

CYCLIN DEPENDENT KINASE [CDK] INHIBITORS

Inhibitor	Structure	Prod. No.	Size	CDK1	CDK2	CDK4	CDK5	GSK-3	Other Enzymes
Aloisine [RP106; 7- <i>n</i> -Butyl-6-(4-methoxyphenyl)[5H]pyrrolo[2,3- <i>b</i>]pyrazine]		ALX-270-386-M001	1 mg	700nM (B) [1]			1.5µM (p35) [1]	920nM [1]	
Aloisine A [RP107; 7- <i>n</i> -Butyl-6-(4-hydroxyphenyl)[5H]pyrrolo[2,3- <i>b</i>]pyrazine]		ALX-270-385-M001	1 mg	150nM (B) [1]	120nM (A) [1] 400nM (E) [1]	>100µM (D1) [1]	160nM (p35) [1] 200nM (p25) [1]	500nM (α) [1] 1.5µM (β) [1]	erk1: 18µM [1] erk2: 22µM [1] JNK: 3.3-10µM [1] IRTK: 60µM [1] PI3K: >10µM [1]
Alsterpaullone [9-Nitro-7,12-dihydroindolo-[3,2- <i>d</i>]1]benzazepin-6(5 <i>H</i>)-one]		ALX-270-275-M001 ALX-270-275-M005	1 mg 5 mg	35nM (B) [2]	15nM (A) [2] 80nM (A) [3] 200nM (E) [2]	>10µM (D1) [2]	40nM (p25) [2]	4nM (α/β) [2] 110nM (β) [3]	erk1: 22µM [2] erk2: 4.5µM [2] LCK: 470nM [3]
3-Amino-1<i>H</i>-pyrazolo[3,4-<i>b</i>]oxazoline		ALX-270-387-M001 ALX-270-387-M005	1 mg 5 mg	600nM (B) [4]			400nM (p25) [4]	1µM (β) [4]	
Aminopurvalanol A [2 <i>R</i>]-2-(6-((3-Amino-5-chlorophenyl)amino-3-(1-methylethyl)-9 <i>H</i> -purin-2-yl)amino)-3-methyl-1-butanol; [NC-27]		ALX-270-249-M001 ALX-270-249-M005	1 mg 5 mg	33nM (B) [5]	33nM (A) [5] 28nM (E) [5]		20nM (p35) [5]		
Arcyriaflavin A		ALX-350-375-M001	1 mg			59nM [6]			
3-ATA [3-Amino-9-thio(10 <i>H</i>)-acridone]		ALX-350-273-M001 ALX-350-273-M005	1 mg 5 mg	88µM (A) [7]	>100µM (A) [7] 100µM (E) [7]	3.1µM (D1) [7]			
Benfluorene [Ethyl 6-hydroxy-4-phenylbenzoate, 5-fluoro-2,3-dipyrindine-3-carboxylate; 1-Aza-9-oxafluorene 5a]		ALX-270-388-M001 ALX-270-388-M005	1 mg 5 mg	28.8µM [8]	>100µM (E) [8]	>100µM (D1) [8]	6% inhibition at 10µM [8]		
BML-259 [N-(5-Isopropylthiazol-2-yl)phenylacetamide]		ALX-270-487-M005	5 mg		98nM (E) [9]		64nM (p25) [9]		
Bohemine [6-Benzylamino-2-(3-hydroxypropylamino)-9-isopropylpurine]		ALX-270-390-M001 ALX-270-390-M005	1 mg 5 mg	1.1µM (B) [10]	600nM (E) [10]				
Bohemine, 2-Hydroxy- [6-(3-Hydroxybenzylamino)-2-(hydroxypropylamino)-9-isopropylpurine]		ALX-270-395-M001 ALX-270-395-M005	1 mg 5 mg	100nM [11]	80nM [11]				
Borrelidin [2-(7-Cyano-8,16-dihydroxy-9,11,15-tetramethyl-18-oxooxacyclooctadeca-4,6-dien-2-yl)cyclopentanecarboxylic acid; Treponemycin; U 785-48; C2989]		ALX-380-102-MC05 ALX-380-102-M001	0.5 mg 1 mg					Cdc28/Cln2 (yeast CDK) : 24µM [12]	
