Consensus nomenclature rules for radiopharmaceutical chemistry – setting the record straight (22nd November, 2016)

Preface

The primary function of nomenclature is to ensure that spoken or written scientific terms and concepts leave no ambiguity in their interpretation. The ultimate intent of generating consensus nomenclature is therefore to create common conventions for terms and definitions, enabling effective and unambiguous communication and understanding within a scientific community.

In order to achieve these goals, the international natural science community agreed to abide by and adopt the use of SI units (1960) and IUPAC rules for chemistry (1921). As the field of radiopharmaceutical chemistry is part of this wider community, it behoves us to also adopt these conventions: to ignore this would be impractical.

Over recent years, within our community, there has been an increased incidence of incorrect usage of established scientific terms and conventions, and even the emergence of 'self-invented' terms.

In order to address these concerns, an initiative was triggered by the 'Drug Development Committee' (DDC) of the 'European Association of Nuclear Medicine' (EANM). A Working Group (WG) on '*Nomenclature in Radiopharmaceutical Chemistry and related areas*' was established in 2015 to achieve clarification of terms and to generate consensus on the utilisation of a standardised nomenclature pertinent to our field.

The WG is internationally composed by Michael Adam (Canada), Gunnar Antoni (Sweden), Heinz H. Coenen (Germany) (chair), Cathy S. Cutler (USA), Yasuhisa Fujibayashi (Japan), Antony D. Gee (UK) (co-chair), Jae Min Jeong (Korea), Robert H. Mach (USA), Tom Mindt (Austria), Victor W. Pike (USA) and Albert Windhorst (Netherlands) as active members of relevant scientific societies (e.g., EANM, 'Society of Radiopharmaceutical Sciences', 'Society of Nuclear Medicine', 'Society of Radiopharmacology' and related national societies).

After conducting a worldwide survey by a questionnaire, aims of primary importance were agreed, and the WG produced a summary of its initial recommendations. These were used as 'guidelines to authors' for the submission of abstracts for the ISRS 2017 meeting. Other societies have also responded positively to using this summary as guidelines for future abstract submissions at their meetings. In addition to the summary document, a full text has now been prepared by the WG, which is presented here in a wider context.

It is now crucial to have a period of open consultation with peers in the field in order to gain feedback about the proposed recommendations and to discuss any further issues requiring clarification. The feedback of all of you on the newly posted nomenclature recommendation document is solicited, as some issues and terms are still unresolved, and suggestions for topics not addressed in the document are particularly welcome. We kindly request you to send any comments on the newly posted document to Heinz H. Coenen (<u>h.h.coenen@web.de</u>) and Antony Gee (<u>antony.gee@kcl.ac.uk</u>) before 31st Jan 2017.

After this deadline, the views expressed during the consultation period will be discussed and the resulting consensus will be incorporated into a pre-final document, which will be again posted on the SRS website, and any outstanding issues will be discussed at the 22nd ISRS meeting in Dresden (May 14th – 19th, 2017). Subsequently, the resulting consensus nomenclature guidelines will be published in an appropriate journal as a reference source for the field, and as a basis for discussion with journal editors, IUPAC and key stakeholders beyond our immediate peer group.

Scientific concepts, definitions and physical units - IUPAC rules and SI units

Below are summarised a number of terms (and descriptions thereof) of relevance to radiochemistry and related fields, which are already described and agreed by the wider scientific community, but are often used incorrectly in the literature. (for IUPAC see: https://iupac.org/what-we-do/nomenclature/ and for SI Units see: https://iupac.org/what-we-do/nomenclature/ and for https://iupac.org/what-we-do/nomenclature/ and for https://iupac.org/what-we-do/nomenclature/ and for SI Units see: https://iupac.org/what-we-do/nomenclature/ and for https://iupac.org/what-we-do/nomenclature/ and for https://iupac.org/what-we-do/model and for https://iupac.org/what-we-do/model and https://iupac.org/what-we-do/model and <a href="https://iupac.org/what-we-do/m

These are complemented by other terms, not described by international convention, but that have been adopted within our field and which have prompted discussion with a cross-section of experts within the field of radiochemistry in order to clarify and enhance the unambiguous communication of scientific findings and research results within the community.

