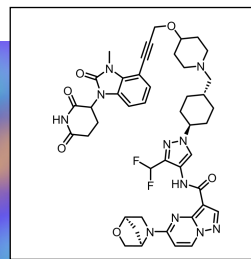


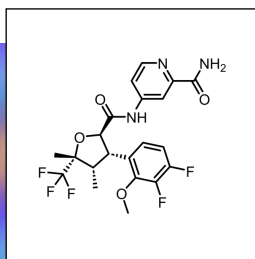
## Merck PCSK9 inhibitor

oral macrocyclic peptide inhibitor of PCSK9  
Ph. III for hypercholesterolemia  
from mRNA display + SBDD  
MK-0616  
MERCK, RAHWAY, NJ



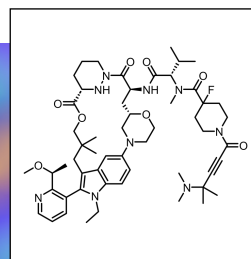
## Kymera/Sanofi IRAK4 degrader

oral IRAK4 degrader  
Ph. II in AD and HS  
from SBDD of ligands for CRBN, IRAK4, and linker  
KT-474  
KYMERA THERAPEUTICS, MA / SANOFI, NJ



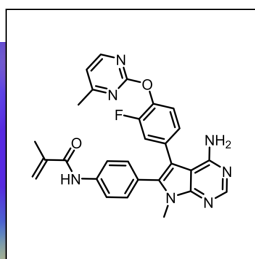
## Vertex NaV1.8 inhibitor

oral NaV1.8 selective inhibitor  
Ph. III for acute pain  
discovery undisclosed  
suzetrigine (VX-548)  
VERTEX PHARMACEUTICALS, SAN DIEGO, CA



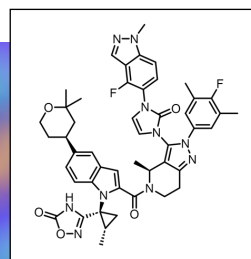
## Revolution Medicines KRAS<sup>G12C</sup> inhibitor

oral KRAS<sup>G12C</sup>(ON) tri-complex inhibitor  
Ph. I/II for adv. solid tumors  
from covalent tethering approach + SBDD  
RMC-6291  
REVOLUTION MEDICINES, REDWOOD CITY, CA



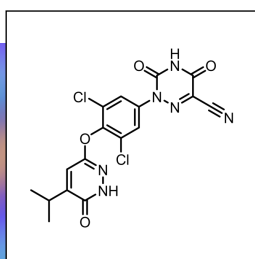
## Relay FGFR2 inhibitor

oral covalent FGFR2 inhibitor  
Ph. II for cholangiocarcinoma  
from rational design with MD simulations  
lirafugratinib (RLY-4008)  
RELAY THERAPEUTICS, CAMBRIDGE, MA



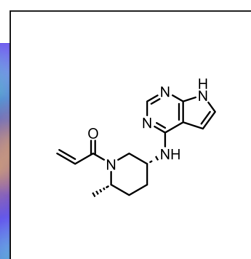
## Chugai/Lilly GLP-1R agonist

oral non-peptide GLP-1R agonist  
Ph. III for obesity + type 2 diabetes  
from LLC-PK1 cell HTS + opt  
orforglipron  
CHUGAI, SHIZUOKA, JP / ELI LILLY, IN



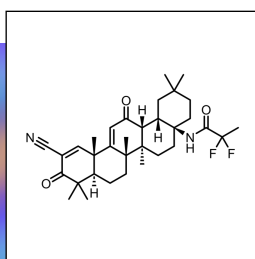
## Roche/Madrigal THR-β-selective agonist

oral THR-β-selective agonist  
Ph. III for NASH  
opt. of triiodothyronine  
resmetirom  
ROCHE, NUTLEY, NJ / MADRIGAL, PA



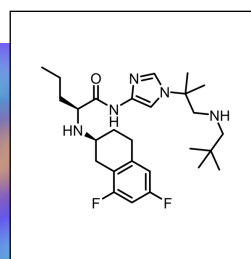
## Pfizer JAK3/TEC-family inhibitor

Oral JAK3/TEC-family kinase inhibitor  
approved for alopecia areata  
structure-based design from tofacitinib  
ritlecitinib (LITFULO®)  
PFIZER, GROTON, CT



## Reata/Biogen Nrf2 activator

oral once-daily Nrf2 activator  
approved to treat Friedreich's ataxia  
from modification of oleanolic acid  
omaveloxolone (SKYCLARYS®)  
REATA PHARMA, TX / BIOGEN, MA



## Pfizer/Springworks γ-secretase inhibitor

oral γ-secretase inhibitor  
approved for desmoid tumors  
single-digit nanomolar potency for Aβ reduction in vitro  
nirogacestat (OGSIVEO®)  
PFIZER / SPRINGWORKS THERAPEUTICS