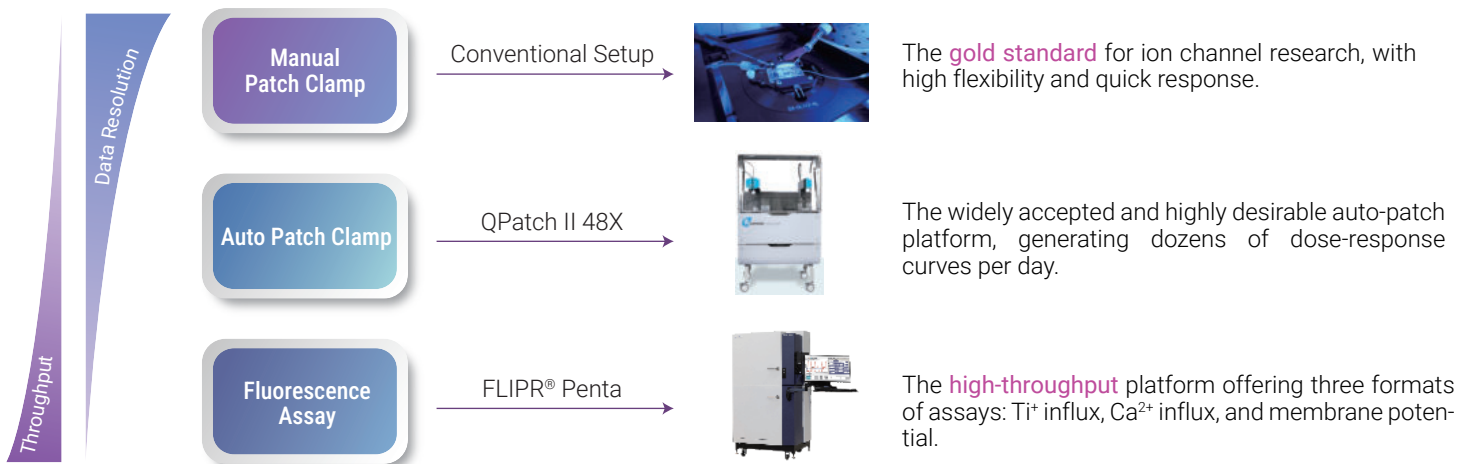


# Electrophysiology Services

Since 2010, our expert electrophysiology team has led the way in early drug discovery, specializing in ion channel screening and cardiac safety assessment. Our reputation is built on comprehensive services from target validation to preclinical candidate identification.

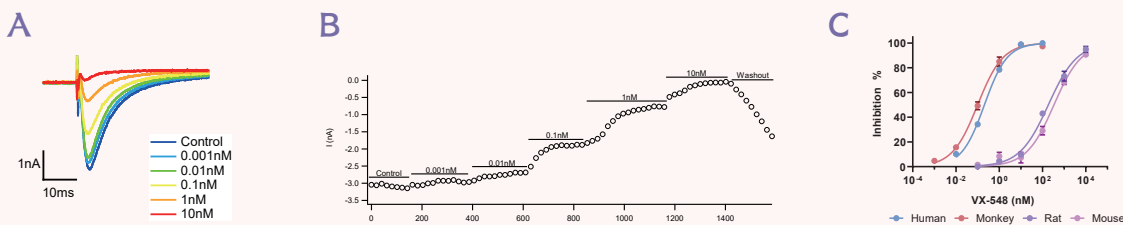
## Ion Channel Screening



### Key Features of Manual Patch Clamp Services

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- ◇ **100+** dose-response curves weekly
- ◇ **35+** scientists with **5+** years of experience
- ◇ **High Flexibility** in MoA exploration: Tailoring specific protocols for agonists, antagonists, PAMs, NAMs, use-dependent drugs, etc.
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- ◇ **17** rigs of manual patch clamp
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### Example Data



Species	IC <sub>50</sub>
Human	0.2067nM
Cyno Monkey	0.1005nM
Rat	186.4nM
Mouse	339.0nM

