

# BRITISH JOURNAL OF PHARMACOLOGY

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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.

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

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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.**

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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.

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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.

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

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

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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).

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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.

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| Line width<br>(axes)                                                              | Line width<br>(graphs)                                                            | Symbol size                                                                       | Figure will reduce<br>to this percentage<br>of the original size |
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|  |  |  | 80                                                               |
|  |  |  | 70                                                               |
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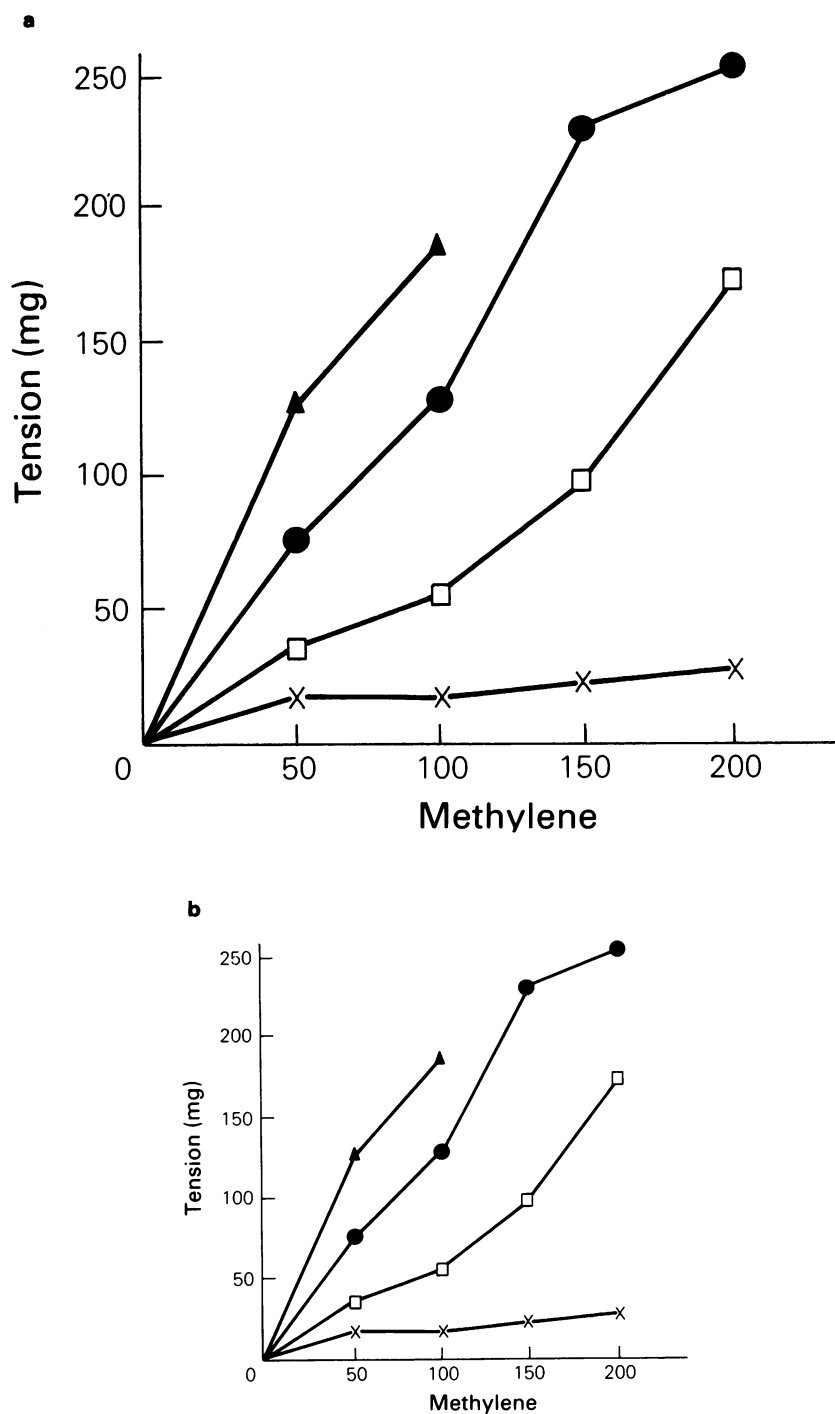
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**Figure 1** (a) Artwork as drawn. (b) Artwork reduced to 60 per cent of its original size for publication in the Journal to fit in single column width.

