



[www.MedChemExpress.com](http://www.MedChemExpress.com)

Inhibitors, Screening Libraries, Proteins

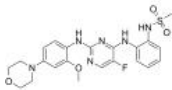
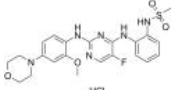
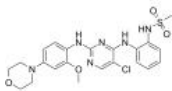
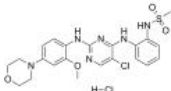
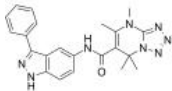
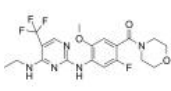
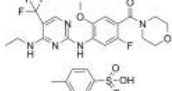
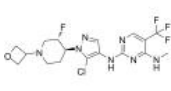
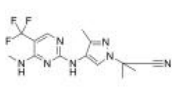
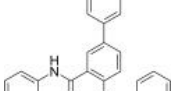
# LRRK2

## Leucine-rich repeat kinase 2

Leucine-rich repeat kinase 2 (LRRK2) is a ubiquitously expressed member of the ROCO protein family. LRRK2 is a complex, multidomain protein containing kinase and GTPase enzymatic activities and multiple protein-protein interaction domains. LRRK2 is the genetic cause of both familial and idiopathic Parkinson's disease (PD), and it is associated with neuronal death, vesicle trafficking, mitochondrial dysfunction, and inflammation.

LRRK2 is a very large protein comprised of 2527 amino acids which has been determined to contain multiple functional domains, including armadillo (ARM), ankyrin-repeats (ANK), leucine-rich repeats (LRR), Ras of complex proteins (ROC), C-terminal of Roc (COR), MAPK-like kinase, and WD40 motifs. Mutations in LRRK2 represent a significant component of both sporadic and familial PD. Pathogenic mutations cluster in the enzymatic domains of LRRK2, and kinase activity seems to correlate with cytotoxicity, suggesting the possibility of kinase-based therapeutic strategies for LRRK2-associated PD. The best-characterized mutation to date, G2019S, leads to increased kinase activity, and mutations in the GTPase domain, such as R1441C and R1441G, have also been reported to influence kinase activity.

## LRRK2 Inhibitors

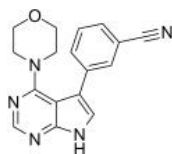
<p><b>CZC-25146</b></p> <p style="text-align: right;">Cat. No.: HY-15800A</p>	<p><b>CZC-25146 hydrochloride</b></p> <p style="text-align: right;">Cat. No.: HY-15800</p>
<p>CZC-25146 is a potent, selective and metabolically stable LRRK2 inhibitor with IC<sub>50</sub> of 4.76 nM/6.87 nM for wild type LRRK2 and G2019S LRRK2 respectively.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> 98.85%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg</p>	<p>CZC-25146 Hcl is a potent, selective and metabolically stable LRRK2 inhibitor with IC<sub>50</sub> of 4.76 nM/6.87 nM for wild type LRRK2 and G2019S LRRK2 respectively.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> &gt;98%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 1 mg, 5 mg</p>
<p><b>CZC-54252</b></p> <p style="text-align: right;">Cat. No.: HY-B0792</p>	<p><b>CZC-54252 hydrochloride</b></p> <p style="text-align: right;">Cat. No.: HY-B0792A</p>
<p>CZC-54252 is a potent and selective LRRK2 inhibitor with IC<sub>50</sub>s of 1.28 nM and 1.85 nM for wild-type and G2019S LRRK2, respectively. CZC-54252 attenuates G2019S LRRK2-induced human neuronal injury with an EC<sub>50</sub> of ~1 nM. CZC-54252 has a neuroprotective activity.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> 99.18%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 10 mM × 1 mL, 10 mg, 50 mg</p>	<p>CZC-54252 hydrochloride is a potent and selective LRRK2 inhibitor with IC<sub>50</sub>s of 1.28 nM and 1.85 nM for wild-type and G2019S LRRK2, respectively. G2019S LRRK2-induced human neuronal injury is attenuated by CZC-54252 hydrochloride with an EC<sub>50</sub> of ~1 nM.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> &gt;98%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 1 mg, 5 mg</p>
<p><b>EB-42486</b></p> <p style="text-align: right;">Cat. No.: HY-142647</p>	<p><b>GENE-7915</b></p> <p style="text-align: right;">Cat. No.: HY-18163</p>
<p>EB-42486 is a novel, potent, and highly selective G2019S-LRRK2 inhibitor (IC<sub>50</sub> &lt; 0.2 nM).</p> <p style="text-align: center;"></p> <p><b>Purity:</b> &gt;98%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 1 mg, 5 mg</p>	<p>GENE-7915 is a potent, selective and brain-penetrant inhibitor of LRRK2 with an IC<sub>50</sub> of 9 nM.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> 99.79%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 100 mg</p>
<p><b>GENE-7915 tosylate</b></p> <p style="text-align: right;">Cat. No.: HY-18163A</p>	<p><b>GENE-9605</b></p> <p style="text-align: right;">Cat. No.: HY-12282</p>
<p>GENE-7915 tosylate is a potent, selective and brain-penetrant inhibitor of LRRK2 with an IC<sub>50</sub> of 9 nM.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> 99.62%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 100 mg</p>	<p>GENE-9605 is a highly potent, selective, and brain-penetrant LRRK2 inhibitor with IC<sub>50</sub> of 19 nM. IC<sub>50</sub> value: Target: LRRK2 GNE-9605 retained excellent predicted human metabolic stability when assayed in human liver microsomes and hepatocytes.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> 99.72%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg</p>
<p><b>GENE0877</b></p> <p style="text-align: right;">Cat. No.: HY-15796</p>	<p><b>GSK2578215A</b></p> <p style="text-align: right;">Cat. No.: HY-13237</p>
<p>GENE0877 is a highly potent, selective, and brain-penetrant aminopyrazole leucine-rich repeat kinase 2 (LRRK2) small molecule inhibitor with an IC<sub>50</sub> of 3 nM.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> 98.89%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 10 mM × 1 mL, 2 mg, 5 mg, 10 mg, 25 mg, 50 mg</p>	<p>GSK2578215A is a potent and highly selective LRRK2 inhibitor, which exhibits IC<sub>50</sub>s of around 10 nM against both wild-type LRRK2 and the G2019S mutant.</p> <p style="text-align: center;"></p> <p><b>Purity:</b> 99.79%  <b>Clinical Data:</b> No Development Reported  <b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg</p>