Radioactivity measurement

SI units for radioactivity measurement, i.e. Bq (MBq, GBq etc.), should be used, while pre-SI units (e.g., imperial units) (e.g., mCi, Ci) can be placed in parentheses after the stated SI units (see Appendix).

The often used lab-jargon "hot" and "cold" must not be used instead of the correct terms "radioactive" and "non-radioactive", respectively, in formal public presentations, manuscripts or official documents.

Specific radioactivity and molar radioactivity

N.B.: Care should be taken when merely using the term 'activity' as it may cause confusion with other terms such as 'chemical activity'; e.g., of ligand binding, enzyme action, etc. (see, for example, the description of 'activity coefficient' in standard Physical Chemistry text books).

According to SI convention, the term 'specific' refers to a physical property as a function of the mass of the material in question; e.g., the specific heat capacity is the heat capacity of an object per kg of mass. Since in chemistry 'mass' is most often denoted in moles, related chemical properties are indicated in 'molar' units; e.g.,

molar volume. Because of this possible source of confusion the following terms are to be used correctly:

Specific radioactivity - the measured radioactivity **per gram** of compound; measured in Bq/g or GBq/mg etc.

Molar radioactivity - the measured radioactivity **per mole** of compound; measured in Bq/mol or GBq/µmol, etc.

In the area of radiopharmaceutical chemistry, dealing with low (nanomolar) masses, the term 'specific radioactivity' or 'specific activity' should only be used in special situations, i.e., in situations where the molecular weight cannot be determined or in the context of nuclide development (such as the activity of irradiated target material). Since in the majority of cases the molecular weight of a radiolabelled probe is known, the term 'molar radioactivity' or 'molar activity' should be used instead of the term 'specific radioactivity'.

Due to radioactive decay, the measurement time for the specific radioactivity or molar radioactivity determination must be stated; e.g., 'the specific radioactivity was 50 GBq/mg' or 'the molar radioactivity was 50 GBq/ μ mol' 2 h after the end of nuclide production, at the end of synthesis, at time of administration, etc.'

Apparent molar radioactivity and apparent specific radioactivity

In cases where masses of other material are present in a radiolabelled compound preparation, the measured specific or molar radioactivity is lower than the true value. This often happens if non-labelled materials present in the synthesis mixture are not entirely removed from the labelled product during purification. Examples are precursor molecules (e.g., spiperone) or a complexing ligand (e.g., DOTA-TATE), which have not been fully removed during the final product purification (e.g., [*N-methyl*-¹¹C]spiperone or [¹⁷⁷Lu]Lu-DOTA-TATE, respectively), but also any other chemically different impurity.

In such cases the terms apparent molar radioactivity or apparent specific radioactivity take into account the mass of the labelled and non-radiolabelled impurities (using moles, or weight, respectively).

The term "pseudo" molar/specific activity must not be used, since the impurity has not been intentionally added.

Another term used in this context is "effective specific or molar radioactivity" which must be differentiated and which is still under discussion. This term is often used to consider other (unkown) material present in a sample prepared, competing with the labelled product in its chemical or biological reactions, for example a complexation process or binding to a target. In this case, however, the "effectivity" must be determined by an additional analytical process, since it is not simply described by the measured "radioactivity per total mass".

'No-carrier-added', 'carrier-free' and 'carrier added'

The non-quantitative terms "carrier-free" (c.f.), "no-carrier-added" (n.c.a.) and "carrier-added" (c.a.) can be used as a general indication of specific/molar radioactivity levels. These terms have already been adequately defined by A.P. Wolf (J. Label. Compd. Radiopharm. **18**, 1-2 (1981)) and are also discussed in the IUPAC draft of 2014. It is advised, however, that, the term "carrier-free" should only be used in cases where an analytical verification has proven this state, i.e., that the theoretical specific or molar activity is reached.

