

Synthesis of Carbon and Hydrogen Isotopomers of L-Tyrosine

W. Augustyniak, P. Suchecki, R. Kański, M. Kańska

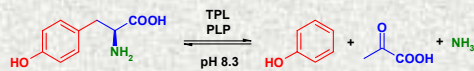
Department of Chemistry, University of Warsaw, Pasteur 1 Str. 02-093 Warsaw, Poland

Summary

[3-¹⁴C]-L-tyrosine was obtained via intermediate [3-¹⁴C]-L-phenylalanine, and its subsequent oxidation using enzyme L-phenylalanine monoxygenase. Isotopomers of L-tyrosine specifically labeled with deuterium and tritium were synthesized using novel enzymatic methods. Cheap commercial radiochemicals were used, i.e. Ba¹⁴CO₃, deuterated and tritiated water as label sources.

Introduction

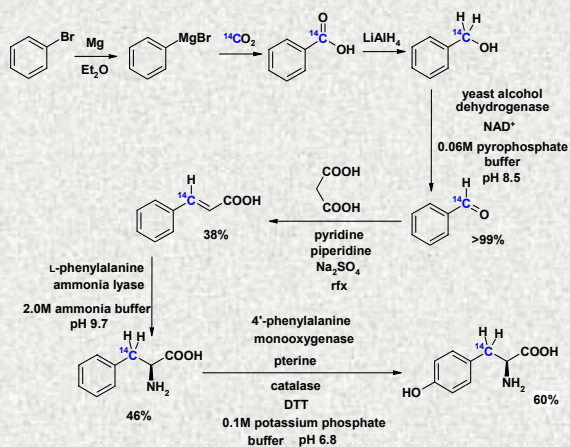
The enzyme *tyrosine phenol-lyase* (TPL, E.C 4.1.99.2) catalyzes reversible decomposition of L-tyrosine to phenol, pyruvate and ammonia [1].



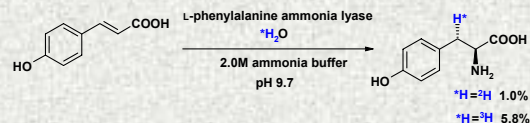
The mechanism of TPL action has already been studied, mostly using spectroscopy methods [2-5]. Our goal was to study the mentioned mechanism using the hydrogen and carbon kinetic isotope effects (KIE). This approach has not been attempted in a full extent yet; however, there is some isotopic data published [6-8].

Specifically labeled isotopomers of L-tyrosine are necessary to perform our research. We have already performed syntheses of 1- and 2- carbon-14 labeled L-tyrosines [9], as well as 2',6'-H₂, 3S-, 3R-, and 2-tritiated analogues [10]. Here we present new methods of their synthesis developed in our group.

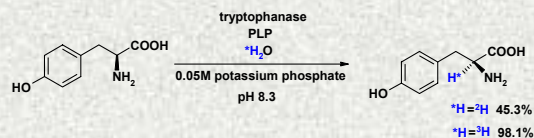
Synthesis of [3-¹⁴C]-L-tyrosine



Synthesis of 3S-hydrogen labeled L-tyrosine



Synthesis of 2-hydrogen labeled L-tyrosine



References

1. N. Brot, Z. Smit, H. Weissbach; *Arch. Biochem Biophys.*, 1-6, 112, 1965.
2. H. Chen; R. S. Phillips; *Biochemistry*, 11591-11599, 32, 1993.
3. R. S. Phillips; R. L. von Tersch; F. Secundo; *Eur. J. Biochem.*, 658-663, 244, 1997.
4. B. Sundararaju; A. A. Antson; R. S. Phillips; T. V. Demidkina; M. V. Barbolina; P. Gollnick; G. Guy Dodson; K. S. Wilson; *Biochemistry*, 6502-6510, 36, 1997.
5. H. Kumagai; S. Kitama; H. Yamada; T. Ueno; H. Fukami; *Pap. Meet. Agric. Chem. Soc. Jap.*, Kyoto, 1975.
6. D. M. Kiick; R. S. Phillips; *Biochemistry*, 7333-7338, 27, 1988.
7. B. S. Axelsson; P. Bjurling; O. Matsson; B. Langstrom; *J. Am. Chem. Soc.*, 1502-1503, 114, 1992.
8. N. G. Faleev; S. N. Spirina; T. V. Demidkina; R. S. Phillips; *J. Chem. Soc., Perkin Trans. 2*, 2001-2004, 1996.
9. W. Augustyniak; R. Kanski; M. Kanska; *J. Lab. Compd. Radiopharm.*, 553-560, 44, 2001.
10. W. Augustyniak; P. Suchecki; J. Jemielny; R. Kanski; M. Kanska; *J. Lab. Compd. Radiopharm.*, 559-567, 45, 2002.