<p><b>HG-10-102-01</b></p> <p>Cat. No.: HY-13488</p>	<p><b>IKK 16</b></p> <p>Cat. No.: HY-13687</p>
<p>HG-10-102-01 is a potent and selective inhibitor of wild-type LRRK2 (IC<sub>50</sub>=23.3 nM) and the G2019S mutant (IC<sub>50</sub>=3.2 nM). IC<sub>50</sub> Value: 23.3 nM (WT LRRK2); 3.2 nM (LRRK2 G2019S). Target: LRRK2. HG-10-102-01 maintains the ability to potently inhibit the biochemical activity of wild-type...</p> <p><b>Purity:</b> 99.34%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg</p>	<p>IKK 16 is a selective IκB kinase (IKK) inhibitor for IKK2, IKK complex and IKK1 with IC<sub>50</sub>s of 40 nM, 70 nM and 200 nM, respectively. IKK16 also inhibits leucine-rich repeat kinase-2 (LRRK2) with an IC<sub>50</sub> of 50 nM.</p> <p><b>Purity:</b> 99.09%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg</p>
<p><b>IKK 16 hydrochloride</b></p> <p>Cat. No.: HY-13687A</p>	<p><b>JH-II-127</b></p> <p>Cat. No.: HY-16936</p>
<p>IKK 16 hydrochloride is a selective IκB kinase (IKK) inhibitor for IKK2, IKK complex and IKK1 with IC<sub>50</sub>s of 40 nM, 70 nM and 200 nM, respectively. IKK16 also inhibits leucine-rich repeat kinase-2 (LRRK2) with an IC<sub>50</sub> of 50 nM.</p> <p><b>Purity:</b> ≥98.0%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 5 mg, 10 mg, 50 mg, 100 mg</p>	<p>JH-II-127 is a highly potent, selective, and brain penetrant LRRK2 inhibitor, with IC<sub>50</sub> of 6.6 nM, 2.2 nM, 47.7 nM for LRRK2-wild-type, LRRK2-G2019S, LRRK2-A2016T.</p> <p><b>Purity:</b> 98.02%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg</p>
<p><b>LRRK2 inhibitor 1</b></p> <p>Cat. No.: HY-111493</p>	<p><b>LRRK2-IN-1</b></p> <p>Cat. No.: HY-10875</p>
<p>LRRK2 inhibitor 1 is a potent, selective and oral LRRK2 inhibitor with an pIC<sub>50</sub> of 6.8.</p> <p><b>Purity:</b> 99.72%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg</p>	<p>LRRK2-IN-1 is a potent and selective LRRK2 inhibitor with IC<sub>50</sub> of 6 nM and 13 nM for LRRK2 (G2019S) and LRRK2 (WT), respectively.</p> <p><b>Purity:</b> 99.19%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 10 mM × 1 mL, 10 mg, 50 mg, 100 mg, 500 mg</p>
<p><b>LRRK2-IN-2</b></p> <p>Cat. No.: HY-145317</p>	<p><b>LRRK2-IN-3</b></p> <p>Cat. No.: HY-145318</p>
<p>LRRK2-IN-2 (compound 22) is a potent, selective, orally active and brain-penetrant inhibitor LRRK2, with IC<sub>50</sub> of &lt;0.6 nM. LRRK2-IN-2 can be used for the research of Parkinson's disease.</p> <p><b>Purity:</b> &gt;98%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 1 mg, 5 mg</p>	<p>LRRK2-IN-3 (compound 24) is a potent, selective, orally active and brain-penetrant inhibitor LRRK2, with IC<sub>50</sub> of 2.6 nM in human PBMCs. LRRK2-IN-3 can be used for the research of Parkinson's disease.</p> <p><b>Purity:</b> &gt;98%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 1 mg, 5 mg</p>
<p><b>LRRK2-IN-4</b></p> <p>Cat. No.: HY-144074</p>	<p><b>MLi-2</b></p> <p>Cat. No.: HY-100411</p>
<p>LRRK2-IN-4 is a potent, selective, CNS-penetrant and orally active leucine-rich repeat kinase 2 (LRRK2) inhibitor with an IC<sub>50</sub> of 2.6 nM. LRRK2-IN-4 has the potential for the research of Parkinson's disease.</p> <p><b>Purity:</b> &gt;98%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 1 mg, 5 mg</p>	<p>MLi-2 is an orally active and highly selective LRRK2 inhibitor with an IC<sub>50</sub> of 0.76 nM. MLI-2 has the potential for Parkinson's disease.</p> <p><b>Purity:</b> 99.74%</p> <p><b>Clinical Data:</b> No Development Reported</p> <p><b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg</p>

