeicosatetraenoic acid which was formerly abbreviated as 12(S)-HPETE now becomes 12(S)-HpETE. In manuscripts, the first use of the full chemical name of any eicosanoid should indicate double bond geometry when this is known.
- (d) *Cell lines* Cell type, species and source should be defined.
- (e) *Molecular biology* Abbreviations pertaining to molecular biological techniques need to be defined or presented in such a way that they can be recognised by the non-specialist, e.g. the oligonucleotide sequence, TAGC.
- (f) *Tension* Tension is force and should be calibrated in newtons ($1 \text{ newton} = 1 \text{ kg ms}^{-1}$) or in kg weight, g weight, or mg weight etc. It should not be calibrated in units of mass (e.g. kg). (See Miller D.J. (1988) *Trends Pharmacol. Sci.*, **9**, 124–5).
- (g) *Ions* When referring to ions, the charge should be indicated, e.g. Na^+ , Ca^{2+} , $2Na^+/Ca^{2+}$ exchange, etc.
- (h) *Inhibitors of nitric oxide synthase* The most commonly used and currently accepted abbreviations for N^G -nitro-L-arginine and N^G -nitro-L-arginine methyl ester are L-NOARG and L-NAME respectively.

7 Enzymes

The International Union of Biochemistry and Molecular Biology Enzyme Commission (EC) number and full name