### Species-Selective Inhibition of Nav1.8 Channels by VX-548

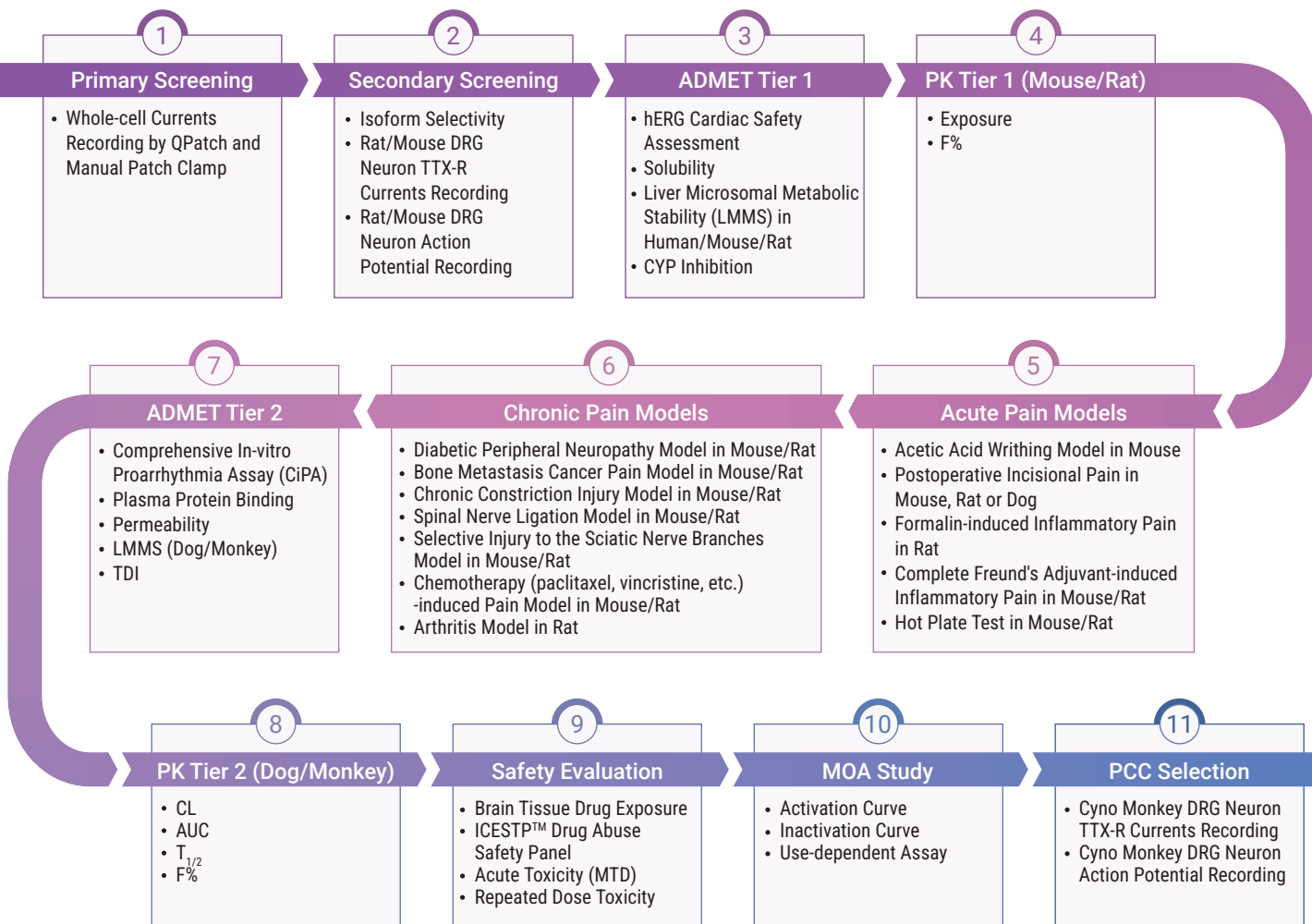
- (A) Representative traces of TTX-resistant (TTX-R) whole-cell currents recorded from cyno monkey DRG neurons by a 50 ms step of -10 mV from the holding potential of -80 mV, before (Control) and after the application of various concentrations of VX-548, as indicated.
- (B) The time-dependent changes in peak currents in (A) upon exposure to the indicated concentrations of VX-548.
- (C) Concentration-response relationship of VX-548 on TTX-R currents in human Nav1.8-CHO stable cell line, as well as in DRG neurons isolated from cynomolgus, rat, and mouse.