**PF-06447475**

Cat. No.: HY-12477

PF-06447475 is a highly potent, selective and brain penetrant **LRRK2** inhibitor with an  $IC_{50}$  of 3 nM.

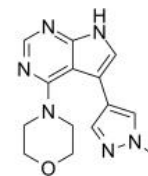


**Purity:** 99.88%  
**Clinical Data:** No Development Reported  
**Size:** 10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 100 mg, 200 mg

**PF-06454589**

Cat. No.: HY-112855

PF-06447475 is a highly potent, selective, brain penetrant **LRRK2 kinase** inhibitor with  $IC_{50}$  values of 3 nM and 11 nM for WT LRRK and G2019S LRRK2, respectively. PF-06447475 can be used for parkinson's disease (PD) research.

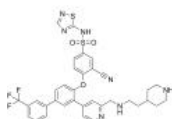


**Purity:** >98%  
**Clinical Data:** No Development Reported  
**Size:** 5 mg, 10 mg, 25 mg, 50 mg, 100 mg

**PF-06456384**

Cat. No.: HY-118952

PF-06447475 is a highly potent, selective, brain penetrant **LRRK2 kinase** inhibitor with  $IC_{50}$  values of 3 nM and 11 nM for WT LRRK and G2019S LRRK2, respectively. PF-06447475 can be used for parkinson's disease (PD) research.

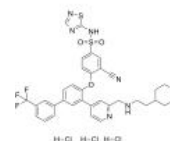


**Purity:** >98%  
**Clinical Data:** No Development Reported  
**Size:** 1 mg, 5 mg

**PF-06456384 trihydrochloride**

Cat. No.: HY-118952A

PF-06447475 trihydrochloride is a highly potent, selective, brain penetrant **LRRK2 kinase** inhibitor with  $IC_{50}$  values of 3 nM and 11 nM for WT LRRK and G2019S LRRK2, respectively. PF-06447475 trihydrochloride can be used for parkinson's disease (PD) research.



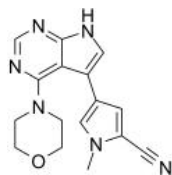
**Purity:** >98%  
**Clinical Data:** No Development Reported  
**Size:** 25 mg

**PFE-360**

(PF-06685360)

Cat. No.: HY-120085

PFE-360 (PF-06685360) is a potent, selective, brain penetrated and orally active **leucine-rich repeat kinase 2 (LRRK2)** inhibitor with a mean  $IC_{50}$  of 2.3 nM in vivo.



**Purity:** 98.94%  
**Clinical Data:** No Development Reported  
**Size:** 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg