

### Cytoskeleton

The cytoskeleton is a filamentous network of F-actin, microtubules, and intermediate filaments (IFs) composed of one of three chemically distinct subunits, actin, tubulin, or one of several classes of IF protein. Cytoskeleton not only helps cells maintain their shape and internal organization, but also provides mechanical support that enables cells to carry out essential functions like division and movement.

The cytoskeleton is involved in intracellular signal transduction at least two ways. First, individual proteins of the cytoskeleton might participate directly in signal transduction by linking two or more signaling proteins. Second, the cytoskeleton might provide a macromolecular scaffold, which spatially organizes components of a signal transduction cascade. Cell migration is a complex and multistep process involved in homeostasis maintenance, morphogenesis, and disease development, such as cancer metastasis, and requires coordination of cytoskeletal dynamics and reorganization, cell adhesion, and signal transduction, and takes a variety of forms. Many signaling pathways including Rho-family GTPases, Paxillin/FAK signaling and PI3K signaling is involved in the process by regulating cytoskeletal activity.

Since the cytoskeleton is involved in virtually all cellular processes, abnormalities in this essential cellular component frequently result in disease. Drugs that modulate microtubule stability, inhibitors of posttranslational modifications of cytoskeletal components, specifically compounds affecting the levels of tubulin acetylation, and compounds targeting signaling molecules which regulate cytoskeleton dynamics, constitute the mostly addressed therapeutic interventions for the diseases including cancer and neurodegenerative disorders.

#### References:

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### Arp2/3 Complex

Actin-related protein 2/3 complex

The Arp2/3 complex is originally identified in Acanthamoeba and consists of seven proteins (actin-related proteins; Arp2 and Arp3, and Arp2/3 complex subunits; ARPC1-5) that are conserved in all eukaryotes, with the exception of some algae, microsporidia and protists. The complex plays an essential role in a wide variety of cellular processes including lamellipodia-mediated cell migration, endocytosis and phagocytosis, by virtue of its ability to generate branched actin filament networks

Activation of Arp2/3 requires interaction with actin nucleation-promoting factors (NPFs). Regulation of Arp2/3 activity is achieved by endogenous inhibitory proteins through direct binding to Arp2/3 and competition with NPFs or by binding to Arp2/3-induced actin filaments and disassembly of branched actin networks. Arp2/3 inhibition has recently garnered more attention as it has been associated with attenuation of cancer progression, neurotoxic effects during drug abuse, and pathogen invasion of host cells

### Arp2/3 Complex Inhibitors, Activators & Chemicals

| · · · · ·   |  |  |   |
|---|--|--|---|
| 187-1, N-WASP inhibitor   | <b>Cat. No.</b> : HY-P1045                           | 187-1, N-WASP inhibitor TFA  | <b>Cat. No.</b> : HY-P1045A                                   |
| 187-1, N-WASP inhibitor, a 14-aa cyclic peptide,<br>is an allosteric neural Wiskott-Aldrich syndrome<br>protein (N-WASP) inhibitor. 187-1, N-WASP<br>inhibitor potently inhibits actin assembly induced<br>by phosphatidylinositol 4,5-bisphosphate (PIP2)<br>with an IC <sub>50</sub> of 2 $\mu$ M.Purity:>98%Clinical Data:No Development Reported<br>Size:1mg, 5 mg  | Opstolik-49 Photol-69 Photol-69 Photol-69 Photol-002 | 187-1, N-WASP inhibitor TFA, a 14-aa cyclic peptide, is an allosteric neural Wiskott-Aldrich syndrome protein (N-WASP) inhibitor.     Purity:   >98%     Clinical Data:   No Development Reported     Size:   1 mg, 5 mg   | Cynlofe (18 Pine) (19 Pine) (19 Pine) (1-30 Pine) (20 (19 A w |
| Benproperine phosphate  | <b>Cat. No.:</b> HY-114657A                          | СК-636<br>(СК-0944636)   | <b>Cat. No.:</b> HY-15892                                     |
| Benproperine phosphate is an orally active, potentactin-related protein 2/3 complex subunit 2(ARPC2) inhibitor. Benproperine phosphateattenuates the actin polymerization rate of actionpolymerization nucleation by impairing Arp2/3function.Purity:99.23%Clinical Data:LaunchedSize:10 mM × 1 mL, 100 mg  | N<br>HO-P-OH<br>OH                                   | CK-636 is a cell permeable inhibitor of Arp2/3<br>complex, that could inhibit actin<br>polymerization, with $IC_{50}$ values of 4 $\mu$ M, 24 $\mu$ M<br>and 32 $\mu$ M for human, fission yeast and bovine,<br>respectively.Purity:98.43%<br>Clinical Data:No Development Reported<br>Size:10 mM × 1 mL, 10 mg, 50 mg, 100 mg |   |
| CK-666  | <b>Cat. No</b> .: HY-16926                           | СК-869   | <b>Cat. No.:</b> HY-16927                                     |
| CK-666 is a cell-permeable actin-related protein<br><b>Arp2/3 complex</b> inhibitor ( $IC_{50}$ =12 µM). CK-666<br>binds to Arp2/3 complex, stabilizes the inactive<br>state of the complex, blocking movement of the<br>Arp2 and Arp3 subunits into the activated<br>filament-like (short pitch) conformation.<br><b>Purity</b> : 99.79%<br><b>Clinical Data:</b> No Development Reported<br><b>Size:</b> 10 mM × 1 mL, 5 mg, 10 mg, 50 mg |  | CK-869 is an Actin-Related Protein 2/3 (ARP2/3) complex inhibitor, with an IC <sub>50</sub> of 7 μM.     Purity:   99.74%     Clinical Data:   No Development Reported     Size:   10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg,  | Br S S  |
| Cytochalasin B<br>(Phomin)  | <b>Cat. No</b> .: HY-16928                           | Cytochalasin D<br>(Zygosporin A; NSC 209835)   | <b>Cat. No.</b> : HY-N6682                                    |
| Cytochalasin B is a cell-permeable mycotoxin binding to the barbed end of <b>actin</b> filaments, disrupting the formation of actin polymers, with $K_d$ value of 1.4-2.2 nM for F-actin.   |  | Cytochalasin D (Zygosporin A; NSC 209835) is a<br>potent and cell-permeable inhibitor of actin<br>polymerization derived from fungus, inhibits the<br>G-actin–cofilin interaction by binding to G-actin.   | HN CON  |
| Purity:99.84%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 1 mg, 5 mg, 10 mg  | 0  | Purity:99.75%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 1 mg, 5 mg, 10 mg   | ő   |
| Dihydrocytochalasin B   | <b>Cat. No.:</b> HY-N6701                            | Jasplakinolide   | Cat. No.: HY-P0027  |
| Dihydrocytochalasin B (H2CB) is a <b>Cytokinesis</b><br>inhibitor and changes the morphology of the cells,<br>similar to that of cytochalasin B; does not<br>inhibit glucose transport.   |  | Jasplakinolide is a potent <b>actin polymerization</b><br>inducer and stabilizes pre-existing actin<br>filaments. Jasplakinolide binds to F-actin<br>competitively with phalloidin with a K <sub>d</sub> of 15<br>nM.  | Br H H OH   |
| Purity: >98%<br>Clinical Data: No Development Reported<br>Size: 1 mg, 5 mg  | 2000 - 2000  | Purity: ≥98.0%   Clinical Data: No Development Reported   Size: 100 µg   |   |

| Latrunculin A<br>(LAT-A)  | <b>Cat. No.:</b> HY-16929 | Phalloidin-TRITC  | <b>Cat. No.:</b> HY-P2270                             |
|---|---------------------------|---|---|
| Latrunculin A (LAT-A) is a toxin isolated from the<br>red sea sponge Latrunculia magnifica, binds to<br>actin monomers, inhibits polymerization of actin,<br>with K <sub>d</sub> s of 0.1, 0.4, 4.7 $\mu$ M and 0.19 $\mu$ M for<br>ATP-actin, ADP-Pi-actin, ADP-actin and G-actin,<br>respectively.<br>Purity: $\geq$ 97.0%<br>Clinical Data: No Development Reported<br>Size: 100 $\mu$ g (237.2 $\mu$ M * 1 mL in Ethanol) |                           | Phalloidin-TRITC is a TRITC labeled, red     fluorescence probe for F-actin. Phalloidin, bound     to actin filaments, reacts covalently with amino     acids Glu-IIT, Met-II9, and Met355, which are very     close to the nucleotide binding site.     Purity:   >98%     Clinical Data:   No Development Reported     Size:   1 mg, 5 mg | a ya Terj G (Haq (Her) (Terj j Dunier i Waya Cyra Tyd |
| SMIFH2  |                           | Wiskostatin   |   |
|   | Cat. No.: HY-16931        |   | Cat. No.: HY-12534                                    |
| SMIFH2 is a <b>formin</b> specific inhibitor. SMIFH2 inhibits actin polymerization by Formins and affects the actin cytoskeleton.   | OF N S Br                 | Wiskostatin is a potent and selective inhibitor of<br>neuronal Wiskott-Aldrich syndrome protein<br>(N-WASP)-mediated actin polymerization.<br>Wiskostatin causes a rapid, profound, and<br>irreversible decrease in cellular ATP levels.  | N<br>OH   |
| Purity:98.22%Clinical Data:No Development ReportedSize:5 mg, 10 mg, 50 mg, 100 mg   | 0.00                      | Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg   | Br  |



## Dynamin

Dynamins are large superfamily GTPase proteins that are involved in various cellular processes including budding of transport vesicles, division of organelles, cytokinesis, and pathogen resistance. Dynamins are involved in scission (cleavage of the vesicle from the parent membrane) of nascent vesicles from parent membranes in eukaryotic cells. Dynamins interact directly with the lipid bilayer at the necks of clathrin-coated pits to sever and release coated vesicles. Dynamins contain five domains, including GTPase domain, middle domain, PH domain, GTPase effector domain (GED), and proline rich domain (PRD), while the dynamin-related proteins (DRPs) lack one or more of these domains or have additional domains. Dynamins and DRPs participate in a wide variety of cellular processes, including budding mitochondrial fission (mammalian Dlp1 and Saccharomyces cerevisiae Dnm1) and fusion (mammalian OPA1, S.cerevisiae Mgm1 and Schizosaccharomycespombe Msp1), vacuolar fission (S. cerevisiae Vps1), interferon-induced anti-viral protection (fish Mx proteins), plant cell cytokinesis and membrane fission (Arabidopsis thalianaDRP proteins), as well as pathogen resistance.