As Eckelman, Volkert and Bonardi (Nucl. Med. Biol. **35**, 523–527 (2008)) and Lapi and Welch (Nucl. Med. Biol. **40**, 314-320 (2013)) pointed out, routinely used radionuclides (e.g., ¹¹C, ¹⁸F, ^{99m}Tc, ¹²³I, etc.) are never "carrier-free". Thus, it is recommended to generally avoid the term "carrier-free" altogether. N.B.: While 'carrier-free' implies 'no-carrier-added', the reverse is not true!

Radionuclide and radioisotope descriptors

The enrichment of a chemical compound with an isotope (stable or radioactive) of one or more of the elements of which it is constituted, is indicated by the symbol of the element, together with its mass number, in square brackets immediately preceding (i.e., with no hyphen or space) the compound's name or chemical formula; e.g., $[^{2}H, ^{14}C]$ benzene, or $[^{2}H, ^{14}C]C_{6}H_{6_{n}}$ for benzene enriched or labelled with stable deuterium and radioactive carbon-14 (see footnote).

If a symbol of an element is given in a chemical formula, or in combination with the name of a chemical compound, together with a mass number "**AE**", but without square brackets, according to IUPAC (see footnote ref. of Red Book), this indicates an isotopically substituted compound, having a composition such that all molecules of the compound only consist of the indicated (radio)nuclide. This means that the theoretical specific or molar activity of the atom or compound is attained; i.e., it is strictly 'carrier-free', a state rarely achieved in practice (see 'molar radioactivity' above).

Examples of correct and incorrect descriptions of isotopically labelled compounds: L-[¹¹C]methionine or [¹¹C]L-methionine are correct, however, L-¹¹C-methionine or L-[¹¹C]-methionine are incorrect!

Rules for designating labelling positions (e.g., L-[*methyl*-¹¹C]methionine or L-[*carboxyl*-¹¹C]methionine), are described in the IUPAC document: <u>http://goldbook.iupac.org/index.html</u>.

In the case of "fluorobenzene" labelled with fluorine-18, [¹⁸F]fluorobenzene is correct, while [¹⁸F]benzene is incorrect, since benzene does not contain a fluorine atom. Likewise, for technetium-III, forming a 1:1 complex with DTPA, [^{99m}Tc]TcDTPA²⁻ is correct, while [^{99m}Tc]DTPA²⁻ or ^{99m}Tc-DTPA²⁻ are incorrect, since the chelator itself does not contain a technetium atom.

According to these conventions, isotope symbols in square brackets in combination with nouns or verbs are meaningless and are to be avoided in a published chemical text or presentation. Instead, the element symbol together with the mass number must be used without any brackets. There is no contradiction or likelihood of confusion with the indication of a carrier-free state (see above), since nouns and verbs cannot be enriched or labelled with the indicated isotope.

Consequently, for example [¹¹C]compound, [¹²⁵I]-substitution, [¹⁸F]-derivative and [⁶⁸Ga]conjugate, are incorrect terms, since these nouns are not names of "chemical compounds". These should instead read as follows: ¹¹C-compound, ¹²⁵I-substitution, ¹⁸F-derivative and ⁶⁸Ga-conjugate (note: with hyphen!), or preferably: ¹¹C-labelled compound, substitution with iodine-125, ¹⁸F-tagged derivative, and ⁶⁸Ga-labelled conjugate. Analogously, ⁶⁸Ga-complex of a chelator, ¹¹¹In-chelate, ¹²⁴I-iodinated antibody, or ^{99m}Tc-labelled conjugate are to be used.

Equally, terms commonly found in literature such as [¹¹C]labelling, [⁶⁴Cu]-labelling, or [¹⁸F]-(radio)fluorination are erroneous, because 'labelling' and 'radiofluorination' are verbs and nouns and of course do not contain "chemical elements". These should instead read: ¹¹C-labelling, ⁶⁴Cu-labelling and ¹⁸F-fluorination, while the prefix 'radio' is redundant here.

Correspondingly, e.g., fluorine-18, technetium-99m, etc., should be used, rather than "¹⁸F" and "^{99m}Tc" (at least not without definition) as this would, strictly speaking, infer a carrier-free status for the radionuclide (see above). Generally, terms such as 18F, F18, F-18, or 99mTc, Tc99m, Tc-99m must not be used.

Likewise, "⁷⁶Br-" (the bromine-76 anion) is more accurately described by the terms [⁷⁶Br]bromide ion or [⁷⁶Br]Br-, and by analogy, [¹⁷⁷Lu]Lu³⁺ is correct for the description of the [¹⁷⁷Lu]lutetium cation rather than "¹⁷⁷Lu³⁺".

Compounds labelled with metallic radionuclides and their complexes follow of course the same conventions as given above for covalently labelled compounds. Examples: [²²³Ra]RaCl₂, [^{99m}Tc]NaTcO₄, [^{99m}Tc]Tc-MDP, and [^{99m}Tc]Tc-MIBI. Examples of radiometal-labelled conjugates: "[⁶⁸Ga]Ga-chelator-Z" (where "Z" is a place holder for a molecule to which the [metal(ligand)_n] complex is attached to, e.g., a peptide or antibody such as in [⁶⁸Ga]Ga-DOTA-TATE and [⁸⁹Zr]Zr-DFO-trastuzumab. It should be mentioned that square brackets are of course also used to denote metal complexes, and care should be taken to avoid confusion (see also the above cited: Nomenclature of Inorganic Chemistry, IUPAC Recommendations 2005 (www.**iupac**.org/fileadmin/user upload/databases/Red Book 2005.pdf). Illustrative examples include [^{99m}Tc][Tc(CO)₃(OH₂)₃]⁺, [¹¹¹In][In(DTPA)]²⁻, [¹¹¹In][In(oxyquinoline)₃] ([¹¹¹In]In-oxine), or [⁶⁴Cu][Cu(ATSM)].

The terms '(radio)isotope' and '(radio)nuclide' are often used incorrectly in texts, e.g., erroneously inferring that "isotope" means "radioactive nuclide" or even "labelled compound".

N.B.: All (radio)isotopes are (radio)nuclides, while the reverse is not true! For example, both the nuclear reactions ¹⁷⁶Yb(n,g)¹⁷⁷Yb (induced by thermal neutrons) and ¹²⁴Xe(p,2n)¹²³Cs (induced by charged particles) produce radionuclides, but only the first one leads to a radioisotope of the starting material. For the sake of clarification the definition of these terms are repeated below:

- 'Nuclide' indicates an atom, characterised by its numbers of protons (identifying its elemental nature) and of nucleons (indicating its mass). There are isobaric, isotonic, isodiapheric, and isotopic nuclides (see chart of the nuclides), which can be stable or radioactive, e.g., ¹H and ²H are stable, ³H is radioactive.
- 'Isotopes' are nuclides of the same element (same proton number), but having different numbers of neutrons (hence different atomic mass). Isotopic nuclides of different energy state are called isomeric nuclides, isomeric isotopes or isomers, such as technetium-99g and -99m.
- 'Isobars' are nuclides with the same mass number, such as ruthenium-100, technetium-100, molybdenum-100, etc.
- 'Isotones' are nuclides with the same number of neutrons, but different numbers of protons, such as hydrogen-2 (deuterium) and helium-3, or lithium-8, beryllium-9, boron-10, carbon-11, nitrogen-12 and oxygen-13.
- 'Isodiapheres' are nuclides with the same difference of neutrons and protons, such as boron-10, carbon-12, nitrogen-14, oxygen-16, fluorine-18, neon-20, etc. (difference: zero), or titanium-49, vanadium-51, chromium-53, manganese-55, iron-57, etc. (difference: five excess neutrons).