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## 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:

| Multiplier | Prefix | Symbol |
|------------|--------|--------|
| $10^{-1}$  | deci   | d      |
| $10^{-2}$  | centi  | c      |
| $10^{-3}$  | milli  | m      |
| $10^{-6}$  | micro  | $\mu$  |
| $10^{-9}$  | nano   | n      |
| $10^{-12}$ | pico   | p      |
| $10^{-15}$ | femto  | f      |
| $10^{-18}$ | atto   | a      |
| Multiplier | Prefix | Symbol |
| $10^3$     | kilo   | k      |
| $10^6$     | mega   | M      |
| $10^9$     | giga   | G      |
| $10^{12}$  | tera   | T      |

Thus, micron =  $\mu\text{m}$ ; ångström = 0.1 nm. Mixed prefixes are not permissible, thus  $\text{m}\mu\text{g}$  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).

### Physico-chemical quantities

| Quantity                       | Preferred unit             | Symbol                                          |
|--------------------------------|----------------------------|-------------------------------------------------|
| Amount (of substance)          | mole                       | mol                                             |
| Capacitance                    | farad                      | F                                               |
| Concentration                  | moles per litre            | M or $\text{mol l}^{-1}$                        |
| Current                        | ampere                     | A                                               |
| Electrical conductance         | siemens                    | S                                               |
| Electromotive force            | volt                       | V                                               |
| Flow (blood or other liquid)   | litres per second (or min) | $1 \text{ s}^{-1}$ or $1 \text{ min}^{-1}$      |
| Flow (air or other gas)        | litres per second (or min) | $1 \text{ s}^{-1}$ or $1 \text{ min}^{-1}$      |
| Force                          | newton                     | N                                               |
| Frequency of regular event     | hertz                      | Hz                                              |
| Length                         | metre                      | m                                               |
| Mass                           | gram                       | g                                               |
| Power                          | watt                       | W                                               |
| Pressure (or partial pressure) | pascal*                    | Pa                                              |
| Radioactivity                  | becquerel or curie         | Bq (60 d.p.m.) or Ci ( $3.7 \times 10^{10}$ Bq) |
| Resistance (electrical)        | ohm                        | $\Omega$                                        |
| Temperature                    | degree celsius             | $^{\circ}\text{C}$                              |
| 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.

### Use of the solidus

The solidus should be avoided as far as possible and the negative index substituted, e.g.  $\text{mg kg}^{-1}$  rather than  $\text{mg/kg}$ ;  $\text{pmol mm}^{-2} \text{ min}^{-1}$  rather than  $\text{pmol/mm}^2/\text{min}$ .

## 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.  $P_{A, \text{CO}_2}$  denotes partial pressure of  $\text{CO}_2$  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.