### **Dynamin Inhibitors**

| Drp1-IN-1   | Cat. No.: HY-125222  | Dynamin inhibitory peptide   | Cat. No.: HY-P1083          |
|---|--|--|-----------------------------|
| Drp1-IN-1 (comp A-7) is a <b>dynamin-1-like</b><br><b>protein (Drp1)</b> inhibitor, with an $IC_{s0}$ of 0.91<br>$\mu$ M.   | A REAL   | Dynamin inhibitory peptide competitively blocks<br>binding of <b>dynamin</b> to amphiphysin, thus<br>preventing endocytosis. Dynamin inhibitory peptide<br>blocks the dopamine D3 effect on GABAA receptors.   |                             |
| Purity:>98%Clinical Data:No Development ReportedSize:5 mg, 10 mg  | N Ö<br>N-NH  | Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  | in a for the form           |
| Dynamin inhibitory peptide TFA  | <b>Cat. No.:</b> HY-P1083A   | DynaMin inhibitory peptide, myristoylated  | <b>Cat. No.</b> : HY-P1369  |
| Dynamin inhibitory peptide TFA competitively<br>blocks binding of <b>dynamin</b> to amphiphysin, thus<br>preventing endocytosis. Dynamin inhibitory peptide<br>TFA blocks the dopamine $D_3$ effect on GABA <sub>A</sub><br>receptors.  | Contraction of the second seco | DynaMin inhibitory peptide, myristoylated is a <b>DynaMin</b> inhibitor to interfere with the binding of amphiphysin with dynamin. DynaMin inhibitory peptide, myristoylated is a membrane-permeant form of the peptide that prevents endocytosis.   | Myristoyl-QVPSRPNRAP-NH     |
| Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg   | HAZTON FLOR  | Purity:>98%Clinical Data:No Development ReportedSize:5 mg, 10 mg, 25 mg  |                             |
| DynaMin inhibitory peptide, myristoylated TFA   | <b>Cat. No.:</b> HY-P1369A   | Dynasore   | <b>Cat. No.</b> : HY-15304  |
| DynaMin inhibitory peptide, myristoylated TFA is a <b>DynaMin</b> inhibitor to interfere with the binding of amphiphysin with dynamin. DynaMin inhibitory peptide, myristoylated TFA is a membrane-permeant form of the peptide that prevents endocytosis.  | Myrstoy-OVPSRPNRAP-NH2 (TFA sat)   | Dynasore is a cell-permeable $dynamin$ inhibitor with an $IC_{so}$ of 15 $\mu M.$  | CUC OH No CO                |
| Purity: > 98%   Clinical Data: No Development Reported   Size: 1 mg, 5 mg   |  | Purity:98.70%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 10 mg, 50 mg  | 9623.                       |
| Dynole 2–24   | <b>Cat. No.</b> : HY-145080  | Dynole 34-2  | <b>Cat. No</b> .: HY-107545 |
| Dynole 2–24 is an indole-based <b>dynamin GTPase</b><br>inhibitor (IC <sub>s0</sub> =0.56 $\mu$ M for dynamin I). Dynole<br>2–24 is nontoxic and shows increased potency<br>against dynamin I and II in vitro and in cells<br>(IC <sub>s(CME)</sub> =1.9 $\mu$ M). Dynole 2–24 also shows<br>4.4-fold selectivity for dynamin I.<br><b>Purity:</b> >98%<br>Clinical Data: No Development Reported<br>Size: 1 mg, 5 mg | tut5   | Dynole 34-2 is a <b>dynamin GTPase</b> inhibitor<br>(IC <sub>so</sub> s=6.9 and 14.2 µM for dynamin1 and dynamin2<br>GTPase activity, respectively) with antimitotic<br>effect. Dynole 34-2 induces apoptosis, as revealed<br>by cell blebbing, DNA fragmentation, and PARP<br>cleavage.<br>Purity: >98%<br>Clinical Data: No Development Reported<br>Size: 1 mg, 5 mg |                             |
| Hydroxy-Dynasore<br>(Dyngo-4a)  | <b>Cat. No.:</b> HY-13863  | Mdivi-1<br>(Mitochondrial division inhibitor 1)  | <b>Cat. No.:</b> HY-15886   |
| Hydroxy Dynasore (Dyngo-4a), a structural analog<br>of Dynasore (HY-15304), is an potency improved,<br>low cytotoxicity and nonspecific binding<br><b>dynamin</b> inhibitor with $IC_{so}$ values of 0.38<br>$\mu$ M and 2.3 $\mu$ M for brain dynamin I and<br>recombinant rat dynamin II, respectively.   | CTC+CH N OH  | Mdivi-1 is a selective dynamin-related protein 1<br>( <b>Drp1</b> ) inhibitor. Mdivi-1 is a mitochondrial<br>division/ <b>mitophagy</b> inhibitor.   |                             |
| Purity:     98.08%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 10 mg, 50 mg, 100 mg  |  | Purity:99.73%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg  | эл                          |

| Cat. No.: HY-N0703                      |
|---|
| HO + |
|   |



# **Gap Junction Protein**

Gap junction (GJ) channels span the plasma membranes of adjacent cells and are formed by the docking of two hemichannels (connexons) oligomerized from connexin (Cx) proteins, which consist of 21 distinct isoforms. GJs provide a direct pathway for cell-to-cell electrical signaling and metabolic communication, allowing the passage of small ions, amino acids, metabolites, tetraethylammonium and signaling molecules such as cAMP, IP3, siRNA and small peptide.

Gap junction channels provide the basis for intercellular communication in the cardiovascular system for maintenance of the normal cardiac rhythm, regulation of vascular tone and endothelial function as well as metabolic interchange between the cells. In the heart, GJs mediate electrical coupling between cardiac myocytes, forming the cell-to-cell pathways for orderly spread of the wave of electrical excitation responsible for synchronous contraction. Gap junctions also play an important role in the control of bladder contractile response and in the regulation of various immune inflammatory processes.

### Gap Junction Protein Inhibitors & Modulators

| AT-1002   |  | AT-1002 TFA  |  |
|---|--|--|--|
|   | Cat. No.: HY-114426  |  | Cat. No.: HY-114426A   |
| AT-1002, a 6-mer synthetic peptide, is a tight <b>junction</b> regulator and absorption enhancer.   | white the first of | AT-1002 TFA, a 6-mer synthetic peptide, is a tight <b>junction</b> regulator and absorption enhancer.  | <sup>wyłt</sup><br>S <sup>1</sup> f <sup>2</sup> f <sup>2</sup> f <sup>2</sup> f <sup>2</sup> f <sup>2</sup><br>w <sup>w</sup> w   |
| Purity: >98%   Clinical Data: No Development Reported   Size: 1 mg, 5 mg  |  | Purity:     99.72%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg   | <b>F</b> (0)   |
| Carbenoxolone disodium  | <b>Cat. No.:</b> HY-B1367  | Danegaptide<br>(GAP-134; ZP 1609)  | <b>Cat. No.:</b> HY-10913  |
| Carbenoxolone disodium is the active <b>metabolite</b> of Glycyrrhizic acid (HY-N0184) and the inhibitor of human $11\beta$ -HSD and bacterial $3\alpha$ , $20\beta$ -HSD. Carbenoxolone disodium is an uncoupling agent for <b>gap junctions</b> and a potent inhibitor of Vaccinia virus replication. | NUC - C - C - C - C - C - C - C - C - C -  | Danegaptide (GAP-134) is a potent, selective and orally active <b>gap-junction</b> modifier with an antiarrhythmic effect.   |  |
| Purity:     99.88%       Clinical Data:     Launched       Size:     10 mM × 1 mL, 25 mg, 50 mg, 100 mg   |  | Purity:>98%Clinical Data:Phase 2Size:1 mg, 5 mg  |  |
| Danegaptide Hydrochloride<br>(GAP-134 Hydrochloride; ZP 1609 Hydrochloride)   | <b>Cat. No.</b> : HY-10913A  | Gap 26   | <b>Cat. No.:</b> HY-P1082  |
| Danegaptide Hydrochloride (GAP-134 Hydrochloride)<br>is a potent, selective and orally active<br>gap-junction modifier with an antiarrhythmic<br>effect.  |  | Gap 26 is a connexin mimetic peptide, composed of<br>residue numbers 63-75 of the first extracellular<br>loop of connexin 43 (gap junction blocker),<br>containing the SHVR amino acid motif.  | VCYDKSFPISHVR  |
| Purity:     99.75%       Clinical Data:     Phase 2       Size:     10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 1  | н-сі<br>.00 mg   | Purity:99.64%Clinical Data:No Development ReportedSize:1 mg, 5 mg, 10 mg   |  |
| Gap 26 TFA  | <b>Cat. No.:</b> HY-P1082A   | Gap 27   | <b>Cat. No.:</b> HY-P0139  |
| Gap 26 TFA is a connexin mimetic peptide, composed<br>of residue numbers 63-75 of the first<br>extracellular loop of connexin 43 (gap junction<br>blocker), containing the SHVR amino acid motif.   | VCYDKSFPISHVR (TFA Sali)   | Gap 27, a synthetic connexin43 mimetic peptide, is<br>a gap junction inhibitor. Gap 27 possesses<br>conserved sequence homology to a portion of the<br>second extracellular loop leading into the fourth<br>transmembrane connexin segment.  | SRPTEKTIFII  |
| Purity:99.03%Clinical Data:No Development ReportedSize:1 mg, 5 mg, 10 mg  |  | Purity:98.07%Clinical Data:No Development ReportedSize:1 mg, 5 mg  |  |
| Gap19   | <b>Cat. No.</b> : HY-P1136   | Gap19 TFA  | <b>Cat. No.:</b> HY-P1136A   |
| Gap19, a peptide derived from nine amino acids of<br>the Cx43 cytoplasmic loop (CL), is a potent and<br>selective <b>connexin 43 (Cx43) hemichannel</b><br>blocker. Gap19 inhibits hemichannels caused by<br>preventing intramolecular interactions of the<br>C-terminus (CT) with the CL.              |  | Gap19 TFA, a peptide derived from nine amino acids<br>of the Cx43 cytoplasmic loop (CL), is a potent and<br>selective <b>connexin 43 (Cx43) hemichannel</b><br>blocker. Gap19 TFA inhibits hemichannels caused by<br>preventing intramolecular interactions of the<br>C-terminus (CT) with the CL. | and the second s |
| Purity: >98%   Clinical Data: No Development Reported   Size: 1 mg, 5 mg  |  | Purity:95.11%Clinical Data:No Development ReportedSize:1 mg, 5 mg, 10 mg   |  |

