Hydrodealkenylative C(Sp3)–C(Sp2) Bond Scission

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SUMMARY

UCLA researchers in the Department of Chemistry and Biochemistry have developed a new chemical reaction that combines ozone, an iron salt, and a hydrogen atom donor to enable hydrodealkenylative cleavage of C(sp3)–C(sp2) bonds in a widely applicable manner.

BACKGROUND

The commercial accessibility of complex molecules very often relies on starting material cost. There is a limit to the natural products, drugs and agrochemicals available for research and commercialization due to the complexity and cost of synthesizing chiral molecules. Traditionally, the assembly of complex molecules from simple precursors is accomplished, more often than not, by the need to install carbon centers with precisely defined stereochemical arrangements. Despite the bevy of methods available to accomplish such goals, sometimes it can be more efficient and cost-effective to reorganize starting materials already containing the required complexity and/or stereochemistry into the desired molecular structures. Such, deconstructive strategies can provide access to challenging, or otherwise inaccessible, molecular structures. While there are some examples of C(sp3)–C(sp2) bond scissions and C(sp3)–C(sp3) bond fragmentations, general methods for the functionalization of C(sp3)–C(sp2) bonds remain elusive. Given the profuse number of organic molecules containing these linkages, activation of such bonds in a controllable manner would be extremely useful.

INNOVATION

UCLA researchers targeted the less common deconstructive strategy toward complexity—particularly one involving C–C bond scission. The hydrodealkenylative cleavage of C(sp3)–C(sp2) bonds, performed using oxygen, an iron salt, and a hydrogen atom donor, has been developed by UCLA researchers. These reactions are simple to operate and reach completion within 30 minutes, delivering their products in high yields—even on decagram scales. Researchers have used this transformation to produce desirable synthetic intermediates and applied it in the total syntheses of complex molecules. With broad substrate scope and high functional group compatibility, this methodological advance lays new paths for the syntheses of organic molecules with utility in chemistry, biology, and medicine.

APPLICATIONS

- Cost-effective synthesis of chiral building blocks
- Medicinal chemistry methodology

ADVANTAGES

- Mild conditions
- High yielding
- Scalable method
- Broad substrate scope
- High functional group compatibility

STATE OF DEVELOPMENT

The developed methodology has been applied in the synthesis of desirable synthetic intermediates in high yields and up to decagram scale. Expensive chiral building blocks have been prepared from up to 10^3-fold cheaper starting materials.

PATENT STATUS

Patent Pending

RELATED MATERIALS


ADDITIONAL TECHNOLOGIES BY THESE INVENTORS

- Novel Non-Peptidomimetic Prenyltransferase Inhibitors
- Compound Library Made Through Phosphine-Catalyzed Annulation/Tebbe/Diels-Alder Reaction
Small Molecule Inhibitor of Cholesterol Biosynthesis and Venous Angiogenesis

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