2-Bromo-12,13-dihydro-5<i>H</i>-indolo[2,3-<i>a</i>]pyrrolo[3,4-<i>c</i>]carbazole-5,7(6<i>H</i>)-dione		ALX-270-403-M001	1 mg	2.1µM (B) [13]	520nM (E) [13]	76nM (D1) [13]		>20µM [13]	CaMKII : 12.4µM [13] PKA : >20µM [13]
Butyrolactone I		ALX-350-250-C200	200 µg	0.68µM (B) [14]					
CGP 74514A [N-(<i>cis</i> -2-Aminocyclohexyl)-N-(3-chlorophenyl)-9-ethyl-9 <i>H</i> -purine-2,6-diamine]		ALX-270-391-M001 ALX-270-391-M005 ALX-270-391-M025	1 mg 5 mg 25 mg	25nM [16]				PKC-α: 6µM [16] PKA: 12.5µM [16] EGFR: >10µM [16]	
CDK1/2 Inhibitor III [5-Amino-3-(4-(aminosulfonyl)phenylamino)-N-(2,6-difluorophenyl)-1 <i>H</i> -1,4-triazole-1-carbothioamide]		ALX-270-442-M001	1 mg	600pM (B) [17]	500pM (A) [17]			32nM (B) [17]	VEGF-R2: 140nM [17]
3-(2-Chloro-3-indolyl-methylene)-1,3-dihydroindol-2-one		ALX-270-392-M001 ALX-270-392-M005	1 mg 5 mg	5.8µM (B) [18]		25µM [18]		>100µM (β) [18]	
Compound 52 [2-(2-Hydroxyethylamino)-6-(3-chloroanilino)-9-isopropylpurine]		ALX-270-248-M001	1 mg	340nM [19]				Cdc28c: 7µM [19] Pho85p: 2µM [19]	
CR8		INQUIRE!		0.09µM (B) [20]	0.072µM (A) [20] 0.041µM (E) [20]		0.11µM (p25) [20]	12.0µM (α/β) [20]	CDK2: 1.10µM (H) [20] CDK9: 0.18µM (T) [20] CK1α/c: 0.40µM [20] DYRK1A: 3.60µM [20] ERK2: 3.60µM [20]
CVT-313 [6-(4-Methoxybenzylamino)-2-(bis-(2-Hydroxyethylamino))-9-isopropylpurine]		ALX-270-393-M001 ALX-270-393-M005	1 mg 5 mg	4.2µM (B) [21]	500nM (A) [21] 500nM (E) [21]	215µM (D1) [21]			
9-Cyanopaullone [9-Cyano-7,12-dihydroindolo-[3,2- <i>d</i>]1]benzazepin-6(5 <i>H</i>)-one]		ALX-270-282-M001 ALX-270-282-M005	1 mg 5 mg	24nM (B) [22, 23]			44nM (p25) [2]	10nM (β) [2]	
Cyclin-dependent Kinase 2 Inhibitor [CDK2/Cyclin Inhibitor]		ALX-153-012-M001 ALX-153-012-M005	1 mg 5 mg		<100nM (E) [24]				
Cyclin-dependent Kinase 2 Inhibitory Peptide I [CDK2/Cyclin Inhibitory Peptide I]		ALX-153-041-C500	500 µg						
Cyclin-dependent Kinase 2 Inhibitory Peptide II [CDK2/Cyclin Inhibitory Peptide II]		ALX-153-042-C500	500 µg						
Debromohymenialdisine [DBH; 4-(2-Amino-4-oxo-2-imidazolin-5-ylidene)-4,5,6,7-tetrahydropyrrolo[2,3- <i>c</i>]azepin-6-one]		ALX-350-290-C100	100 µg					Chk1: 3µM [26] Chk2: 3.5µM [26] MEK1: 824nM [27]	
6-Dimethylaminopurine [6-DMAP; N ⁶ ,N ⁶ -Dimethyladenine]		ALX-480-050-M100	100 mg	120µM (B) [28]			120µM (p35) [28]		Non-selective protein kinase inhibitor.