| Larazotide acetate   | Cat. No.: HY-106268A        | Meclofenamic acid<br>(Meclofenamate)   | Cat. No.: HY-117275   |
|--|-----------------------------|--|---|
| Larazotide acetate is a synthetic peptide.<br>Larazotide acetate acts as a tight <b>junction</b><br>regulator and reverses leaky junctions to their<br>normally closed state.  |                             | Meclofenamic Acid (Meclofenamate), a<br>non-steroidal, anti-inflammatory agent, is a<br>highly selective <b>fat mass and obesity-associated</b><br>( <b>FTO</b> ) <b>enzyme</b> inhibitor. Meclofenamic Acid<br>competes with FTO binding for the m(6)A-containing<br>nucleic acid.  | CI NOCOH  |
| Purity:     99.68%       Clinical Data:     Phase 3       Size:     10 mM × 1 mL, 1 mg, 5 mg, 10 mg, 25 mg   |                             | Purity:>98%Clinical Data:LaunchedSize:1 mg, 5 mg   |   |
| Meclofenamic acid sodium<br>(Meclofenamate sodium)   | <b>Cat. No.</b> : HY-B1320  | Meclofenamic acid-d4<br>(Meclofenamate-d4)   | <b>Cat. No.</b> : HY-117275S  |
| Meclofenamic acid (Meclofenamate) sodium is a<br>nonsteroidal anti-inflammatory drug (NSAID)<br>approved for use in arthritis (osteo and<br>rheumatoid), analgesia (mild to moderate pain),<br>dysmenorrhea, and heavy menstrual blood loss<br>(menorrhagia).     Purity:   99.86%     Clinical Data:   Launched     Size:   10 mM × 1 mL, 50 mg, 100 mg, 200 mg |                             | Meclofenamic acid-d4 (Meclofenamate-d4) is the deuterium labeled Meclofenamic acid. Meclofenamic Acid (Meclofenamate), a non-steroidal, anti-inflammatory agent, is a highly selective fat mass and obesity-associated (FTO) enzyme inhibitor.     Purity:   >98%     Clinical Data:   No Development Reported     Size:   1 mg, 10 mg |   |
| Peptide5   | <b>Cat. No.:</b> HY-P2275   | Rotigaptide<br>(ZP123)   | <b>Cat. No.</b> : HY-106225   |
| Peptide5, a connexin 43 mimetic peptide, reduce<br>animals swelling, astrogliosis, and neuronal cell<br>death after spinal cord injury.  | alenler y                   | Rotigaptide (ZP123) is a novel and specific<br>modulator of <b>connexin 43 (Cx43</b> ). Rotigaptide<br>prevents the uncoupling of Cx43-mediated <b>gap</b><br><b>junction</b> communication and normalizes cell-to-cell<br>communication during acute metabolic stress.  |   |
| Purity: >98%   Clinical Data: No Development Reported   Size: 1 mg, 5 mg   |                             | Purity:99.63%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 1 mg, 5 mg  | HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>HQ<br>H |
| TAT-Gap19  | <b>Cat. No.</b> : HY-P1136B | TAT-Gap19 TFA  | <b>Cat. No.:</b> HY-P1136C  |
| TAT-Gap19, a Cx mimetic peptide, is a specific<br>connexin43 hemichannel (Cx43 HC) inhibitor.<br>TAT-Gap19 does not inhibits the corresponding Cx43<br>GJCs. TAT-Gap19 traverses the blood-brain barrier<br>and alleviate liver fibrosis in mice.  | YGRKKRRQRRKQIEIKKFK         | TAT-Gap19 TFA, a Cx mimetic peptide, is a specific <b>connexin43 hemichannel (Cx43 HC)</b> inhibitor.<br>TAT-Gap19 TFA does not inhibits the corresponding Cx43 GJCs. TAT-Gap19 TFA traverses the blood-brain barrier and alleviate liver fibrosis in mice.  | YGRIKKRIGRREKCIEIKKFK (TFA sati)  |
| Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  |                             | Purity:98.36%Clinical Data:No Development ReportedSize:5 mg, 10 mg   |   |
| Tonabersat<br>(SB-220453)  | <b>Cat. No</b> .: HY-15204  |  |   |
| Tonabersat (SB-220453) is a <b>gap-junction</b> modulator.<br>Tonabersat prevents inflammatory damage in the<br>central nervous system.  | U HN CO                     |  |   |
| Purity:     98.36%       Clinical Data:     Phase 2       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg   | F CI                        |  |   |



## Integrin

Integrins, a family of heterodimeric adhesion receptors for diverse extracellular matrices, have consistently been implicated as crucial drivers of ovarian cancer development and progression. A number of the RGD-based members of the integrin family, including  $\alpha 5\beta 1$ , and  $\alpha \nu \beta 3$  or  $\alpha \nu \beta 5$  integrins, are markedly elevated in aggressive ovarian tumors. These adhesion receptors appear to promote cell adhesion, survival, motility and invasion during ovarian tumor growth or metastatic progression. Importantly, the functions of these integrins are strongly dependent on the activation of focal adhesion kinase (FAK) and its downstream signaling, including the PI3K/Akt- and Ras/MAPK-dependent pathways.

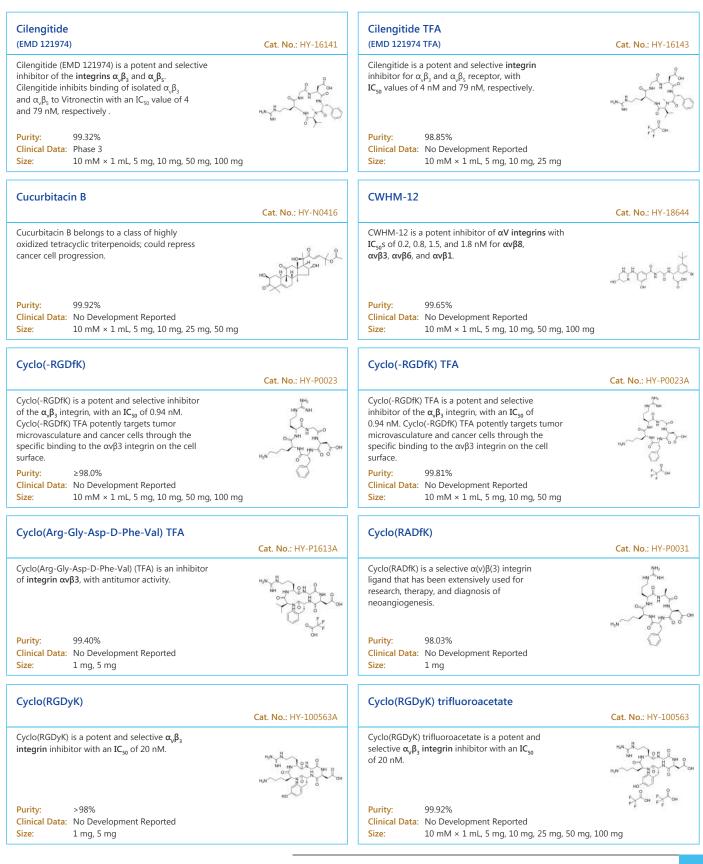
Integrins are transmembrane proteins and are major receptors for cell-extracellular matrix (ECM) and cell-cell adhesion. Modulation of these molecules, particularly  $\alpha v$  integrin family, has exhibited profound effects on fibrosis in multiple organ and disease state. Based on the several studies, the integrins  $\alpha v\beta 3$ ,  $\alpha v\beta 5$ ,  $\alpha v\beta 6$ , and  $\alpha v\beta 8$  have been known to modulate the fibrotic process via activation of latent transforming growth factor (TGF)- $\beta$  in pre-clinical models of fibrosis.

Each integrin is typically formed by the non-covalent pairing of one  $\alpha$  subunit, of which, 18 types are known to exist, and one  $\beta$  subunit, of which 8 types are known to exist. Together, 24 distinct heterodimers have been identified to date. The  $\alpha$ v subunit can form heterodimers with the  $\beta$ 1,  $\beta$ 3,  $\beta$ 5,  $\beta$ 6 or  $\beta$ 8 subunits and  $\beta$ 1 can associate with many different  $\alpha$  subunits from  $\alpha$ 1 to  $\alpha$ 11, and  $\alpha$ v, indicating that not all theoretically possible  $\alpha$  and subunit pairs form. Interestingly, the activation of TGF- $\beta$  appears to be a common function of multiple  $\alpha$ v integrins.

### Integrin Inhibitors, Agonists, Antagonists & Modulators

| A-205804  |  | A-286982   |                           |
|---|--|--|---------------------------|
|   | Cat. No.: HY-100226  |  | Cat. No.: HY-107587       |
| A-205804 is an orally bioavailable, potent and selective lead inhibitor of E-selectin and ICAM-1 expression, with an IC <sub>50</sub> of 20 nM and 25 nM for E-selectin and ICAM-1, respectively.<br>A-205804 can be used in the research of chronic inflammatory diseases. | N S S NH2<br>S S   | A-286982 is a potent and allosteric<br><b>LFA-1/ICAM-1 interaction</b> inhibitor with <b>IC<sub>50</sub>s</b> of<br>44 nM and 35 nM in an LFA-1/ICAM-1 binding and<br>LFA-1-mediated cellular adhesion assay,<br>respectively.   |                           |
| Purity:98.12%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 10 mg, 50 mg   |  | Purity:99.69%Clinical Data:No Development ReportedSize:5 mg, 10 mg, 25 mg, 50 mg   |                           |
| Alicaforsen<br>(ISIS-2302)  | <b>Cat. No.:</b> HY-145728   | Arg-Gly-Asp-Ser<br>(RGDS peptide; Fibronectin tetrapeptide)  | <b>Cat. No.:</b> HY-12290 |
| Alicaforsen is a 20-base antisense<br>oligonucleotide inhibiting <b>ICAM-1</b> production,<br>which is an important adhesion molecule involved<br>in leukocyte migration and trafficking to the site<br>of inflammation.  | Alicaforsen  | Arg-Gly-Asp-Ser is an integrin binding sequence<br>that inhibits <b>integrin receptor</b> function.<br>Arg-Gly-Asp-Ser directly and specifically bind<br>pro-caspase-8, pro-caspase-9 and pro-caspase-3,<br>while it does not bind pro-caspase-1.                              |                           |
| Purity:>98%Clinical Data:Phase 3Size:1 mg, 5 mg   |  | Purity:99.76%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 10 mg, 50 mg, 100 mg  |                           |
| Arg-Gly-Asp-Ser (TFA)<br>(RGDS peptide (TFA); Fibronectin tetrapeptide (TFA))   | <b>Cat. No.:</b> HY-12290A   | ATN-161  | <b>Cat. No.:</b> HY-13535 |
| Arg-Gly-Asp-Ser (TFA) is an integrin binding<br>sequence that inhibits <b>integrin receptor</b> function.<br>Arg-Gly-Asp-Ser (TFA) directly and specifically<br>bind pro-caspase-8, pro-caspase-9 and<br>pro-caspase-3, while it does not bind<br>pro-caspase-1.            | но <sup>2</sup> Н, о <sup>0</sup> 2 Ц, <sup>Мо</sup> 4 , ил,<br>но С <sub>0</sub> Ц, Ц, <sup>Мо</sup> 4 , ил,<br>но С <sub>0</sub> Ц, <sup>1</sup> , <sup></sup> | ATN-161 is a novel integrin $\alpha 5\beta 1$ antagonist, which inhibits angiogenesis and growth of liver metastases in a murine model.  |                           |
| Purity: >98%<br>Clinical Data:<br>Size: 1 mg, 5 mg  | С.   | Purity:>98%Clinical Data:Phase 2Size:1 mg, 5 mg  |                           |
| ATN-161 trifluoroacetate salt   |  | Bexotegrast  |                           |
| (ATN-161 TFA salt)  | Cat. No.: HY-13535A  |  | Cat. No.: HY-137561       |
| ATN-161 trifluoroacetate salt is a novel <b>integrin</b> $\alpha 5\beta 1$ antagonist, which inhibits angiogenesis and growth of liver metastases in a murine model.  |  | Bexotegrast is a potent inhibitor of $\alpha\nu\beta6$<br>integrin. Bexotegrast can be used for researching<br>fibrosis such as idiopathic pulmonary fibrosis<br>(IPF) and nonspecific interstitial pneumonia<br>(NSIP) (extracted from patent WO2020210404A1,<br>compound 5). |                           |
| Purity:     ≥95.0%       Clinical Data:     Phase 2       Size:     5 mg, 10 mg, 50 mg, 100 mg  | ~~   | Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  |                           |
| BI-1950   | <b>Cat. No.:</b> HY-124040   | BIO-1211   | <b>Cat. No.:</b> HY-14126 |
| BI-1950 is a highly potent <b>lymphocyte function</b><br><b>associated antigen-1 (LFA-1)</b> inhibitor. LFA-1 is<br>an essential component in normal immune system<br>function and a target for drug discovery.   |  | BIO-1211 is a highly selective and orally active $\alpha 4\beta 1$ (VLA-4) inhibitor, with IC <sub>so</sub> values of 4 nM and 2 $\mu$ M for $\alpha 4\beta 1$ and $\alpha 4\beta 7$ , respectively.   | ؿؿؿؙڣ۬ڹ <sub></sub> ڹؠؿ   |
| Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg   | à  | Purity:99.64%Clinical Data:No Development ReportedSize:1 mg  |                           |