Radiochemical yield (RCY)

Prior to a discussion of radiochemical yields, two facts should be considered:

- Synthetic chemistry is the science of combining elements and molecules to form compounds in proportion to their components; i.e., in relation to their masses. Since it is the number of atoms/molecules that are generally referred to, masses are usually expressed in moles.
 - (Example: 1 mole of carbon is combusted with 1 mole oxygen gas to form 1 mole of carbon dioxide; $C + O_2 \rightarrow CO_2$)
- Radiochemistry is the chemistry of radioactive materials (elements, molecules). With the exception of the field of 'hot-atom-chemistry', the standard laws and conventions of chemistry still apply, with the exception of accounting for radioactive decay.

The correction for the decay of two (or more) radioactive samples to an identical point in time enables the law of relative masses to be applied; i.e., the application of established chemistry, concepts, definitions and terms.

'Radiochemical yield', calculated using decay-corrected radioactivity values for products and starting compounds, is identical to the concept of 'chemical yield'. Logically, the reference time for correction of decay must be identical for describing a particular reaction, irrespective of whether it is chosen to be the end of radionuclide production, the end of bombardment, the start of synthesis, the end of synthesis, or any convenient reference time point.

Radiochemical yield (RCY) (Definition)

Radiochemical yield is the amount of radioactivity in the product expressed as the percent of related starting radioactivity used in the corresponding synthesis (step). The **quantity of both must be decay corrected to the same point in time** before the calculation is made.

N.B.: Sometimes the amount of radioactivity produced at the end of a nuclide production cannot be (or is not) determined, for example with gaseous compounds such as [¹¹C]CO. It is, however, recommended to measure the radioactivity of resultant product or to determine the 'trapping efficiency' of the labelled starting material and use this to correct the RCY calculation.

If it is accepted that **radiochemical yields are always decay corrected** (as is general convention in the wider nuclear and radiochemistry fields), it makes 'newly created' terms to describe yields with radioactive materials superfluous, enabling an approach that is consistent with mainstream chemistry nomenclature; and it simplifies the understanding of our scientific findings and concepts within and outside the field of radiochemistry.

Furthermore, it is good practise to report if the radiochemical yield refers to an isolated or non-isolated product.

Consider the following example. "The (radio)synthesis of compound Y": The first step of reaction of A and B formed compound C, which was converted into D, oxidised to E and finally hydrolysed to product Y which was isolated by preparative HPLC.

Description of chemical yields: The overall yield of product "Y" was 40 %. While the yield of C after the first step was 90 %, the yield of the conversion of C to D amounted only to 50 %; but the yield of oxidation and hydrolysis were almost quantitative. *Description of radiochemical yields*: The radiochemical yield of product "Y" was 40 %. While the radiochemical yield of C after the first step was 90 %, the radiochemical yield of the conversion to D amounted only to 50 %; but the radiochemical yield of oxidation and hydrolysis were almost quantitative.

If it is understood that radiochemical yield is identical to chemical yield, the sentences are identical and easily comprehended. If non-decay corrected yields were used. The 'yields' would be totally different to the standard chemical description of yields.

By comparison, colloquial terms for "radio-yield" found in literature are neither necessary nor helpful: e.g., expressions such as 'radiochemical conversion', 'analytical radiochemical yield', 'radio-HPLC yield' must not be used as a surrogate for the accepted terms 'radiochemical yield' or 'radiochemical purity'. If these terms were to be used, the previous example would be even more nonsensical. Also, a term such as 'radiochemical conversion' yield might give the impression that there is a nuclear change. Furthermore, other expressions such as 'radioincorporation', 'radio-oxidation', 'radiohydrolytic' yield etc. would have the

same justification.

Equally, 'analytical radiochemical yield', 'radio-HPLC yield' should not be used as a surrogate for the accepted term radiochemical purity (RCP).

The following are examples of good practice when describing radiochemical yields: 'The radiochemical yield of "Y" was 67 % (based on HPLC analysis of the crude **product**)."