## Ion Channel Assay List

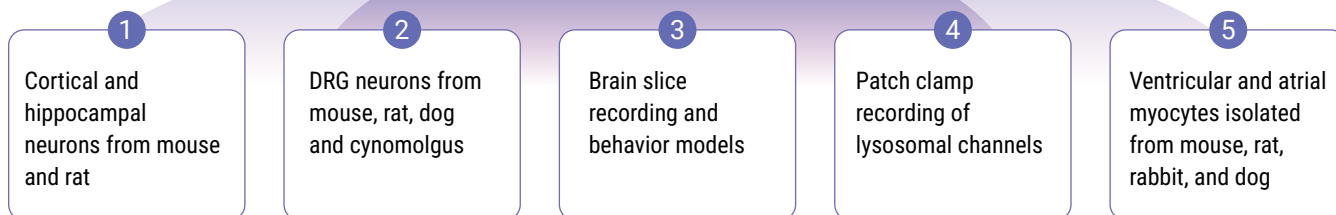
Ion Channel Family	Subtype	Reference Agonist/Activator	Reference Antagonist/Inhibitor
Sodium Channel	Nav1.1	--	TTX
	Nav1.2	--	TTX
	Nav1.3	--	TTX
	Nav1.4	--	TTX
	Nav1.5	--	TTX / Lidocaine
	Nav1.6	--	TTX
	Nav1.7	--	TTX / PF-05089771
	rNav1.7	--	TTX
	Nav1.8	--	A-803467 / VX-548 / PF-01247324
	rNav1.8	--	A-803467 / VX-548 / PF-01247324
Potassium Channel	INa-late	ATXII	Ranolazine
	hERG (IKr, Kv11.1)	--	Cisapride / E-4031 / Terfenadine
	IKs (KCNQ1/KCNE1)	--	Chromanol 293B
	Kv1.1	--	4-AP
	Kv1.2	--	4-AP
	Kv1.3	--	4-AP / PAP-1 / Psora-4 / Margatoxin
	rKv1.3	--	PAP-1
	mKv1.3	--	PAP-1
	dogKv1.3	--	PAP-1
	Kv1.4	--	4-AP/ Psora-4
	Kv1.5	--	4-AP/ Psora-4
	Kv1.6	--	4-AP
	Kv2.1	--	TEA-CI
	Kv3.1	--	TEA-CI
	Kv3.2	--	TEA-CI
	Kv3.4	--	TEA-CI
	Kv4.2/KChIP2.2	--	4-AP
	Kv4.3	--	4-AP
	KCNQ2 (Kv7.2)	Retigabine	--
	KCNQ3 (Kv7.3)	Retigabine	--
	KCNQ4 (Kv7.4)	Retigabine / XE1101	--
	KCNQ5 (Kv7.5)	Retigabine / XE1101	--
	KCNQ2/3 (Kv7.2/Kv7.3)	Retigabine / XE1101	--
	rKCNQ2/3 (Kv7.2/Kv7.3)	Retigabine / XE1101	--
	mKCNQ2/3 (Kv7.2/Kv7.3)	Retigabine / XE1101	--
	KCNQ2/4 (Kv7.2/Kv7.4)	Retigabine	--
	KCNQ3/5 (Kv7.3/Kv7.5)	Retigabine	--
	KCa1.1 (BKαβ1)	--	TEA-CI
	KCa1.1 (BKαβ4)	--	TEA-CI
	KCa2.1 (SK1)	--	Apamin
	KCa2.2 (SK2)	--	Apamin
	KCa2.3 (SK3)	--	Apamin
	KCa3.1 (IK1)	--	Senicapoc / TRAM-34
Kir2.1	--	BaCl <sub>2</sub>	
Kir3.1/3.4 (IKAch)	--	BaCl <sub>2</sub>	
Kir3.2	--	BaCl <sub>2</sub>	
Kir4.1	--	BaCl <sub>2</sub>	
Kir6.2/SUR2A (KATP)	Pinacidil	Glibenclamide	
Kir6.2/SUR1 (KATP)	Pinacidil	Glibenclamide	
TREK1 (K2P2.1)	--	Quinidine	
TASK2 (KCNK5)	--	Quinidine	
Calcium Channel	Cav1.2	--	Nifedipine / Verapamil
	Cav1.3	--	--
	Cav2.1	--	CdCl <sub>2</sub>
	Cav2.2	--	CdCl <sub>2</sub> / Cav2.2 blocker 1
	Cav3.2	--	NiCl <sub>2</sub>
TRP Channel	TRPA1	Cinnamaldehyde / AITC	Ruthenium Red / A-967079
	rTRPA1	AITC	A-967079
	TRPV1	Capsaicine	Capsazepine
	TRPV3	2-APB	Forsythoside B