#### **BIO5192** BIO5192 hydrate Cat. No.: HY-107589 Cat. No.: HY-107589A BIO5192 is a selective and potent integrin $\alpha 4\beta 1$ BIO5192 hydrate is a selective and potent integrin (VLA-4) inhibitor (K<sub>4</sub><10 pM). BIO5192 α4β1 (VLA-4) inhibitor (K\_<10 pM). BIO5192 selectively binds to $\alpha 4\beta 1$ (IC<sub>50</sub>=1.8 nM) over a hydrate selectively binds to $\alpha 4\beta 1$ (IC<sub>50</sub>=1.8 nM) range of other integrins. over a range of other integrins. Purity: > 98% >98.0% Purity: Clinical Data: No Development Reported Clinical Data: No Development Reported Size: 5 mg, 10 mg Size: 1 mg **BIRT 377** BMS-587101 Cat. No.: HY-110117 Cat. No.: HY-120628 BIRT 377 is a potent amd orally bioavailable BMS-587101 is a potent and orally active antagonist of leukocyte function associated inhibitor of the interaction between intercellular adhesion molecule-1 (ICAM-1) and lymphocyte antigen-1 (LFA-1). BMS-587101 has anti-inflammatory effects and can be used for function-associated antigen-1 (LFA-1), with a K, of 25.8 nM. BIRT 377 also inhibits the production rheumatoid arthritis research. of IL-2 in vivo. Purity: > 98% **Purity:** 98 67% Clinical Data: No Development Reported Clinical Data: Phase 2 5 mg, 10 mg, 25 mg, 50 mg, 100 mg Size: 1 mg, 5 mg Size: BMS-688521 **BOP** sodium Cat. No.: HY-10596 Cat. No.: HY-129453 BMS-688521 is a highly potent, orally active BOP sodium is a potent and selective dual inhibitor of the LFA-1/ICAM interaction, with $\alpha 9\beta 1/\alpha 4\beta 1$ integrin inhibitor with K<sub>d</sub> values in an IC<sub>50</sub> of 2.5 nM in the adhesion assay and an the picomolar range. BOP sodium shows the rapid IC<sub>sn</sub> of 60 nM in the MLR assay. BMS-688521 is and preferential mobilization of hematopoietic efficacious in a mouse allergic eosinophilic lung stem cell (HSC) and progenitors. inflammation model. Purity: 98.72% >98% Purity: Clinical Data: No Development Reported Clinical Data: No Development Reported Size: 10 mM × 1 mL, 5 mg, 10 mg Size: 1 mg, 5 mg c(phg-isoDGR-(NMe)k) c(phg-isoDGR-(NMe)k) TFA Cat. No.: HY-111413 Cat. No.: HY-111413A c(phg-isoDGR-(NMe)k) is a selective and potent c(phg-isoDGR-(NMe)k) TFA is a selective and potent $\alpha$ 5 $\beta$ 1-integrin ligand with an IC<sub>50</sub> of 2.9 nM. $\alpha$ 5 $\beta$ 1-integrin ligand with an IC<sub>so</sub> of 2.9 nM. >98% Purity: >98% Purity: Clinical Data: No Development Reported Clinical Data: No Development Reported 1 mg, 5 mg, 10 mg Size: 1 mg, 5 mg, 10 mg Size: Carotegrast Carotegrast methyl Cat. No.: HY-14857 (AJM300) Cat. No.: HY-124290 Carotegrast is an orally available $\alpha 4$ integrin Carotegrast methyl (AJM300) is an orally active receptor inhibitor with anti-inflammatories and selective $\alpha 4$ integrin antagonist. HCA2969, an active metabolite of Carotegrast methyl, is a activities specific and dual $\alpha 4\beta 1/\alpha 4\beta 7$ integrin antagonist. Carotegrast methyl prevents the development of colitis in mice. <br/> Purity: 98.14% **Purity:** 99.72% Clinical Data: No Development Reported Clinical Data: No Development Reported Size: 10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 100 mg Size: 5 mg, 10 mg, 25 mg, 50 mg, 100 mg



| E7820  |                             | Echistatin   |  |
|--|-----------------------------|--|--|
| (ER68203-00)   | Cat. No.: HY-14571          |  | Cat. No.: HY-P1189   |
| E7820 (ER68203-00), an orally active aromatic<br>sulfonamide derivative, is a unique angiogenesis<br>inhibitor suppressing an expression of integrin<br>alpha2 subunit on endothelium. E7820 inhibits rat<br>aorta angiogenesis with an IC <sub>50</sub> of 0.11 $\mu$ g/ml.Purity:99.25%<br>Clinical Data:Phase 2<br>Size:10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg | N SHO                       | Echistatin, the smallest active RGD protein<br>belonging to the family of disintegrins that are<br>derived from snake venoms, is a potent inhibitor<br>of platelet aggregation. Echistatin is a potent<br>inhibitor of bone resorption in culture.Purity:>98%Clinical Data:No Development Reported<br>Size:Size:5 mg, 10 mg, 25 mg | FERRI DE CONTRA DE C |
| Echistatin TFA   | Cat. No.: HY-P1189A         | EMD527040  | <b>Cat. No.:</b> HY-101473   |
| Echistatin TFA, the smallest active RGD protein<br>belonging to the family of disintegrins that are<br>derived from snake venoms, is a potent inhibitor<br>of <b>platelet aggregation</b> . Echistatin is a potent<br>inhibitor of <b>bone resorption</b> in culture.<br><b>Purity:</b> 95.13%   |                             | EMD527040 is a potent and highly selective αvβ6<br>antagonist with antifibrotic activities. EMD527040<br>can be used for carcinoma and liver fibrosis<br>research.   |  |
| Clinical Data: No Development Reported<br>Size: 1 mg, 5 mg   |                             | Clinical Data: No Development Reported<br>Size: 1 mg, 5 mg   |  |
| Eptifibatide   |                             | Eptifibatide acetate   |  |
|  | Cat. No.: HY-B0686          |  | Cat. No.: HY-B0686A  |
| Eptifibatide is a cyclic heptapeptide, acts as a competitive antagonist for the activated platelet <b>glycoprotein IIb/IIIa receptor</b> , with anti-platelet activity.  |                             | Eptifibatide acetate is a cyclic heptapeptide,<br>acts as a competitive antagonist for the activated<br>platelet glycoprotein IIb/IIIa receptor, with<br>anti-platelet activity.   |  |
| Purity:     99.91%       Clinical Data:     Launched       Size:     10 mM × 1 mL, 10 mg, 50 mg, 100 mg  | 2× ae3d1                    | Purity:>98%Clinical Data:LaunchedSize:1 mg, 5 mg   | о<br>Лон   |
| Fibronectin  | <b>Cat. No.:</b> HY-P3160   | Firategrast<br>(SB 683699)   | <b>Cat. No.:</b> HY-14951  |
| Fibronectin, a glycoprotein (~500 kDa) present in<br>blood as well as in cells, is a biomarker of<br>tissue injury. Fibronectin binds to<br>membrane-spanning receptor proteins called<br>integrins.   | Fibronectins                | Firategrast (SB 683699) is an orally active and<br>specific $\alpha 4\beta 1/\alpha 4\beta 7$ integrin antagonist.<br>Firategrast reduces trafficking of lymphocytes<br>into the central nervous system (CNS) and<br>decreases multiple sclerosis (MS) activity.   | GG HLOH<br>F J LOH   |
| Purity:≥95.0%Clinical Data:No Development ReportedSize:1 mg  |                             | Purity:     99.88%       Clinical Data:     Phase 2       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg   | u v v v  |
| Fradafiban<br>(BIBU-52)  | <b>Cat. No</b> .: HY-101720 | GLPG0187   | <b>Cat. No.:</b> HY-100506   |
| Fradafiban is a nonpeptide platelet glycoprotein IIb/IIIa antagonist, which binds to the human platelet GP IIb/IIIa complex with a $K_d$ value of 148 nM.  | HO-C-S-NH O-C-S-NH          | GLPG0187 is a broad spectrum integrin receptor antagonist with antitumor activity; inhibits $\alpha_{v}\beta_{1}$ -integrin with an IC <sub>50</sub> of 1.3 nM.  | to <sup>oth</sup> sta,   |
| Purity:>98%Clinical Data:Phase 1Size:1 mg, 5 mg  |                             | Purity:     99.78%       Clinical Data:     Phase 1       Size:     10 mM × 1 mL, 1 mg, 5 mg, 10 mg, 50 mg, 100  | ) mg   |