Elbfluorene [3-Acetyl-6-hydroxy-4-phenylbenzo[4,5]furo[2,3- <i>b</i>]pyridine; 1-Aza-9-oxafluorene 5b]		ALX-270-389-M001 ALX-270-389-M005	1 mg 5 mg	4.2µM (B) [8]	>100µM (E) [8]	>100µM (D1) [8]	15% inhibition at 10µM [8]		
Fascaplysin		ALX-270-300-M001 ALX-270-300-M005	1 mg 5 mg	>100µM (B) [29]	>50µM (A) [29] >100µM (D2) [29] >100µM (D3) [29]	350nM (D1) [29] 20µM (p35) [29]	20µM (p35) [29]	CDK6 (D1): 3.4µM [29] CDK6 (D2): 700nM [32] v-abl: >10µM [29] c-met: >10µM [29] EGF-IR: >10µM [29] IRTK: >10µM [29]	
Flavopiridol [L-86-8276; NSC-649890]		ALX-430-161-M005 ALX-430-161-M025	5 mg 25 mg	400nM [30]	100nM [31]	[31]			
102-Hymenialdisine [4-(2-Amino-4-oxo-2-imidazolin-5-ylidene)-2-bromo-4,5,6,7-tetrahydropyrrolo[2,3- <i>c</i>]azepin-8-one]		ALX-350-289-C500	500 µg	22nM (B) [32]	70nM (A) [32] 40nM (E) [32]	600nM (D1) [32]	28nM (p25) [32]	10nM (β) [32] 130nM (β) [33]	CDK3: 100nM /cyclin E [32] CDK6 (D2): 700nM [32] CK1: 35nM [32] erk1: 470nM [32] erk2: 2µM [32] JNK: 8.5µM [32] MAPK2: 12µM [32] MARK2: 670nM [33] MEK1: 6nM [27]

Inhibitor	Structure	Prod. No.	Size	CDK1	CDK2	CDK4	CDK5	GSK-3	Other Enzymes
Hymenialdin [2-Debromopaullone]		ALX-350-291-M001	1 mg	>100µM (B) [32]			4µM (p25) [32]	12µM (β) [32]	CK1: >100µM [32]
Indirubin		ALX-270-361-M001 ALX-270-361-M005	1 mg 5 mg	10µM (B) [34]	2.2µM (A) [34] 7.5µM (E) [34]	12µM (D1) [34]	5.5µM (p35) [34]	600nM (β) [34]	CK1: 8.5µM [34] CK2: >100µM [34] erk1: >100µM [34] erk2: 43µM [34] JNK: >100µM [34] MAPK: >100µM [34] c-Raf: >10µM [34] c-Src: 18µM [34] Non-kinases: 8-Lactamase: 20µM [36] Chymotrypsin: 100µM [36] Malate Dehydrogenase: 30µM [36]
Indirubin-5-sulfonic acid · Na [E226]		ALX-270-296-M001 ALX-270-296-M005	1 mg 5 mg	55nM (B) [34]	35nM (A) [34] 150nM (E) [34]	300nM (D1) [34]	65nM (p35) [34]	280nM (β) [35]	CK1: 10µM [34] CK2: 1.5µM [34] erk1: 38µM [34] erk2: >100µM [34] JNK: 5.2µM [34] MAPK: 3µM [34] PKC b2: 6.5µM [34] c-Raf: 5.5µM [34] c-Src: 3.8µM [34]
Indirubin-3'-monoxime		ALX-270-271-M001 ALX-270-271-M005	1 mg 5 mg	180nM (B) [34]	590nM (A) [3] 440nM (A) [34] 250nM (E) [34]	3.33µM (D1) [34]	100nM (p35) [34]	190nM (β) [3] 22nM (β) [35]	AMPK: 220nM [3] CK1: 9µM [34] CK2: 12µM [34] erk1: >100µM [34] erk2: >100µM [34] JNK: NT LCK: 3.80nM [3] MAPK: >100µM [34] p55fgf: 78µM [28] p55ck: 50µM [28]
5-Iodoindirubin-3'-monoxime		ALX-270-424-M001	1 mg	25nM (B) [35]			20nM (p25) [35]	9nM (β) [35]	
N-6-(Δ²-Isopentenyl)-adenine [6-(Δ ² -Dimethylallylamino)purine; Triacanthine]		ALX-350-034-M100	100 mg	45µM (B) [28]	50µM (A) [28] 200µM (D1) [28]		80µM (p35) [28]		CDK6 (D3): >100µM [28] erk1: 90µM [28] IRTK: 140µM [28] p55fgf: 78µM [28] p55ck: 50µM [28]