## 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.                  |
| $\gamma$ -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 <sub>2</sub> |
| boiling point                                                        | b.p.                    | edition                                                | edn                   |
| bovine serum albumin                                                 | BSA                     | editor(s)                                              | ed.                   |
| cardiovascular system                                                | CVS                     | effective concentration                                | EC <sub>50</sub>      |
| catechol- <i>O</i> -methyl transferase                               | COMT                    | effective dose, median                                 | ED <sub>50</sub>      |
| central nervous system                                               | CNS                     | electrocardiogram                                      | ECG                   |
| cerebrospinal fluid                                                  | CSF                     | electrocorticogram                                     | ECog                  |
| chi-squared (statistics)                                             | $\chi^2$                | electroconvulsive therapy                              | ECT                   |
| clearance                                                            | <i>c</i>                | electroencephalogram                                   | EEG                   |
| coenzyme A                                                           | CoA                     | electromyogram                                         | EMG                   |
| concentrated                                                         | conc.                   | electron spin resonance                                | e.s.r.                |
| correlation coefficient                                              | <i>r</i>                | endothelial-derived relaxing factor                    | EDRF                  |
| cubic                                                                | cu.                     | epithelial-derived relaxing factor                     | EpDRF                 |
| degree of freedom (statistics)                                       | d.f.                    | equilibrium constant                                   | <i>K</i>              |
| deoxyribonucleic acid                                                | DNA                     | equivalent (general use)                               | equiv.                |
| deoxyribonuclease                                                    | DNase                   | erythrocyte                                            | r.b.c.                |
| fatty acids, nonesterified                                           | NEFA                    | erythrocyte sedimentation rate                         | ESR                   |
| figure(s) (with reference number)                                    | Figure(s)               | ethylenediaminetetracetic acid                         | EDTA                  |
| figure (diagram)                                                     | figure                  | excitatory postsynaptic potential                      | e.p.s.p.              |
| gas-liquid chromatography                                            | g.l.c.                  | experiment                                             | expt                  |
| glomerular filtration rate                                           | GFR                     | experimental                                           | exptl                 |
| haemoglobin                                                          | Hb                      | page/pages                                             | p./pp.                |
| half-life                                                            | <i>t</i> <sub>1</sub>   | para-                                                  | <i>p</i> -            |
| high-frequency                                                       | h.f.                    | paragraph                                              | para. or ¶            |
| high performance liquid chromatography                               | h.p.l.c.                | parts per million                                      | p.p.m.                |
| human serum albumin                                                  | HSA                     | per cent                                               | %                     |
| hydrogen-ion concentration                                           | [H <sup>+</sup> ]       | platelet activating factor                             | PAF                   |
| hydrogen-ion activity, negative logarithm of (hydrogen-ion exponent) | pH                      | posterior                                              | post.                 |
| 6-hydroxydopamine                                                    | 6-OHDA                  | probability (significance level in a statistical test) | <i>P</i>              |
| N-[2-Hydroxyethyl]piperazine-N'-[2-ethanesulphonic acid]             | HEPES                   | radioimmunoassay                                       | RIA                   |
| 5-hydroxyindoleacetic acid                                           | 5-HIAA                  | rectus (configuration by the sequence rule)            | R                     |
| 5-hydroxytryptamine                                                  | 5-HT                    | red blood corpuscle                                    | RBC                   |
| immunoglobulins                                                      | IgA, IgD, IgE, IgG, IgM | relative band speed to front (chromatography)          | <i>R</i> <sub>F</sub> |
| inhibitor constant                                                   | <i>K</i> <sub>i</sub>   | relative molecular mass                                | <i>M</i> <sub>r</sub> |
| inhibitory concentration                                             | IC <sub>50</sub>        | relative retention time (gas chromatography)           | <i>t</i> <sub>r</sub> |
| inhibitory postsynaptic potential                                    | i.p.s.p.                | renal plasma flow                                      | RPF                   |
| insoluble                                                            | insol.                  | resistance (respiratory)                               | <i>R</i>              |
| international unit                                                   | iu                      | respiratory conductance                                | Sgaw                  |
| intra-arterial                                                       | i.a.                    | revolutions per minute                                 | r.p.m.                |
| intracellular fluid                                                  | ICF                     | ribonucleic acid                                       | RNA                   |
| intradermal                                                          | i.d.                    | section                                                | §                     |
| intramuscular                                                        | i.m.                    | sedimentation coefficient (ultracentrifugation)        | <i>s</i>              |
| intraperitoneal                                                      | i.p.                    | sinister (configuration by the sequence rule)          | S                     |
|                                                                      |                         | soluble                                                | sol.                  |
|                                                                      |                         | solution                                               | soln.                 |
|                                                                      |                         | Spearman rank coefficient                              | <i>r</i> <sub>s</sub> |
|                                                                      |                         | standard deviation                                     | s.d.                  |
|                                                                      |                         | (of observed sample)                                   |                       |

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

- (b) Only well-established and universally accepted subtype names (e.g. muscarinic and nicotinic acetylcholine receptors;  $\alpha$ - and  $\beta$ -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 M<sub>1</sub> 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 ( $\alpha$ 1)<sub>2</sub> $\beta$ 1 $\gamma$  $\delta$ .
- (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  $\beta$ -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 M<sub>1</sub>, M<sub>2</sub>, M<sub>3</sub>, M<sub>4</sub> and M<sub>5</sub>.

*Nicotinic acetylcholine receptors* The principal subgroups are muscle and neuronal nicotinic acetylcholine receptors.