"The radiochemical yield* of "Y" was 67 %", with the following as a footnote:

"*determined by radio-HPLC analysis of the crude product", or

"*non-isolated, estimated by radio-HPLC,

or, in the general experimental section: "All radiochemical yields were determined by radio-HPLC analysis of the crude product, unless stated otherwise."

or alternatively use: 'The radiochemical purity of the crude product was 67 %.", "The radiochemical yield of "Y" determined from an aliquot of the reaction solution amounted to 67 %."

or: "The radiochemical yield of crude "Y" was 32% based on the amount of radioactivity eluted from the HPLC column".

Expressions such as 'conversion' or 'incorporation' may, however, be used in a semantic sense and even are indispensable in context of mechanistic discussions to avoid over-repetition of the same phrase in a text. For example: "The 'conversion' (or 'incorporation') proceeded with 50 % yield. Here it is clear from the context, that the radiochemical yield of the conversion or incorporation is intended. In this case, the prefix "radio" is to be avoided.

Radioactivity yield

Radioactivity yield is the amount of radioactive product expressed in Bq (MBq, GBq), which is obtained from a starting amount of radioactivity (e.g., produced from a cyclotron) and is **not corrected for decay**.

This term is useful, or necessary to indicate the amount of radioactive product obtained from a starting amount of radioactivity. If this is expressed as a "non-decay-corrected radiochemical yield" in %, it is significantly dependent on losses due to the technical manipulations used, and on their duration, in addition to the yield of the labelling reaction.

Thus, if a "radioactivity yield" is stated, e.g., to demonstrate the (economic) efficiency of a production process, the time required for all production steps should be carefully described in order to make results comparable! A rigorous scientific

report/publication will indicate the length of reaction times used in addition to the time required for other technical manipulations. In this case, starting radioactivity levels can be calculated from reported radiochemical yields.

In an experimental section it should also state if a yield is estimated using the measured radioactivity of the isolated product, or if it was estimated, for example, by HPLC analysis of a sample of the crude product.

It is further recommended to specify how radioactivity, specific activity, etc. are analysed and measured; e. g., determined by HPLC. Although normally reported in the experimental section of publications, it may also be useful to include these clarifications in footnotes on slides and electronic presentations.

Definitions of purity

Chemical purity

Chemical purity is the absence of other chemical compounds/species. (N.B.: 1. Chemically pure samples may contain isotopically labelled material! 2. Chemical purity is often erroneously described as ratio of the mass of carrier to the mass of other impurities. However, this is erroneous and would lead to a nonsensical result, i.e., that as the level of carrier decreases (and molar activity increases) the chemical purity would decrease! It is strongly discouraged, to report chemical purity in this manner.)

Radiochemical purity

Radiochemical purity is the absence of other radiochemical compounds/species. (N.B.: Radiochemically pure samples may contain other non-radioactive chemicals.)

Radionuclidic purity

Radionuclidic purity is the absence of other radionuclides.

Radioisotopic purity

Radioisotopic purity is the absence of other radioisotopes. This refers to radioisotopes of the same element, but not to radionuclides of other elements!

N.B.: All these purities are normally expressed as the degree of purity in %, for example the 'radionuclidic purity' as the ratio, given as a percentage, of the radioactivity of the radionuclide concerned to the total radioactivity of the preparation, and so on.

Examples:

- i) If there is a sample of acetyl-salicylic acid, labelled with carbon-11 in both carboxylic acid groups (acetyl- and benzoyl-position), this material is chemically, radionuclidically, and radioisotopically pure, but not radiochemically.
- ii) Iodine-123 labelled iodo-bromo-benzene may be radioisotopically pure (i.e., containing no other radioiodine isotopes), but it may unintentionally contain bromine-77 and would therefore not be radionuclidically nor radiochemically pure.

Physical units

Although it is not specifically the theme of these guidelines on nomenclature, attention must also be paid to the correct use of physical units, such as using the correct term for a given unit (see above molar radioactivity for Bq/mole). An example, often occurring in reports on nuclide production, is the incorrect use of "MBq/ μ Ah" to represent the radioactivity produced per μ A beam current during a 1 hour irradiation. Actually, yield of radioactivity means activity produced per current or per number of charged particles applied.