Kempaulone [9-Bromo-7,12-dihydroindolo-[3,2- <i>d</i>]1]benzazepin-6(5 <i>H</i>)-one]		ALX-270-274-M001 ALX-270-274-M005	1 mg 5 mg	400nM (B) [2]	670nM (A) [3] 7.5µM (E) [37]	>100µM (D1) [37]	850nM (p25) [37] 850nM (p35) [37]	23nM (β) [2] 230nM (β) [3]	CK2: 20µM [37] erk1: 20µM [37] erk2: 9µM [37] LCK: 470nM [3, 37] c-Raf: 38µM [37] c-Src: 15µM [37]
NU2058 [O ⁶ -Cyclohexylmethylguanine]		ALX-270-394-M005 ALX-270-394-M025	5 mg 25 mg	5µM (B) [38]	12µM (A) [38]				
NU6102 [O ⁶ -Cyclohexylmethyl-2-(4'-sulfamoylanilino)purine]		ALX-270-419-M001 ALX-270-419-M005	1 mg 5 mg	9.5nM (B) [39]	5.4nM (A3) [39]	1.6µM (D1) [39]			DYRK1A: 0.9µM [39] PDK1: 0.8µM [39] ROCK1: 0.6µM [39]
NU6140 [4-(Cyclohexylmethoxy-9 <i>H</i> -purin-2-ylamino)-N,N-diethylbenzamide]		ALX-270-441-M005	5 mg	6.6µM (B) [40]	0.41µM (A) [40]	5.5µM (D) [40]	15µM (p25) [40]		CDK7 (H): 3.9µM [40]
Olomoucine (high purity) [6-Benzylamino-2-(2-hydroxyethylamino)-9-methylpurine]		ALX-350-013-M005 ALX-350-013-M025	5 mg 25 mg	50µM (A) [28] 7µM (B) [28] 1µM (E) [28]	7µM (A) [28] 7µM (E) [28]	>1M (D1) [28]	3µM (p25) [28] 3µM (p35) [28]	100µM (β) [35]	CDK6 (D3): >250µM [28] erk1: 50µM [42] erk2: 40µM [42] IRTK: 400µM [42]
Olomoucine II [6-(2-Hydroxybenzylamino)-2(R)-[1-(hydroxymethyl)-propylamino]-9-isopropylpurine]		ALX-270-396-M001 ALX-270-396-M005	1 mg 5 mg	20nM (B) [43]					
Olomoucine, Iso- [6-Benzylamino-2-(2-hydroxyethylamino)-7-methylpurine]		ALX-350-090-M005 ALX-350-090-M025	5 mg 25 mg	>500µM (B) [28]		>1M (D1) [28]	>1M (p35) [28]		
Olomoucine, N⁶-Isopropyl- [6-Benzylamino-2-(2-hydroxyethylamino)-9-isopropylpurine]		ALX-270-397-M001 ALX-270-397-M005	1 mg 5 mg	2µM (B) [28]			3µM (p35) [28]		
Olomoucine, Dimethylamino- [6-Benzylamino-2-(2-dimethylaminoethylamino)-9-methylpurine]		ALX-350-054-M005	5 mg	>100µM (B) [28]			140µM (p35) [28]		
PNU 112455A · HCl [N-(6-Aminopyrimidin-4-yl)sulfanilamide · HCl]		ALX-270-398-M001 ALX-270-398-M005	1 mg 5 mg		K _i =2µM (E) [44]		K _i =2µM (p25) [44]		erk2: >100µM [44]
Purvalanol A [6-(3-Chloroanilino)-2-(1 <i>R</i>)-(isopropyl-2-hydroxyethylamino)-9-isopropylpurine]		ALX-270-246-M001 ALX-270-246-M005	1 mg 5 mg	4nM (B) [19]	70nM (A) [19] 100nM (A) [3] 35nM (E) [19]	850nM (D1) [19]	75nM (p35) [19]	>10µM (β) [19]	CK1: >3.3µM [19] CK2: >10µM [19] DYRK1A: 300nM [3] erk1: 9µM [19] IRTK: 5µM [19] MLCK: 90µM [42] PKC: >100µM [42] c-Src: 250µM [42]
RO-3306 [(5 <i>Z</i>)-2-(Thiophen-2-yl)methylamino)-5-(quinolin-6-yl)methylene]thiazol-4(5 <i>H</i>)-one]		ALX-270-463-M001 ALX-270-463-M005	1 mg 5 mg		K _i =20nM [45]				