- (b) *Adenosine receptors* Known also as P<sub>1</sub> purinoceptors (see Purinoceptors, 3v). (See Fredholm, B.B., *et al.*, (1994) *Pharmacol. Rev.*, **46**, 145–156.)
- (c) *Adrenoceptors* The principal subtypes are  $\alpha$ <sub>1</sub>-,  $\alpha$ <sub>2</sub>-,  $\beta$ <sub>1</sub>-,  $\beta$ <sub>2</sub>- and  $\beta$ <sub>3</sub>-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 AT<sub>1</sub> and AT<sub>2</sub>.
- (e) *Bombesin receptors* Proposed subtypes such as BB<sub>1</sub>, BB<sub>2</sub> may be used but must be fully referenced.
- (f) *Bradykinin receptors* The principal subtypes are B<sub>1</sub> and B<sub>2</sub> receptors.
- (g) *Calcitonin gene-related peptide (CGRP) receptors* Proposed CGRP receptor subtypes must be fully referenced.
- (h) *Cannabinoid receptors* The principal subtypes are CB<sub>1</sub> and CB<sub>2</sub>.
- (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<sub>A</sub> and CCK<sub>B</sub> receptors.
- (k) *Dopamine receptors* The principal subtypes are D<sub>1</sub>, D<sub>2</sub>, D<sub>3</sub>, D<sub>4</sub> and D<sub>5</sub>.
- (l) *Endothelin receptors* The principle subtypes are ET<sub>A</sub> and ET<sub>B</sub> 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<sub>A</sub> and GABA<sub>B</sub> receptors. Modulatory sites on the GABA<sub>A</sub> receptor should be referenced.
- (o) *Histamine receptors* The principal subtypes are H<sub>1</sub>, H<sub>2</sub> and H<sub>3</sub> receptors.
- (p) *5-Hydroxytryptamine (5-HT) receptors* The principal subtypes are 5-HT<sub>1</sub>, 5-HT<sub>2</sub>, 5-HT<sub>3</sub> and 5-HT<sub>4</sub>. 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<sub>n</sub>, (e.g. EP<sub>1</sub>, EP<sub>2</sub>) and referenced. (See Coleman, R.A., *et al.*, (1994) *Pharmacol. Rev.*, **46**, 205–229).
- (v) *Purinoceptors* The principal subtypes are P<sub>1</sub> and P<sub>2</sub> receptors. Subdivision of P<sub>1</sub> into A<sub>1</sub>, A<sub>2</sub> and A<sub>3</sub> subtypes and of P<sub>2</sub> into P<sub>2X</sub> and P<sub>2Y</sub> 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 NK<sub>1</sub>, NK<sub>2</sub> and NK<sub>3</sub> receptors.
- (y) *Vasoactive intestinal peptide (VIP) receptors* Proposed subtypes should be fully referenced.
- (z) *Vasopressin and oxytocin receptors* The principle subtypes are V<sub>1A</sub>, V<sub>1B</sub>, V<sub>2</sub> 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<sup>+</sup> channels, Na<sup>+</sup> channels, Ca<sup>2+</sup> channels and Cl<sup>-</sup> channels). Further subtypes must be fully referenced. For Ca<sup>2+</sup> 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-purinerbic, 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 serotoninergic. The term 5-HTergic is not acceptable.

Likewise, the terms purinerbic (ATP) and purinerbic (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) *EC<sub>50</sub>* 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 EC<sub>50</sub> values are determined in the presence of other agonists or antagonists the concentration of the latter should be stated. Related terms, e.g. EC<sub>25</sub>, are acceptable if accompanied by a full definition.
- (b) *IC<sub>50</sub>* 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.

#### 7 Enzymes

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

(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.  $\text{Na}^+$ ,  $\text{Ca}^{2+}$ ,  $2\text{Na}^+/\text{Ca}^{2+}$  exchange, etc.
- (h) *Inhibitors of nitric oxide synthase* The most commonly used and currently accepted abbreviations for  $\text{N}^G$ -nitro-L-arginine and  $\text{N}^G$ -nitro-L-arginine methyl ester are L-NOARG and L-NAME respectively.