Thus, it is strongly encouraged to report either physical yields in units activity per charge (Bequerel per Coulomb, MBq/C) or saturation yields as activity per current (Bequerel per Ampere, MBq/ μ A), since everything else can be calculated from those parameters.

This issue has been under discussion for many years. In some instances 'measures' using other units may also be justified. If this is the case, definitions must be properly given. Such terms and their utility are explained on pp. 281-283 in the appendix of the IAEA-Tecdoc-1211 (2001): "Charged particle cross-section database for medical radioisotope production: diagnostic radioisotopes and monitor reactions". The topic was recently dealt with in great detail by Otuka and Takacs in "Definitions of radioisotope thick target yields" (Radiochim. Acta **103**, 1-6 (2015)).

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Footnote

The convention for naming specifically and selectively labelled compounds are concisely described in 'Nomenclature of Inorganic Chemistry, IUPAC Recommendations 2005': (www.iupac.org/fileadmin/user_upload/databases/Red_Book_2005.pdf). A more detailed treatment of these conventions are provided in chapter II-2 of 'Nomenclature of Inorganic Chemistry II, IUPAC Recommendations 2000' (Red Book II), and in the IUPAC Nomenclature of Organic Chemistry (Blue Book) prepared by Advanced Chemistry Development; found on the ACD website: http://acdlabs.com/iupac/nomenclature. Additionally, a new draft IUPAC document on "Terminology on carrier, specific activity, and purities in nuclear and radio-chemistry, radioanalytical and radiopharmaceutical chemistry" by Bonardi et al. (2014) is currently under discussion.

(*Text adapted from IUPAC Red Book:* "Hydrogen is an exception in that the three isotopes ¹H, ²H and ³H can have the alternative names protium, deuterium and tritium, respectively, and the symbols D and T may be used for deuterium and tritium.

However, ²H and ³H are preferred, because D and T can disturb the alphabetical ordering in formulae. These names give rise to the names proton, deuteron, triton for the cations ¹H⁺, ²H⁺ and ³H⁺, respectively. Because the word 'proton' is often used in contradictory senses, i.e., for isotopically pure ¹H⁺ ions on the one hand, and for the naturally occurring undifferentiated isotope mixture on the other, it is recommended that the undifferentiated mixture be designated by the name hydron, derived from hydrogen.")

It has to be pointed out, that the lower-case characters p, d, t and the symbol α are also valid descriptors for the ions of hydrogen and helium, respectively, and are generally used when describing nuclear reactions, e.g., ¹⁴N(p, α)¹¹C, or isotopically substituted solvents, such as DMSO-d₆ in NMR-spectroscopy.

Appendix

Definitions and (SI*) Units of Radiological Quantities

Quantity	Definition	SI-Unit	Old Unit	Conversion Factor
Activity	Number of radioactive disintegrations per time	Becquerel 1 Bq = 1 s ⁻¹	Curie 1 Ci = 3.7 · 10 ¹⁰ s ⁻¹	1 Ci = $3.7 \cdot 10^{10}$ Bq 1 Bq = $2.7 \cdot 10^{-11}$ Ci
Energy dose	Total absorbed radiation energy per mass	Gray 1 Gy = 1 J/kg	Rad 1 rad = 10 ⁻² J/kg	1 rad = 10 ⁻² Gy 1 Gy = 100 rad
Equivalent dose	Energy dose · quality factor of type of radiation	Sievert 1 Sv = 1 J/kg	Rem 1 rem = 10 ⁻² J/kg	1 rem = 10 ⁻² Sv 1 Sv = 100 rem
lon dose	Electrical charge of ions produced in 1 kg air by radiation	Coulomb/kg	Röntgen R	1 R = $2.58 \cdot 10^{-4}$ C/kg 1 C/kg = $3.876 \cdot 10^{3}$ R

*SI Units = International System of Units