(R)-Roscovitine [2 <i>R</i>]-[2-Ethyl-2-hydroxyethylamino)-6-benzylamino-9-isopropylpurine]		ALX-350-251-M001 ALX-350-251-M005 ALX-350-251-M050	1 mg 5 mg 50 mg	200nM [41]	700nM (A) [42] 250nM (A) [3] 700nM (E) [42]	>100µM (D1) [42]	160nM (p35) [42]	130µM [46]	CDK6 (D3): >100µM [D3] [42] CK1: 17µM [3] DYRK1A: 3.1µM [3] erk1: 34µM [42] erk2: 14µM [42] IRTK: 70µM [42] MLCK: 90µM [42] PKC: >100µM [42] c-Src: 250µM [42]
(S)-Roscovitine [2 <i>S</i>]-[2-Ethyl-2-hydroxyethylamino)-6-benzylamino-9-isopropylpurine]		ALX-350-293-M001	1 mg	800nM [41]					
Scytonemin [SCY]		ALX-350-376-M001	1 mg	3.0µM [47]					PKCβ2: 2.7µM [47] Myt1: 1.2µM [47] Chk1: 1.4µM [47] PLK1: 2.0µM [47]
Staurosporine		ALX-380-014-C100 ALX-380-014-C250 ALX-380-014-M005	100 µg 250 µg 5 mg	6nM (B) [48] 5nM (B) [35]	7nM (A) [48]	<10µM (D1) [48]	4nM (p25) [35]	15nM (β) [35]	Non-selective protein kinase inhibitor. Chk1: 8nM [49] PKC: 5nM [49]
SU9516 [3-[1-(3 <i>H</i>)-Imidazo[4-yl]-meth-(<i>Z</i>)-ylidene]-5-methoxy-1,3-dihydro-indol-2-one]		ALX-270-400-M005 ALX-270-400-M025	5 mg 25 mg	40nM (B) [50]	22nM (A) [50]	200nM (D1) [50]			EGFR: >100µM [50] FLK1: 130nM [51] p38: >10µM [50] PDGFR: 18µM [50] PKC: >10µM [50] c-Src: 11µM [51]
UCN-01 [7-Hydroxystaurosporine]		ALX-380-222-MC25 ALX-380-222-M001	0.25 mg 1 mg	31nM [52]	42nM [52]	32nM [52]			CDK6: 58nM [52]

LITERATURE REFERENCES:

[1] Aloisine, a new family of CDK/GSK-3 inhibitors. SAR study, crystal structure in complex with CDK2, enzyme selectivity, and cellular effects: Y. Mettrey, et al.; J. Med. Chem. **46**, 222 (2003). • [2] Paulones are potent inhibitors of glycogen synthase kinase-3beta and cyclin-dependent kinase 5alpha. M. Leost, et al.; Biol. Biochem. **267**, 5983 (2000). • [3] The specificities of protein kinase inhibitors: an update: J. Bain, et al.; Biochem. J. **371**, 199 (2003). • [4] Pyrazolo[3,3-*b*]quinoxalines. A new class of cyclin-dependent kinases inhibitors: M.A. Ortega, et al.; Bioorg. Med. Chem. **10**, 2177 (2002). • [5] Synthesis and application of functionally diverse 2,6,9-trisubstituted purine libraries as CDK inhibitors: Y.T. Chang, et al.; Chem. Biol. **6**, 361 (1999). • [6] Synthesis of quinolinyloquinolinyllalpyrrolo [3,4-*c*] carbazoles as cyclin D1/CDK4 inhibitors: G. Zhu, et al.; Bioorg. Med. Chem. Lett. **13**, 1231 (2003). • [7] The p16 status of tumor cell lines identifies small molecule inhibitors specific for cyclin-dependent kinase 4: A. Kubo, et al.; Clin. Cancer Res. **5**, 4279 (1999). • [8] Evaluation of the first cytostatically active 1-aza-9-oxafluorenes as novel selective CDK1 inhibitors with a glycoprotein modulating properties: K. Brachwitz, et al.; J. Med. Chem. **46**, 876 (2003). • [9] Discovery and SAR of 2-aminothiazole inhibitors of cyclin-dependent kinase 5/p25 as a potential treatment for Alzheimer's disease: C.J. Heisl, et al.; Bioorg. Med. Chem. Let