## SPECIAL REPORTS

- A. Jovanovic & A. Terzic.** Diadenosine tetraphosphate-induced inhibition of ATP-sensitive K<sup>+</sup> channels in patches excised from ventricular myocytes 233
- A. Kawabata.** Evidence that endogenous nitric oxide modulates plasma fibrinogen levels in the rat 236
- P.G. McLean & I.M. Coupar.** Stimulation of cyclic AMP formation in the circular smooth muscle of human colon by activation of 5-HT<sub>4</sub>-like receptors 238
- PAPERS**
- P. Picard & R. Couture.** Intracerebroventricular responses to neuropeptide  $\gamma$  in the conscious rat: characterization of its receptor with selective antagonists 241
- P. Poulat, J. de Champlain & R. Couture.** Cardiovascular responses to intrathecal neuropeptide  $\gamma$  in conscious rats: receptor characterization and mechanism of action 250
- S. Baudet, E. Do, J. Noireaud & H. Le Marec.** Alterations in the force-frequency relationship by *tert*-butylbenzohydroquinone, a putative SR Ca<sup>2+</sup> pump inhibitor, in rabbit and rat ventricular muscle 258
- S. Baudet, A. Khammari, J. Noireaud & H. Le Marec.** Differential effects of *tert*-butyl-benzohydroquinone, a putative SR Ca<sup>2+</sup> pump inhibitor, on isometric relaxation during the staircase in the rabbit and rat ventricle 268
- J.H. St Lambert, M.R. Dashwood & K.M. Spyer.** Role of brainstem adenosine A<sub>1</sub> receptors in the cardiovascular response to hypothalamic defence area stimulation in the anaesthetized rat 277
- R. Verheggen, S. Freudenthaler, F. Meyer-Dulheuer & A.J. Kaumann.** Participation of 5-HT<sub>1</sub>-like and 5-HT<sub>2A</sub> receptors in the contraction of human temporal artery by 5-hydroxytryptamine and related drugs 283
- Z. Wang, M. Ma & R. Wang.** Enhanced vasocontraction of rat tail arteries by toxoflavin 293
- M. Hirafuji, A. Nezu, H. Shinoda & M. Minami.** Involvement of platelet cyclic GMP but not cyclic AMP suppression in leukocyte-dependent platelet adhesion to endothelial cells induced by platelet-activating factor *in vitro* 299
- P.J. White, R.B. Rose-Meyer & W. Hope.** Functional characterization of adenosine receptors in the nucleus tractus solitarius mediating hypotensive responses in the rat 305
- G.W. Bisset & K.M. Fairhall.** Release of vasopressin and oxytocin by excitatory amino acid agonists and the effect of antagonists on release by muscarine and hypertonic saline, in the rat *in vivo* 309
- C. Richer, V. Domergue, M.-P. Vincent & J.-F. Giudicelli.** Involvement of nitric oxide, but not prostaglandins, in the vascular sympatho-inhibitory effects of losartan in the pithed spontaneously hypertensive rat 315
- L.J.M. Cross, A.G. Beck-Sickinger, M. Bienert, W. Gaida, G. Jung, E. Krause & M. Ennis.** Structure activity studies of mast cell activation and hypotension induced by neuropeptide Y (NPY), centrally truncated and C-terminal NPY analogues 325
- M. Connor & G. Henderson.**  $\delta$ - and  $\mu$ -opioid receptor mobilization of intracellular calcium in SH-SY5Y human neuroblastoma cells 333
- F.J. Dowell, D. Henrion, M. Duriez & J.-B. Michel.** Vascular reactivity in mesenteric resistance arteries following chronic nitric oxide synthase inhibition in Wistar rats 341
- R.A. Silvestre, M. Salas, J. Rodríguez-Gallardo, O. García-Hermida, T. Fontela & J. Marco.** Effect of (8–32) salmon calcitonin, an amylin antagonist, on insulin, glucagon and somatostatin release: study in the perfused pancreas of the rat 347
- R. Ogata, Y. Inoue, H. Nakano, Y. Ito & K. Kitamura.** Oestradiol-induced relaxation of rabbit basilar artery by inhibition of voltage-dependent Ca channels through GTP-binding protein 351
- S. Illiano, R. Marsault, J.-J. Descombes, T. Verbeuren & P.M. Vanhoutte.** Regulation of nitric oxide-like activity by prostanoids in smooth muscle of the canine saphenous vein 360
- P.I. Aaronson, W. McKinnon & L. Poston.** Mechanism of butyrate-induced vasorelaxation of rat mesenteric resistance artery 365
- N. Maeda, T. Tamagawa, I. Niki, H. Miura, K. Ozawa, G. Watanabe, K. Nonogaki, K. Uemura & A. Iguchi.** Increase in insulin release from rat pancreatic islets by quinolone antibiotics 372
- R. Lemmens-Gruber, A. Karkhaneh, C. Studenik & P. Heistracher.** Cardiotoxicity of emetine dihydrochloride by calcium channel blockade in isolated preparations and ventricular myocytes of guinea-pig hearts 377
- C. Roberts, J. Watson, M. Burton, G.W. Price & B.J. Jones.** Functional characterization of the 5-HT terminal autoreceptor in the guinea-pig brain cortex 384
- Instructions to Authors** 389
- Nomenclature Guidelines** 397