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| Gly-Arg-Gly-Asp-Ser  | <b>Cat. No.:</b> HY-P0295                | Gly-Arg-Gly-Asp-Ser TFA  | <b>Cat. No.:</b> HY-P0295A  |
|--|--|--|-----------------------------|
| Gly-Arg-Gly-Asp-Ser is a pentapeptide that forms the cell-binding domain of a glycoprotein, osteopontin. Gly-Arg-Gly-Asp-Ser binds to integrin receptors $\alpha\nu\beta3$ and $\alpha\nu\beta5$ with estimated IC <sub>50</sub> of 5 and 6.5 $\mu$ M. |  | Gly-Arg-Gly-Asp-Ser (TFA) is a pentapeptide that forms the cell-binding domain of a glycoprotein, osteopontin. Gly-Arg-Gly-Asp-Ser binds to <b>integrin receptors</b> $\alpha\nu\beta3$ and $\alpha\nu\beta5$ with estimated <b>IC</b> <sub>so</sub> of 5 and 6.5 $\mu$ M. |                             |
| Purity:95.05%Clinical Data:No Development ReportedSize:2 mg, 5 mg, 10 mg, 25 mg  |  | Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  | P} OH                       |
| GRGDSP   | <b>Cat. No.:</b> HY-P0290                | GRGDSP TFA   | <b>Cat. No.</b> : HY-P0290A |
| GRGDSP, a synthetic linear RGD peptide, is an integrin inhibitor.  |  | GRGDSP (TFA) is an <b>integrin</b> inhibitor.  |                             |
| Purity:>98%Clinical Data:No Development ReportedSize:5 mg, 10 mg, 25 mg  | 0 0 X <sub>01</sub>                      | Purity:≥98.0%Clinical Data:No Development ReportedSize:1 mg, 5 mg, 10 mg   | a di                        |
| HSDVHK-NH2   | <b>Cat. No.:</b> HY-P1187                | HSDVHK-NH2 TFA   | <b>Cat. No.:</b> HY-P1187A  |
| HSDVHK-NH2 is an antagonist of the integrin $\alpha\nu\beta3\text{-vitronectin}$ interaction, with an $IC_{s0}$ of 1.74 pg/mL (2.414 pM).<br>.   |  | HSDVHK-NH2 TFA is an antagonist of the <b>integrin</b> $\alpha \gamma \beta$ 3-vitronectin interaction, with an IC <sub>50</sub> of 1.74 pg/mL (2.414 pM).<br>br/>.  | ารรู้ใหญ่หนัก-<br>น้ำ       |
| Purity:99.63%Clinical Data:No Development ReportedSize:5 mg, 10 mg   |  | Purity: >98%   Clinical Data: No Development Reported   Size: 1 mg, 5 mg   | <u>11</u>                   |
| ICAM-1-IN-1  | <b>Cat. No.:</b> HY-U00003               | ILK-IN-2<br>(OSU-T315 analog)  | <b>Cat. No.</b> : HY-18676B |
| ICAM-1-IN-1 is a potent and selective inhibitor of <b>E-selectin</b> and <b>ICAM-1</b> with <b>IC</b> <sub>50</sub> values of 7 and 5 nM, respectively.  | N S HN-                                  | ILK-IN-2 (OSU-T315 analog) is a ILK inhibitor.   | ۳۵<br>۲۰                    |
| Purity:     99.94%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 100   | mg                                       | Purity:99.41%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 2 mg, 5 mg, 50 mg   | ÷0.0                        |
| ILK-IN-3   | <b>Cat. No.:</b> HY-115677               | Integrin Antagonists 27  | <b>Cat. No.</b> : HY-18668  |
| ILK-IN-3 is an <b>integrin linked kinase</b> inhibitor with antitumor activity.  | H <sub>2</sub> N NH<br>N <sub>N</sub> NN | Integrin Antagonists 27 is a small molecule integrin $\alpha v \beta 3$ antagonist with binding affinity of 18 nM, as s novel anticancer agent.  | O'ris-y-                    |
| Purity:     99.57%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 100   | mg                                       | Purity: ≥98.0%   Clinical Data: No Development Reported   Size: 10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg   |                             |

| Integrin modulator 1   |                                  | Integrin-IN-2  |  |
|--|----------------------------------|--|--|
|  | Cat. No.: HY-134130              |  | Cat. No.: HY-130119  |
| Integrin modulator 1 is a potent and selective $\alpha 4\beta 1$ integrin agonist, with an IC <sub>50</sub> of 9.8 nM for RGD-binding $\alpha 4\beta 1$ . Integrin modulator 1 increases cell adhesion mediated by $\alpha 4\beta 1$ integrin, with an EC <sub>50</sub> of 12.9 nM.  | CT I S COH                       | Integrin-IN-2 (compound 39) is an orally<br>bioavailable pan $\alpha v$ integrin inhibitor.<br>Integrin-IN-2 can increases the $\alpha v \beta 6$ , $\alpha v \beta 3$ ,<br>$\alpha v \beta 5$ and $\alpha v \beta 8$ binding affinities with pIC <sub>50</sub><br>values of 7.8, 8.4, 8.4 and 7.4, respectively.  | A A A A A A A A A A A A A A A A A A A  |
| Purity:99.43%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg,  | 100 mg                           | Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  |  |
| iRGD peptide<br>(c(CRGDKGPDC))   | Cat. No.: HY-P0122               | Irigenin   | <b>Cat. No.</b> : HY-N2587   |
| iRGD peptide is a 9-amino acid cyclic peptide,<br>triggers tissue penetration of drugs by first<br>binding to <b>av integrins</b> , then proteolytically<br>cleaved in the tumor to produce CRGDK/R to<br>interact with neuropilin-1, and has<br>tumor-targeting and tumor-penetrating properties.<br><b>Purity:</b> 99.03%<br><b>Clinical Data:</b> No Development Reported<br><b>Size:</b> 1 mg, 5 mg, 10 mg | CROKOPDC (Deuffet brigs:Cys-Cys) | Irigenin is a is a lead compound, and mediates its<br>anti-metastatic effect by specifically and<br>selectively blocking $\alpha 9\beta 1$ and $\alpha 4\beta 1$ integrins<br>binding sites on C-C loop of Extra Domain A (EDA).<br>Irigenin shows anti-cancer properties.Purity:99.84%<br>Clinical Data:<br>Size:90.82%<br>Size:  | HO OH O OH   |
| LDV  | <b>Cat. No.:</b> HY-P2267        | Leukadherin-1  | <b>Cat. No.:</b> HY-15701  |
| LDV, a tripeptide, is a non-fluorescent analog of LDV-FITC. LDV is a $\alpha 4\beta 1$ integrin (VLA-4) ligand, and binds $\alpha 4\beta 1$ integrin in leukemia cells.  | othouthoffice of the             | Leukadherin-1, a specific agonist of the leukocyte surface integrin CD11b/CD18, increases CD11b/CD18-dependent cell adhesion to fibrinogen with an EC <sub>50</sub> of 4 $\mu$ M.  | Cryfs Clor   |
| Purity:>98%Clinical Data:Phase 4Size:1 mg, 5 mg  |                                  | Purity:     ≥98.0%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg, 2  | 200 mg   |
| Lifitegrast  |                                  | LXW7   |  |
| (SAR 1118; SHP-606)  | Cat. No.: HY-19344               |  | Cat. No.: HY-P0178   |
| Liftegrast (SAR 1118) is an integrin lymphocyte<br>function-associated antigen-1 (LFA-1; $\alpha$ L $\beta$ 2)<br>antagonist; inhibits Jurkat T cell attachment to<br>ICAM-1 with an IC <sub>50</sub> of 2.98 nM.<br>Purity: 99.58%<br>Clinical Data: Launched<br>Size: 10 mM × 1 mL, 10 mg, 50 mg, 100 mg   | S. Lippos                        | LXW7, a cyclic peptide containing Arg-Gly-Asp<br>(RGD), is an <b>integrin</b> $\alpha\nu\beta3$ inhibitor. LXW7 has a<br>high binding affinity to $\alpha\nu\beta3$ <b>integrin</b> with an<br>IC <sub>50</sub> of 0.68 $\mu$ M. LXW7 increases phosphorylation of<br>VEGFR-2 and activation of ERK1/2.<br>Anti-inflammatory effect.<br>Purity: >98%<br>Clinical Data: No Development Reported<br>Size: 1 mq, 5 mg | HAN SNH<br>NH<br>HN SH<br>HN SCOOH<br>HN<br>HN<br>HN<br>HN<br>HN<br>HN<br>HN<br>HN<br>HN<br>HN<br>HN<br>HN<br>HN |
|  |                                  |  |  |
| LXW7 TFA   | <b>Cat. No.:</b> HY-P0178A       | MK-0429<br>(L-000845704)   | <b>Cat. No.</b> : HY-15102   |
| LXW7 TFA, a cyclic peptide containing Arg-Gly-Asp<br>(RGD), is an integrin $\alpha\nu\beta3$ inhibitor. LXW7 has a<br>high binding affinity to $\alpha\nu\beta3$ integrin with an<br>IC <sub>50</sub> of 0.68 $\mu$ M. LXW7 TFA increases<br>phosphorylation of VEGFR-2 and activation of<br>ERK1/2. Anti-inflammatory effect.   |                                  | MK-0429 (L-000845704) is an orally active, potent, selective and nonpeptide <b>pan-integrin</b> antagonist with <b>IC</b> <sub>50</sub> values of 1.6 nM, 2.8 nM, 0.1 nM, 0.7 nM, 0.5 nM and 12.2 nM for $\alpha$ v $\beta$ 1, $\alpha$ v $\beta$ 3, $\alpha$ v $\beta$ 5, $\alpha$ v $\beta$ 6, $\alpha$ v $\beta$ 8 and $\alpha$ 5 $\beta$ 1, respectively.  | 5  |
| Purity:   99.17%     Clinical Data:   No Development Reported     Size:   10 mM × 1 mL, 1 mg, 5 mg, 10 mg  |                                  | Purity:     99.84%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg   |  |

