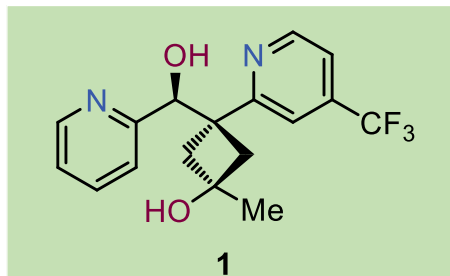


Background

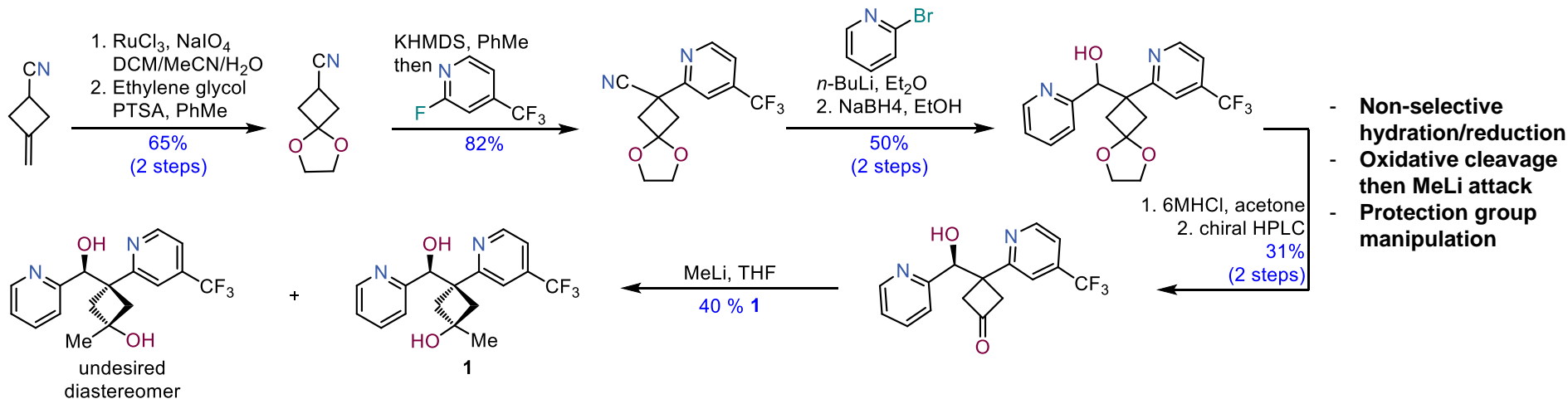
- Transient receptor potential vanilloid-3 (TRPV3) is one of the non-selective cation channels, found in neuronal and non-neuronal tissues, including dorsal root ganglia, spinal cord, and keratinocytes, could serve as the new target for analgesic agent.
- **1** show analgesic effect toward neuropathic pain in preclinical study.
- TRPV3 FLIPR $K_b = 0.56 \mu\text{M}$ ($0.62 \mu\text{M}$ plasma adjusted), $\text{Cl}_{\text{int,u}} = 2 \text{ (L/h/Kg)}$



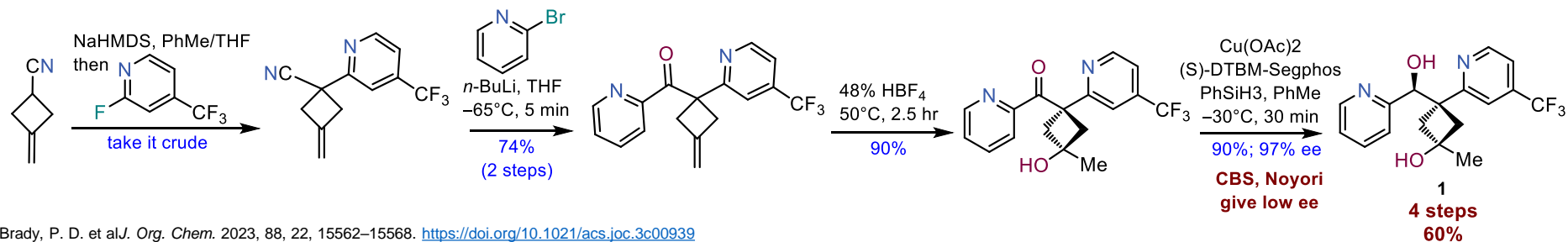
abbvie

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Initial Med. Chem Route



Optimized Route



Brady, P. D. et al. *J. Org. Chem.* 2023, 88, 22, 15562–15568. <https://doi.org/10.1021/acs.joc.3c00939>