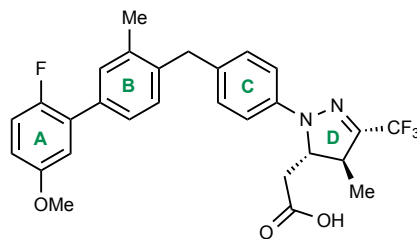
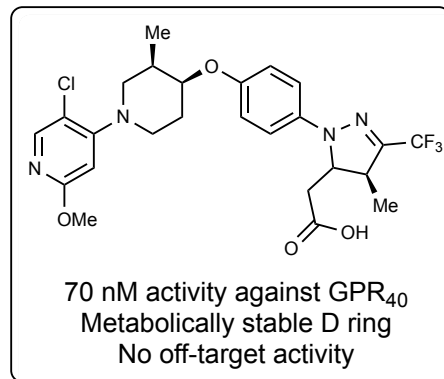
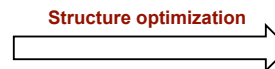


## BMS-986118:

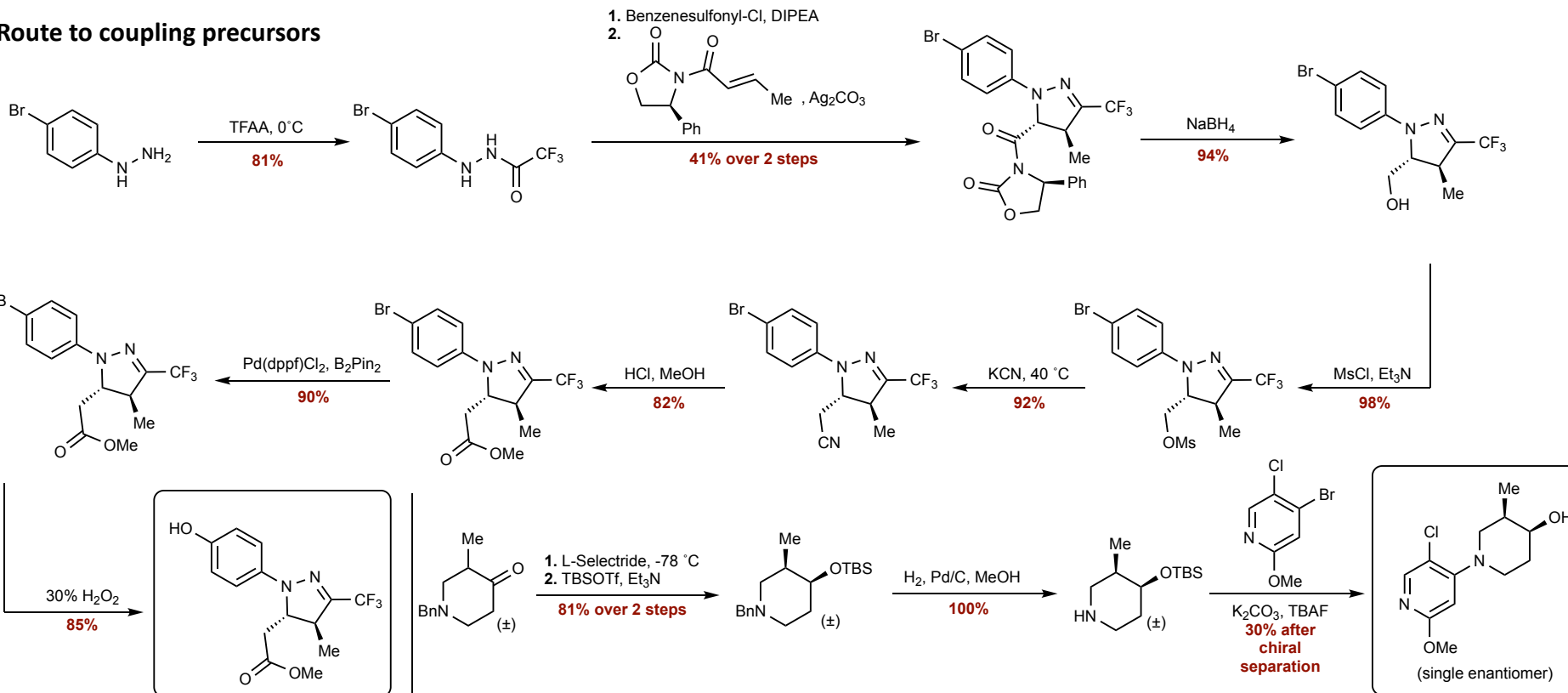
- G protein-coupled receptor inhibitor
- Kilograms of API required for clinical trials as diabetes treatment
- Initial route used Mitsunobu for etherification
  - Low yields, difficult to purify
- Pd etherification used for scaleup route
  - Higher yielding
  - Telescoping into ester cleavage allowed for easier purification

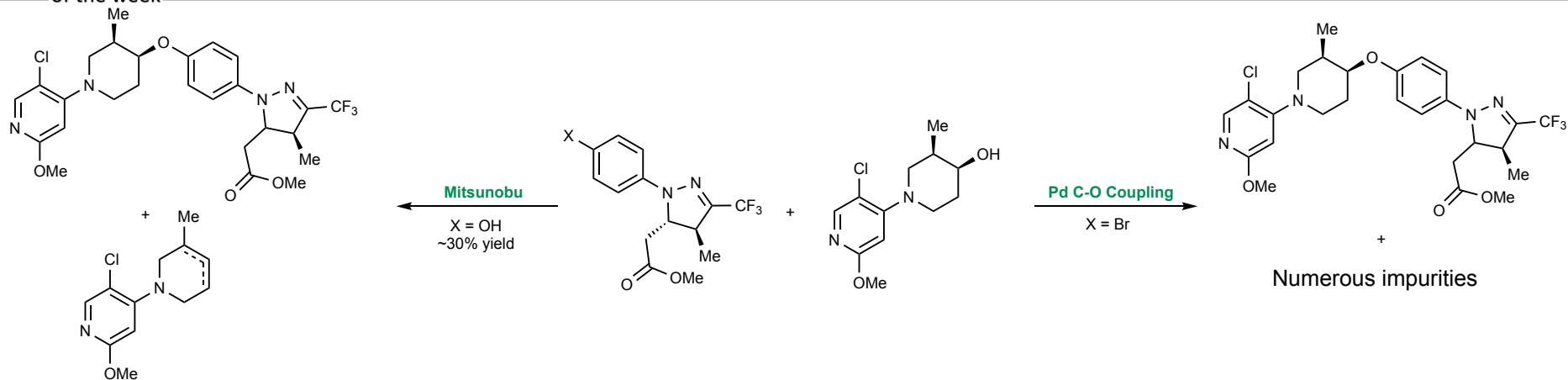


33 nM activity against GPR40  
A ring metabolically unstable  
Off-target activity *in vivo*



## Route to coupling precursors





## Coupling optimization