| Natalizumab   |                            | OSU-T315   |                             |
|---|----------------------------|--|-----------------------------|
| Natalizumab   | Cat. No.: HY-108831        | 050-1515   | Cat. No.: HY-18676          |
| Natalizumab is a recombinant, humanized IgG4<br>monoclonal antibody, binds to $\alpha 4\beta 1$ -integrin and<br>blocks its interaction with vascular cell adhesion<br>molecule-1 (VCAM-1). Natalizumab can be used for<br>the treatment of relapsing remitting multiple<br>sclerosis and Crohn's disease.<br>Purity: 99.10%<br>Clinical Data: Launched<br>Size: 10 mg, 25 mg | Natalizumab                | OSU-T315 (ILK-IN-1) is a small Integrin-linked<br>kinase (ILK) inhibitor with an $IC_{50}$ of 0.6 $\mu$ M,<br>inhibiting PI3K/AKT signaling by dephosphorylation<br>of AKT-Ser473 and other ILK targets (GSK-3 $\beta$ and<br>myosin light chain).<br>Purity: 99.88%<br>Clinical Data: No Development Reported<br>Size: 10 mM × 1 mL, 5 mg, 10 mg, 50 mg | 70-0-44 <sup>4</sup> 4      |
|   |                            |  |                             |
| Pyrintegrin   | <b>Cat. No.</b> : HY-13306 | R-BC154 acetate  | Cat. No.: HY-136214         |
| Pyrintegrin is an <b>β1-integrin</b> agonist and a 2,4-disubstituted pyrimidine that promotes embryonic stem cells survival. Pyrintegrin enhances cell-extracellular matrix (ECM) adhesion-mediated <b>integrin</b> signaling.  |                            | R-BC154 acetate is a selective fluorescent $\alpha_{9}\beta_{1}$ integrin antagonist. R-BC154 acetate acts as auseful high affinity, activation dependentintegrin probe, which can be used to investigate $\alpha 9\beta_{1}$ and $\alpha 4\beta_{1}$ integrin binding activity.Purity:>98%  |                             |
| Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg   |                            | Clinical Data: No Development Reported<br>Size: 1 mg, 5 mg   |                             |
| RGD   |                            | RGD peptide (GRGDNP)   |                             |
|   | Cat. No.: HY-P0278         | - Politica ( 2007)   | Cat. No.: HY-P1740          |
| RGD is a tripeptide that effectively triggers cell<br>adhesion, addresses certain cell lines and elicits<br>specific cell responses; binds to <b>integrins</b> .  |                            | RGD peptide (GRGDNP) acts as an inhibitor of<br>integrin-ligand interactions and plays an<br>important role in cell adhesion, migration,<br>growth, and differentiation.   | MANT HIT HIT AND            |
| Purity:>98%Clinical Data:Phase 2Size:1 mg, 5 mg   |                            | Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  | HNLIN                       |
| RGD peptide (GRGDNP) (TFA)  |                            | RGD Trifluoroacetate   |                             |
| RGD peptide (GRGDNP) (TFA) acts as an inhibitor of<br>integrin-ligand interactions and plays an<br>important role in cell adhesion, migration,<br>growth, and differentiation.  | Cat. No.: HY-P1740A        | RGD Trifluoroacetate is a tripeptide that<br>effectively triggers cell adhesion, addresses<br>certain cell lines and elicits specific cell<br>responses; RGD Trifluoroacetate binds to <b>integrins</b> .  | Cat. No.: HY-P0278A         |
| Purity:99.25%Clinical Data:No Development ReportedSize:1 mg, 5 mg   | н,л <sup>⊥</sup> ын ⊊¦он   | Purity:99.29%Clinical Data:No Development ReportedSize:5 mg, 10 mg, 25 mg, 50 mg, 100 mg   | 51 v.                       |
| RO0270608   | <b>Cat. No.:</b> HY-138542 | RWJ 50271  | <b>Cat. No.</b> : HY-110086 |
| RO0270608, the active metabolite of R411, is a<br>dual <b>alpha4beta1-alpha4beta7</b> (α4β1/α4β7) integrin<br>antagonist. Antiinflammatory activity.  |                            | RWJ 50271 is an selective and orally active<br>inhibitor of <b>lymphocyte function-associated</b><br><b>antigen-1/intercellular adhesion</b><br><b>molecule-1(LFA-1/ICAM-1)</b> interaction with an<br><b>IC</b> <sub>so</sub> of 5.0 μM (HL60 cells), RWJ 50271 inhibits<br>LFA-1/ICAM-1-mediated cell adhesion.  | холи у Цтон                 |
| Purity:     98.69%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg   | , 100 mg                   | Purity:99.51%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 5 mg  |                             |

#### SB-267268 SB-273005 Cat. No.: HY-19306 Cat. No.: HY-19307 SB-267268 is a selective and nonpeptidic SB-273005 is a potent nonpeptide and orally active alpha(v)beta3 ( $\alpha v\beta 3$ ) and alpha(v)beta5 ( $\alpha v\beta 5$ ) integrin antagonist with K s of 1.2 nM and 0.3 nM for αvβ3 receptor and αvβ5 receptor, respectively. integrins antagonist, with K<sub>i</sub>s of 0.9, 0.5 and 0.7 nM for human $\alpha\nu\beta3$ , monkey $\alpha\nu\beta3$ and human $\alpha\nu\beta5$ , respectively. Purity: > 98% Purity: >98% Clinical Data: No Development Reported Clinical Data: No Development Reported Size: 1 mg, 5 mg Size: 1 mg, 5 mg Sibrafiban SR121566A (RO 48-3657) Cat. No.: HY-10309 Cat. No.: HY-U00235 Sibrafiban (RO 48-3657) is the orally active, SR121566A is a novel non-peptide Glycoprotein nonpeptide, double-prodrug of Ro 44-3888 and a IIb/IIIa (GP IIb-IIIa) antagonist, which can selective glycoprotein IIb/IIIa receptor inhibit ADP-, arachidonic acid- and of Flogen antagonist. Sibrafiban inhibits platelet collagen-induced human platelet aggregation with IC<sub>so</sub>s of 46±7.5, 56±6 and 42±3 nM, respectively. aggregation. Purity: > 98% **Purity:** >98% Clinical Data: No Development Reported Clinical Data: No Development Reported 1 mg, 5 mg Size: 5 ma Size: TC-I 15 TC113 Cat. No.: HY-107588 Cat. No.: HY-145314 TC113 is a c(RGDyK)-Based conjugate of Gemcitabine TC-I 15 (TC-I-15) is an allosteric, collagen-binding integrin $\alpha 2\beta 1$ inhibitor with (GEM). TC113 could be internalized by A549 cells IC values of 26.8 $\mu$ M and 0.4 $\mu$ M for GFOGER and through integrin $\alpha$ $\beta_{2}$ . TC113 shows potent GLOGEN, respectively. TC-I 15 inhibits platelet antiproliferative properties against WM266.4 and adhesion to collagen and thrombus deposition. A549 cells Purity: >98% Purity: >98% Clinical Data: No Development Reported Clinical Data: No Development Reported Size: 1 mg, 5 mg Size: 1 mg, 5 mg **TCS 2314** Tetrac (Tetraiodothyroacetic acid; 3,3',5,5'-Tetraiodothyroacetic acid) Cat. No.: HY-12308 Cat. No.: HY-W008859 TCS 2314 (compound 3) is orally active and Tetrac (Tetraiodothyroacetic acid), a derivative selective very late antigen-4 (VLA-4, α4β1, of L-thyroxine (T4), is a thyrointegrin receptor CD49d/CD29) antagonist with an IC<sub>50</sub> of 4.4 nM. antagonist. Tetrac blocks the actions of T4 and 3,5,3'-triiodo-L-thyronine (T3) at the cell surface receptor for thyroid hormone on integrin ανβ3. ≥99.0% Purity: ≥95.0% Purity: Clinical Data: No Development Reported Clinical Data: No Development Reported 10 mM × 1 mL, 10 mg, 25 mg Size: 5 ma Size: THI0019 Tirofiban Cat. No.: HY-117388 (L700462; MK383) Cat. No.: HY-17369B THI0019 is a potent integrin $\alpha 4\beta 1$ (VLA-4) agonist Tirofiban(L700462;MK383) is a potent non-peptide, with an $\text{EC}_{\text{so}}$ range of 1-2 $\mu\text{M}.$ THI0019 induces glycoprotein IIb/IIIa (integrins alphaIIbbetaIII) stem/progenitor cells adhesion. THI0019 also antagonist Target: integrin IIb/IIIa Tirofiban hydrochloride monohydrate blocks platelet regulates adhesion mediated by $\alpha 4\beta 7,\,\alpha 5\beta 1$ and αLβ2. aggregation and thrombus formation. Purity: 98.31% Purity: 98.37% Clinical Data: No Development Reported Clinical Data: Launched 5 mg, 10 mg, 25 mg, 50 mg, 100 mg Size: Size: 5 mg, 10 mg, 50 mg, 100 mg

| Tirofiban hydrochloride monohydrate  |  | Tirofiban-d9  |  |
|--|--|---|--|
| nonban nyurochionde mononyurate  | Cat. No.: HY-17369   | (L700462-d9; MK383-d9)  | Cat. No.: HY-17369BS                                       |
| Tirofiban hydrochloride monohydrate is a potent<br>non-peptide, glycoprotein IIb/IIIa (integrins<br>alphaIIbbetaIII) antagonist IC50 value: Target:<br>integrin IIb/IIIa Tirofiban hydrochloride<br>monohydrate blocks platelet aggregation and<br>thrombus formation.   | ~~ <sup>0,0</sup><br>остра<br>остра<br>ино ио <sup>н</sup> | Tirofiban-d9 is deuterium labeled Tirofiban.  | eq eee t for   |
| Purity:     99.34%       Clinical Data:     Launched       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg  |  | Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg   |  |
| Tirofiban-d9 hydrochloride   | <b>Cat. No.</b> : HY-17369AS                               | TR-14035  | <b>Cat. No.:</b> HY-15770                                  |
| Tirofiban-d9 (L700462-d9) hydrochloride is the<br>deuterium labeled Tirofiban. Tirofiban(L700462) is<br>a potent non-peptide, glycoprotein IIb/IIIa<br>(integrins alphaIIbbetaIII) antagonist.   | Contraction Contraction Contraction                        | TR-14035 is a orally active dual $\alpha_4\beta_7/\alpha_4\beta_1$<br>integrin antagonist, with IC <sub>50</sub> s of 7 nM and 87 nM<br>for $\alpha_4\beta_7$ and $\alpha_4\beta_{12}$ respectively. TR-14035 can<br>be used for the research of inflammation and<br>autoimmune diseases. | C L HOLDO  |
| Purity:>98%Clinical Data:Size:1 mg, 10 mg  |  | Purity:     95.81%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg  |  |
| Valategrast<br>(R-411 free base)   | <b>Cat. No.</b> : HY-14190                                 | Valategrast hydrochloride<br>(R-411)  | <b>Cat. No.:</b> HY-14189                                  |
| Valategrast (R-411 free base) is a potent and<br>orally active integrin $\alpha 4\beta 1$ (VLA-4) and $\alpha 4\beta 7$ dual<br>antagonist. Valategrast has the potential for<br>Chronic obstructive pulmonary disease (COPD) and<br>asthma treatment.<br>Purity: 98.57%<br>Clinical Data: No Development Reported |  | Valategrast hydrochloride (R-411) is a potent<br>integrin α4β1 (VLA-4) and α4β7 dual antagonist.<br>Valategrast hydrochloride has the potential for<br>Chronic obstructive pulmonary disease (COPD) and<br>asthma treatment.<br>Purity: >98%<br>Clinical Data: No Development Reported    |  |
| Size: 5 mg, 10 mg, 25 mg, 50 mg, 100 mg  |  | Size:     5 mg, 10 mg, 25 mg, 50 mg, 100 mg   |  |
| Vedolizumab<br>(Anti-Human lymphocyte α4β7 integrin, Humanized Antibo  | ody) Cat. No.: HY-P9911                                    | Vedolizumab (anti-α4β7-integrin)  | Cat. No.: HY-P9911A  |
| Vedolizumab is a humanized IgG1 monoclonal antibody that targets the $\alpha 4\beta7$ <b>integrin</b> for the treatment of ulcerative colitis and Crohn's disease.   | Vedolizumab  | Vedolizumab (anti- $\alpha 4\beta 7$ -integrin) is a humanized IgG1 monoclonal antibody that targets the $\alpha 4\beta 7$ integrin for the treatment of ulcerative colitis and Crohn's disease.  | Vedolizumab (anti-α <sub>4</sub> β <sub>7</sub> -integrin) |
| Purity:99.64%Clinical Data:LaunchedSize:1 mg, 5 mg, 25 mg, 50 mg   |  | Purity:>98%Clinical Data:LaunchedSize:1 mg, 5 mg  |  |
| XVA143   | <b>Cat. No.:</b> HY-139202                                 | Zaurategrast<br>(CT7758)  | <b>Cat. No.:</b> HY-70073                                  |
| XVA143, an $\alpha/\beta$ I-like allosteric antagonist,<br>inhibits LFA-1 dependent firm adhesion, while at<br>the same time it enhances adhesion in shear flow<br>and rolling both in vitro and in vivo.  | "Qinjiqira"  | Zaurategrast (CT7758) is a potent and oral-effective $\alpha_4\text{-}integrin$ inhibitor.  | HN CO OH   |
| Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  |  | Purity:98.03%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 5 mg, 10 mg, 25 mg   |  |