Ion Channel Family	Subtype	Reference Agonist/Activator	Reference Antagonist/Inhibitor
TRP Channel	TRPV4	GSK1016790A	GSK2193874
	TRPC4	Rosiglitazone	ML204
	TRPC5-WT	Rosiglitazone	ML204 / GFB887
	TRPC5-T478C	Rosiglitazone	ML204
	TRPC6	OAG / Carbachol	SAR7334
	TRPM3	CIM-0216	Isosakuranrtin
	TRPM4/SUR1	A23187	9-Phenanthrol
	TRPM8	Menthol	2-APB
	TRPML1-ΔNC	ML-SA1	ML-SI3
	TRPML1 <sup>L15/16A L577/578A</sup>	--	--
Chloride Channel	CFTR	Forskolin	CFTR(inh)-172
	TMEM16A (ANO1)	--	T16A(inh)-A01
	TMEM16B (ANO2)	--	--
ASIC (Acid-Sensing Ion Channel)	ASIC1a	pH=6.0	Amiloride / Benzamil
	ASIC1b	pH=5.0	Amiloride / Benzamil
	ASIC3a	pH=5.0	Amiloride / Benzamil
HCN Channel	HCN1	--	Ivabradine
	HCN2	--	Ivabradine
	HCN4	--	Ivabradine
Lysosomal Two-Pore Channel	TPC2	TPC2-A1-N	YM-201636
P2X Receptor	P2X1	ATP / BZ-ATP	NF449
	P2X2	ATP / BZ-ATP	PPADS
	P2X3	αβ-methylene ATP	AF219 / BLU-5937 / S-600918
	rP2X3	αβ-methylene ATP	AF219 / BLU-5937
	gpP2X3	αβ-methylene ATP	AF219 / BLU-5937
	rP2X2/3	αβ-methylene ATP	AF219
	P2X2/3	αβ-methylene ATP	AF219
	gpP2X2/3	αβ-methylene ATP	AF219
	P2X4	ATP / BZ-ATP	BX430
	P2X5	BZ-ATP	PPADS
	P2X7	ATP / BZ-ATP	AZ10606120
mP2X7	--	--	
GABA <sub>A</sub> Receptor	GABA <sub>A</sub> (α1β2γ2)	GABA / Diazepam / Etomidate / Propofol	Bicuculline / PTX
	GABA <sub>A</sub> (α1β3γ2)	GABA	Bicuculline / PTX
	GABA <sub>A</sub> (α2β2γ2)	GABA / Diazepam	Bicuculline / PTX
	GABA <sub>A</sub> (α2β3γ2)	GABA	Bicuculline / PTX
	GABA <sub>A</sub> (α3β2γ2)	GABA / Diazepam	Bicuculline / PTX
	GABA <sub>A</sub> (α3β3γ2)	GABA	Bicuculline / PTX
	GABA <sub>A</sub> (α4β3γ2)	GABA	Bicuculline / PTX
	GABA <sub>A</sub> (α4β3δ)	GABA / Zuranolone (SAGE-217) /Brexanolone (SAGE-547)	Bicuculline / PTX
	GABA <sub>A</sub> (α5β3γ2)	GABA	Bicuculline / PTX / α5IA
	GABA <sub>A</sub> (α5β2γ2)	GABA / Diazepam	Bicuculline / PTX
	GABA <sub>A</sub> (α6β2γ2)	GABA	Bicuculline / PTX
	GABA <sub>A</sub> (α6β3γ2)	GABA	Bicuculline / PTX
GABAA (α6β3δ)	GABA / Brexanolone (SAGE-547)	Bicuculline / PTX	
NMDA Receptor	NR1/NR2A	L-Glu+Gly / NMDA+Gly	D-AP5 / Memantine
	rNR1/NR2A	L-Glu+Gly / NMDA+Gly	D-AP5 / Memantine
	NR1/NR2B	L-Glu+Gly / NMDA+Gly	D-AP5 / Memantine
	rNR1/NR2B	L-Glu+Gly / NMDA+Gly	Ifenprodil
	NR1/NR2C	L-Glu+Gly / NMDA+Gly	D-AP5 / Memantine
	NR1/NR2D	L-Glu+Gly / NMDA+Gly	D-AP5 / Memantine
AMPA Receptor	GluA1-TARPγ2	--	--
	GluA1-TARPγ8	L-Glu	NBQX
Glycine Receptor	GlyR α1β	Glycine	Strychnine
nACh Receptor	nAChR α1β1δε	Acetylcholine (ACh)	Adiphenine
	nAChR α3β4	Acetylcholine (ACh)	MLA
	nAChR α4β2	Acetylcholine (ACh)	DHβE
	nAChR α7/RIC3	Acetylcholine (ACh)	MLA
5-HT3 Receptor	5-HT3A	Serotonin (5-HT)	Ondansetron

## Comprehensive Study Workflow for Nav1.8 Modulator Development



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