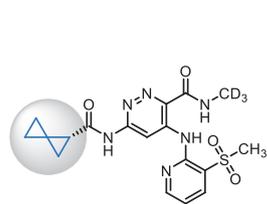


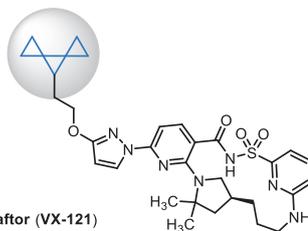
Metabolically Stabilized Lipophilic Groups

Introduction

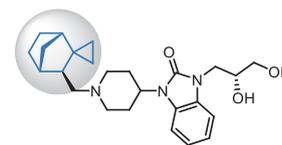
Protein binders frequently contain lipophilic groups that interact with hydrophobic surfaces within the protein interior. However, incorporating such groups into drug candidates often results in reduced metabolic stability, posing significant challenges for advancing effective bioactive molecules to clinics. Spirocyclic groups provide a viable solution to this issue. They increase volume and lipophilicity while being much less prone to metabolic clearance.¹⁻⁴ Enamine's collection of spirocyclic groups features a diverse range of ring sizes and molecular topologies as exemplified by the collection of spiroalkyl amines.



Lomeceutininib (BMS-986322)
treatment of psoriasis
Phase 2 trials
Bristol-Myers Squibb

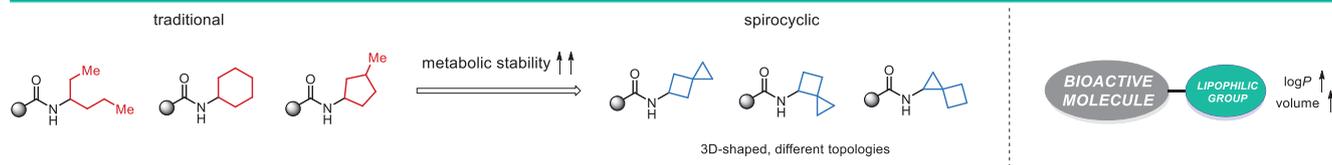


Vanzacaftor (VX-121)
treatment of cystic fibrosis
FDA approved 2024
Vertex Pharmaceuticals

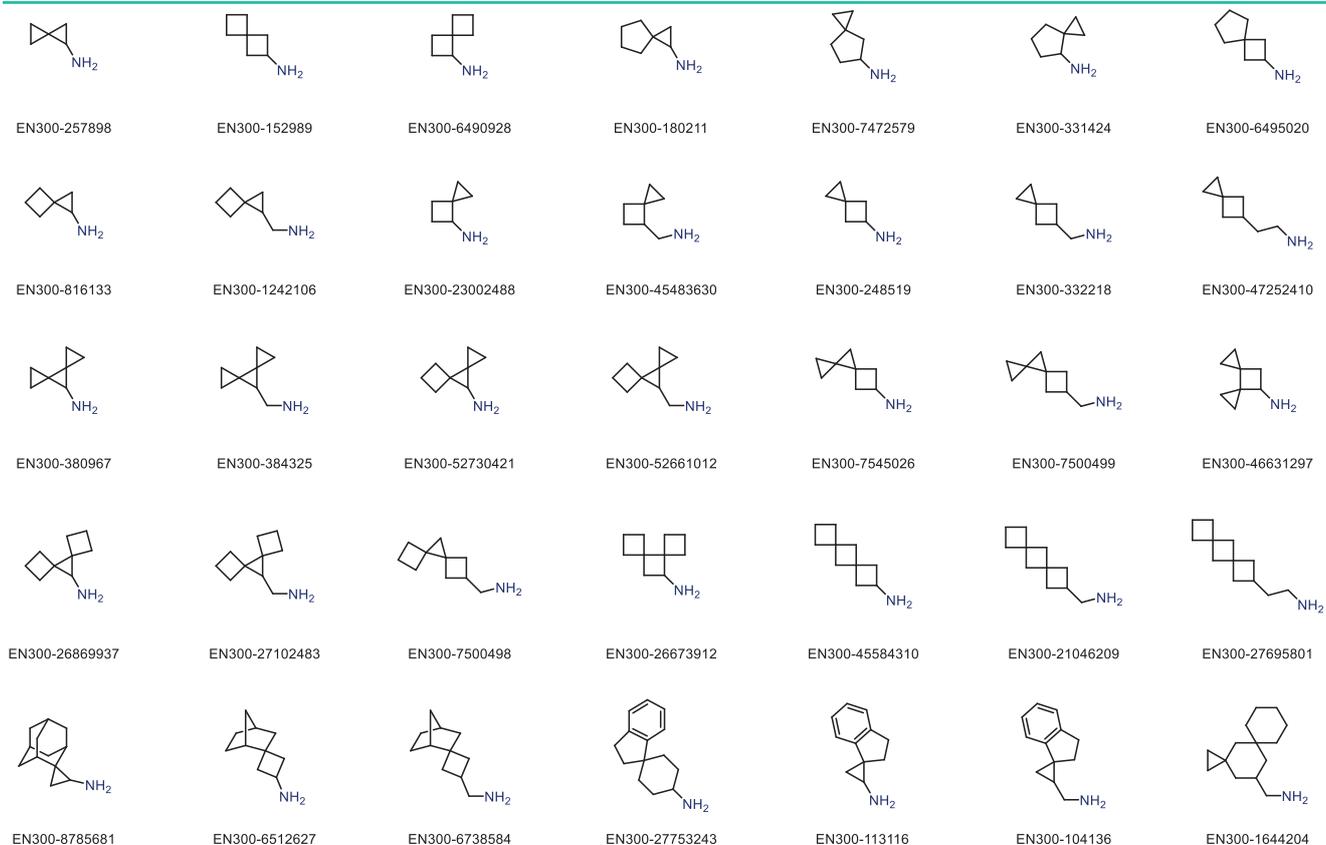


MK-5757
treatment of schizophrenia
Phase 2 trials
Merck

Concept



We offer: over 100 spiroalkyl amines from stock on 5-10 gram scale.



References

1. E. Benedetti and L. Micouin. *Expert Opin. Drug Discov.* **2024**, *19*, 263.
2. D. Candito et al. *J. Med. Chem.* **2022**, *65*, 16801.
3. D. Candito et al. *J. Med. Chem.* **2022**, *65*, 16801.
4. K. Prysiazniuk et al. *Angew. Chem. Int. Ed.* **2024**, *63*, e202316557.



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