| Zaurategrast ethyl ester   |   | Zaurategrast ethyl ester sulfate  |  |
|--|---|---|--|
| (CDP323; UCB1184197)   | Cat. No.: HY-75385  | (CDP323 sulfate; UCB1184197 sulfate)  | Cat. No.: HY-75385A  |
| Zaurategrast ethyl ester (CDP323), the ethyl ester<br>prodrug of CT7758, is a $\alpha 4\beta 1/\alpha 4\beta 7$ integrin<br>antagonist used for the treatment of inflammatory<br>and autoimmune disorders.               |   | Zaurategrast ethyl ester sulfate (CDP323 sulfate),<br>the ethyl ester prodrug of CT7758, is a<br>$\alpha 4\beta 1/\alpha 4\beta 7$ integrin antagonist used for the<br>treatment of inflammatory and autoimmune<br>disorders.         |  |
| Purity:     99.06%       Clinical Data:     Phase 2       Size:     10 mM × 1 mL, 10 mg, 25 mg, 50 mg, 100 mg  |   | Purity:>98%Clinical Data:Phase 2Size:1 mg, 5 mg   | ŎН   |
| α2β1 Integrin Ligand Peptide   | <b>Cat. No.:</b> HY-P1868   | $\alpha 2\beta 1$ Integrin Ligand Peptide TFA   | <b>Cat. No.</b> : HY-P1868A                                  |
| $\alpha 2\beta 1$ Integrin Ligand Peptide interacts with the $\alpha 2\beta 1$ integrin receptor on the cell membrane and mediates extracellular signals into cells. It is a potential antagonist of collagen receptors. | но Страна Стр | $\alpha 2\beta 1$ Integrin Ligand Peptide TFA interacts with<br>the $\alpha 2\beta 1$ integrin receptor on the cell membrane<br>and mediates extracellular signals into cells. It<br>is a potential antagonist of collagen receptors. | HOLO<br>HOLO<br>HOLO<br>HOLO<br>HOLO<br>HOLO<br>HOLO<br>HOLO |
| Purity: >98%   Clinical Data: No Development Reported   Size: 1 mg, 5 mg, 10 mg  |   | Purity:99.33%Clinical Data:No Development ReportedSize:1 mg, 5 mg, 10 mg  | F.J. CO.   |
| α5β1 integrin agonist-1  |   | αvβ1 integrin-IN-1  |  |
|  | Cat. No.: HY-139702   |   | Cat. No.: HY-100445  |
| α5β1 integrin agonist-1, acting as <b>α5β1</b><br><b>integrin</b> agonist, is able to selectively deliver<br>5-FU into tumor cells, successfully leading to<br>cancer cell death.<br>Purity: >98%                        | C C C C C C C C C C C C C C C C C C C   | ανβ1 integrin-IN-1 (Compound C8) is a potent and selective ανβ1 integrin inhibitor with an IC <sub>50</sub> of 0.63 nM. Antifibrotic effects.     Purity:   >98%  | HALL - LACOU   |
| Clinical Data: No Development Reported<br>Size: 1 mg, 5 mg   |   | Clinical Data: No Development Reported<br>Size: 5 mg, 10 mg, 50 mg, 100 mg  |  |
| ανβ1 integrin-IN-1 TFA   |   | αvβ5 integrin-IN-1  |  |
|  | Cat. No.: HY-100445A  |   | Cat. No.: HY-145363  |
| ανβ1 integrin-IN-1 TFA (Compound C8) is a potent and selective <b>ανβ1 integrin</b> inhibitor with an IC <sub>50</sub> of 0.63 nM. Antifibrotic effects.   | 2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2<br>2                     | $\alpha\nu\beta5$ integrin-IN-1 is a first potent and selective $\alpha\nu\beta5$ integrin inhibitor (pIC_{50} = 8.2) .   | ~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~                       |
| Purity:     98.30%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg   |   | Purity:>98%Clinical Data:No Development ReportedSize:5 mg, 10 mg, 25 mg, 50 mg, 100 mg  |  |



## **Kinesin**

Kinesins are a family of molecular motors that use the energy of ATP hydrolysis to move along the surface of, or destabilize, microtubule filaments. The kinesin motor protein family consists of 14 distinct subclasses and 45 kinesin proteins in humans. A large number of these proteins, or their orthologues, have been shown to possess essential function(s) in both the mitotic and the meiotic cell cycle. Kinesins also can be classified into three groups based on the position of their motor domains: N-terminal, C-terminal and internal kinesins. Conventional kinesin operates as a dimer, walking in a co-ordinated, hand-over-hand fashion along a microtubule protofilament.

Kinesins have important roles in chromosome separation, microtubule dynamics, spindle formation, cytokinesis and cell cycle progression. Roles of kinesins in diseases typically involve defective transport of cell components, transport of pathogens, or cell division.



## **Microtubule/Tubulin**

Microtubules are a component of the cytoskeleton, found throughout the cytoplasm. These tubular polymers of tubulin can grow as long as 50 micrometres, with an average length of 25  $\mu$ m, and are highly dynamic. The outer diameter of a microtubule is about 24 nm while the inner diameter is about 12 nm. Microtubules are found in eukaryotic cells and are formed by the polymerization of a dimer of two globular proteins, alpha and beta tubulin. Tubulin is one of several members of a small family of globular proteins. The tubulin superfamily includes five distinct families, the alpha-, beta-, gamma-, delta-, and epsilon-tubulins and a sixth family which is present only in kinetoplastid protozoa. The most common members of the tubulin family are  $\alpha$ -tubulin and  $\beta$ -tubulin, the proteins that make up microtubules. Microtubules are very important in a number of cellular processes. They are involved in maintaining the structure of the cell.





Monopolar spindle 1 (Mps1/TTK) is a serine/threonine kinase conserved from yeast to human. It has been shown to function as the key kinase that activates the spindle assembly checkpoint (SAC) to secure proper distribution of chromosomes to daughter cells.

MPS1, a dual specificity protein kinase, is also one of the main components of the SAC and ensures cells do not progress from metaphase to anaphase until the kinetochores are properly attached to the microtubules and under the appropriate tension at the metaphase plate. Cancer cells heavily rely on MPS1 to cope with aneuploidy resulting from aberrant numbers of chromosomes. The kinase has been found to be upregulated in a large number of tumor types. Mps1 is an attractive oncology target due to its high expression level in cancer cells as well as the correlation of its expression levels with histological grades of cancers.



# Myosin

Myosins are mechanoenzymes that interact with actin filaments and hydrolyse ATP to generate movement and force. This enables myosins to propel the sliding of actin filaments, to produce tension on actin filaments and to walk along these filaments. As a result, myosins can regulate the structure and dynamics of the actin cytoskeleton and affect the localization and transport of cellular components. The different myosins are grouped into classes on the basis of their motor domains. There are 35 known classes of myosin, and humans have 40 myosin genes that fall into 13 classes (I, II, III, V, VI, VI, XV, XVI, XVII, XIX and XXXV).

Myosins are actin-dependent motors that participate in a diverse range of crucial activities, including muscle contraction, intracellular trafficking, cell division, motility, actin cytoskeletal organisation and cell signaling. Myosin malfunction has been implicated in a variety of disorders including deafness, hypertrophic cardiomyopathy, Usher syndrome, Griscelli syndrome and cancer.

Myosin light chain kinase (MLCK) is an enzyme that activates the myosin light chain to exert its function related to cytoskeleton contraction and tight junction regulation. In most cells, MLCK is a transducer for signalling MLC phosphorylation in response to Ca <sup>2+</sup> binding to MLCK-associated calmodulin. MLCK-mediated MLC phosphorylation and actomyosin contractility is important in muscle contraction, cell migration, and endo/exocytic processes, and is recognized for its central role in signalling endothelial cell-cell adhesion and barrier function.

### Myosin Inhibitors, Activators & Modulators

| (+)-Blebbistatin   | Cat No. HV 107657            | (-)-Blebbistatin  | <b>Cat. No.:</b> HY-13441    |
|--|------------------------------|---|------------------------------|
|  | Cat. No.: HY-107657          | ((S)-(-)-Blebbistatin)  | Cat. No.: HY-13441           |
| (+)-Blebbistatin is the inactive enantiomer of<br>(-)-Blebbistatin. (-)-Blebbistatin is a selective<br>inhibitor of myosin II ATPase.  |                              | (-)-Blebbistatin is a selective inhibitor of the ATPase activity of non-muscle <b>myosin II</b> .   | OH<br>N<br>N                 |
| Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  |                              | Purity:99.42%Clinical Data:No Development ReportedSize:5 mg, 10 mg, 50 mg   |                              |
| Aficamten  |                              | ATM-3507  |                              |
| (CK-274; CK-3773274)   | Cat. No.: HY-139465          |   | Cat. No.: HY-100948          |
| Aficamten (CK-274) is a potent <b>cardiac myosin</b> inhibitor with an $IC_{so}$ of 1.4 $\mu$ M. Aficamten can be used for the research of hypertrophic cardiomyopathy (HCM).  | N CON                        | ATM-3507 is a potent $tropomyosin$ inhibitor with $IC_{s_0}s$ from 3.83-6.84 $\mu M$ in human melanoma cell lines.  | <i>a da</i> atoroto          |
| Purity:99.86%Clinical Data:No Development ReportedSize:5 mg, 10 mg, 25 mg, 50 mg, 100 mg   | 0                            | Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg   |                              |
| ATM-3507 trihydrochloride  | <b>Cat. No.</b> : HY-100948B | Blebbistatin  | <b>Cat. No.:</b> HY-13813    |
| ATM-3507 trihydrochloride is a potent tropomyosin inhibitor with $IC_{so}$ s from 3.83-6.84 $\mu$ M in human melanoma cell lines.  | a d'arche                    | Blebbistatin is a selective <b>non-muscle myosin</b><br><b>II</b> ( <b>NMII</b> ) inhibitor, promotes directional<br>migration of corneal endothelial cells (CECs) and<br>accelerates wound healing, and better preserves<br>cell junctional integrity and barrier function.  | ()<br>N<br>N<br>N            |
| Purity:     98.10%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 50 mg, 100 mg   |                              | Purity:     99.64%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg   | 0 он                         |
| BTS  |                              | Danicamtiv  |                              |
| (N-Benzyl-p-toluenesulfonamide; N-Tosylbenzylamine)  | Cat. No.: HY-16690           | (MYK-491)   | Cat. No.: HY-109128          |
| BTS (N-Benzyl-p-toluenesulfonamide) is a potent<br>and selective inhibitor of <b>skeletal muscle myosin</b><br><b>II subfragment 1 (S1) ATPase</b> activity, with an<br>$IC_{so}$ s of ~5 µM for actin- and Ca <sup>2+</sup> -stimulated<br>myosin S1 ATPase. BTS specifically inhibits the<br>contraction of fast skeletal muscle fibers. |                              | Danicamtiv (MYK-491), an inotropic agent, is a<br>selective allosteric activator of <b>cardiac</b><br><b>myosin</b> . Danicamtiv increases cardiac systolic<br>function and preserves mechanical efficiency.  | -N SP N N                    |
| Purity:99.51%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 500 mg  |                              | Purity:     99.49%       Clinical Data:     Phase 2       Size:     10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 1  | 00 mg                        |
| HA-100   | <b>Cat. No.:</b> HY-100984   | HA-100 hydrochloride  | <b>Cat. No.</b> : HY-100984A |
| HA-100 is a potent <b>protein kinase</b> inhibitor, with $IC_{so}^{S}$ of 4 $\mu$ M, 8 $\mu$ M, 12 $\mu$ M and 240 $\mu$ M for cGMP-dependent protein kinase (PKG), cAMP-dependent protein kinase (PKA), protein kinase C (PKC) and MLC-kinase, respectively. HA-100 also used as a ROCK inhibitor.  |                              | HA-100 hydrochloride is a potent <b>protein kinase</b><br>inhibitor, with IC <sub>so</sub> s of 4 $\mu$ M, 8 $\mu$ M, 12 $\mu$ M and 240<br>$\mu$ M for cGMP-dependent protein kinase ( <b>PKG</b> ),<br>cAMP-dependent protein kinase ( <b>PKA</b> ), protein<br>kinase C ( <b>PKC</b> ) and <b>MLC-kinase</b> , respectively. |                              |
| Purity:     99.77%       Clinical Data:     No Development Reported       Size:     10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 1   | 1)<br>00 mg                  | Purity: >98%   Clinical Data: No Development Reported   Size: 1 mg, 5 mg  | H-CI                         |

| Mavacamten   |                             | ML-7 hydrochloride  |                             |
|--|-----------------------------|---|-----------------------------|
| (MYK461; SAR439152)  | Cat. No.: HY-109037         |   | Cat. No.: HY-15417          |
| Mavacamten (MYK461) is an orally active modulator<br>of <b>cardiac myosin</b> , with <b>IC</b> <sub>50</sub> <b>s</b> of 490, 711 nM for<br>bovine cardiac and human cardiac, respectively.  |                             | ML-7 hydrochloride is a naphthalene sulphonamide<br>derivative, potently inhibits <b>MLCK (IC<sub>50</sub>=300</b><br>nM). ML-7 hydrochloride also inhibits <b>YAP/TAZ</b> .  | HN<br>O=S=O<br>H-CI         |
| Purity:     99.90%       Clinical Data:     Phase 3       Size:     10 mM × 1 mL, 1 mg, 5 mg, 10 mg, 25 mg, 50   | mg                          | Purity:99.75%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 10 mg, 50 mg   | Y                           |
| ML-9   | <b>Cat. No.:</b> HY-100932  | ML-9 Free Base  | <b>Cat. No.:</b> HY-100932A |
| $\label{eq:main_set} \begin{array}{l} \text{ML-9 is a selective and potent inhibitor of Akt} \\ \textbf{kinase, inhibits myosin light-chain kinase (MLCK) \\ and stromal interaction molecule 1 (STIM1) \\ activity. ML-9 inhibits inhibits MLCK, PKA and PKC \\ activity with K, values of 4, 32 and 54 \muM, respectively. \\ \hline Purity: 99.89\% \\ \hline Clinical Data: No Development Reported \\ \hline Size: 10 mM \times 1 mL, 25 mg, 50 mg, 100 mg, 250 mg \\ \hline \end{array}$ |                             | ML-9 (Free Base) is a selective and potent inhibitor of Akt kinase, inhibits myosin light-chain kinase (MLCK) and stromal interaction molecule 1 (STIM1) activity.     Purity:   >98%     Clinical Data:   No Development Reported     Size:   1 mg, 5 mg |                             |
| MLCK inhibitor peptide 18  | <b>Cat. No</b> .: HY-P1029  | MS-444<br>(BE-34776)  | <b>Cat. No.:</b> HY-100685  |
| MLCK inhibitor peptide 18 is a myosin light chain kinase (MLCK) inhibitor with an $IC_{50}$ of 50 nM, and inhibits <b>CaM kinase II</b> only at 4000-fold higher concentrations.   | RKKYKYRRK-NH <sub>2</sub>   | MS-444 inhibits the activity of purified smooth muscle myosin light chain kinase (MLCK) with an $IC_{\rm 50}$ value of 10 $\mu M.$  | OH O                        |
| Purity:99.66%Clinical Data:No Development ReportedSize:1 mg, 5 mg, 10 mg, 25 mg  |                             | Purity:99.42%Clinical Data:No Development ReportedSize:10 mM × 1 mL, 5 mg, 10 mg  | I<br>ОН                     |
| MT-134   | <b>Cat. No.:</b> HY-141810  | Omecamtiv mecarbil<br>(CK-1827452)  | <b>Cat. No.:</b> HY-14233   |
| MT-134 is a <b>SkMII</b> -specific inhibitor and has excellent exposure in muscles.  | N SO                        | Omecamtiv mecarbil (CK-1827452) is a selective cardiac myosin activator.  | Vilgo <sup>i.</sup>         |
| Purity:>98%Clinical Data:No Development ReportedSize:1 mg, 5 mg  | O OH                        | Purity:     98.89%       Clinical Data:     Phase 3       Size:     10 mM × 1 mL, 5 mg, 10 mg, 25 mg, 50 mg, 10   | )0 mg                       |
| Omecamtiv mecarbil-d8<br>(CK-1827452-d8)   | <b>Cat. No.</b> : HY-14233S | para-Nitroblebbistatin  | <b>Cat. No.:</b> HY-120870  |
| Omecamtiv mecarbil-d8 (CK-1827452-d8) is the<br>deuterium labeled Omecamtiv mecarbil. Omecamtiv<br>mecarbil (CK-1827452) is a selective <b>cardiac</b><br><b>myosin</b> activator.   |                             | para-Nitroblebbistatin is a non-cytotoxic,<br>photostable, fluorescent and specific <b>Myosin II</b><br>inhibitor, usd in the study of the specific role<br>of myosin II in physiological, developmental, and<br>cell biological studies.                 | O'N+O'                      |
| Purity: >98%   Clinical Data: No Development Reported   Size: 1 mg, 5 mg   |                             | Purity: >98%   Clinical Data: No Development Reported   Size: 500 μg  | C OH                        |

#### W-7 hydrochloride Pentachloropseudilin (Antibiotic A 15104 Y; PCIP) Cat. No.: HY-115669 Cat. No.: HY-100912 Pentachloropseudilin (Antibiotic A 15104 Y; PCIP) W-7 hydrochloride is a selective calmodulin is a reversible and allosteric potent inhibitor of antagonist. W-7 hydrochloride inhibits the $\begin{array}{l} \mbox{magnitude} \ \mbox{my order number of the transmission of transmissi$ Myols (class 1 myosins) with $IC_{50}$ s range from 1 to 5 $\mu\text{M}$ for mammalian class-1 myosins and greater H-CI than 90 $\mu$ M for class-2 and class-5 myosins. induces apoptosis and has antitumor activity. ≥98.0% Purity: 99.65% Purity: Clinical Data: No Development Reported Clinical Data: No Development Reported Size: 5 mg Size: 10 mM × 1 mL, 25 mg, 50 mg





PAKs (p21-activated kinases) are a family of six serine/threonine kinases that act as key effectors of RHO family GTPases in mammalian cells. PAKs are subdivided into two groups: group I (PAK1, PAK2, and PAK3) and group II (PAK4, PAK5, and PAK6), based on their domain architecture and regulation. Group I PAKs are activated by GTPases such as Cdc42, Rac, TC10, CHP, and Wrch-1, as well as in a GTPase-independent manner. Group II PAKs are generally not activated by Cdc42/Rac binding. PAK plays important roles in cytoskeletal organization, cellular morphogenesis, and survival, and members of this family have been implicated in many diseases including cancer, infectious diseases, and neurological disorders.

PAKs participate in various signaling networks. PAKs activate the MAPK pathway by phosphorylating Raf1 in addition to NF-κB. PAKs also phosphorylate a number of regulators of the cytoskeleton such as MLCK, LIMK, filamin A, ILK, merlin, and Arpc1b. In addition, PAKs regulate survival and apoptotic pathways through phosphorylation of its effectors such as DLC1 and BimL. On translocation to the nucleus, PAKs directly affect gene transcription. Several transcription factors and transcriptional co-regulators such as FKHR, SHARP, CTBP1 and SNAI1 are substrates to PAK1. PAKs also regulate cell cycle progression through phosphorylation of histone H3, Aurora A and PIK1.



## ROCK

#### Rho-associated protein kinase; Rho-associated kinase; Rho-kinase; ROK

ROCK (Rho-associated protein kinase) is a kinase belonging to the AGC (PKA/ PKG/PKC) family of serine-threonine kinases. ROCKs (ROCK1 and ROCK2) occur in mammals, zebrafish, Xenopus, invertebrates and chicken. Human ROCK1 has a molecular mass of 158 kDa and is a major downstream effector of the small GTPase RhoA. Mammalian ROCK consists of a kinase domain, acoiled-coil region and a Pleckstrin homology (PH) domain, which reduces the kinase activity of ROCKs by an autoinhibitory intramolecular fold if RhoA-GTP is not present. ROCK plays a role in a wide range of different cellular phenomena, as ROCK is a downstream effector protein of the small GTPase Rho, which is one of the major regulators of the cytoskeleton.