

SUPPLEMENTARY MATERIAL

corresponding to:

A screen of kinase inhibitors reveals a potential role of Chk1 in regulating *Hydra* head regeneration and maintenance

YUNJIN LEE, VARUN MUDDALURU, SHIRAZ ANWAR, JOANNA YVONNE WILSON and ANA REGINA CAMPOS

Chk1 (HYDRA)	1 VFEPFVEGWDFVETLGE <ins>GAYGEVRLAINRKTQEAVAVKIVNADKLAGNKD</ins>	50
Chk1 (HUMAN)	1 MAVPFVEDWDLVQTLGE <ins>GAYGEVQLAVNRVTEAVAVKIVDMKRAVDCPE</ins>	50
Chk1 (HYDRA)	51 <ins>CLKKEVCIHKMLQHSIIKFYGQRTEKDRVYLFLEYAAGGELFDRIEPDV</ins>	100
Chk1 (HUMAN)	51 NIKKEICINKMLNHENVVFKFYGHREGNIQYL <ins>FLEYCSGGELFDRIEPDI</ins>	100
Chk1 (HYDRA)	101 <ins>GMPIPQACRYFKQLINGLEYIHSKGVTHRDIKPENILLVDGNLKITDFG</ins>	150
Chk1 (HUMAN)	101 GMPEPDAQRFFHQLMAGVVYLHGIGITHRDIKPENLLLDERDNLKISDFG	150
Chk1 (HYDRA)	151 <ins>LSTVFRYKDVERLLERCCGTPPYVAPEVLQKKEYKAEPAYIWSCGIVLTA</ins>	200
Chk1 (HUMAN)	151 LATVFRYNNRERLLNPKMCGTLPYVAPELLKRREFHAEPVDVWSCGIVLTA	200
Chk1 (HYDRA)	201 <ins>MLAGELPWDEPIESCKEYLDWSHSKLIHTPWNKLNTTSIGFLKKLLHPVP</ins>	250
Chk1 (HUMAN)	201 MLAGELPWDPQPSDQCQEYSDWKEKKTLYLNPKKIDSAPLALLHKILVENP	250
Chk1 (HYDRA)	251 <ins>SKRYTIAEIKKDKWFNG--NYGSILTKSPLNGLTNFETSSLKKHCSS--D</ins>	296
Chk1 (HUMAN)	251 SARITIPDIKKDRWYNKPLKKGAKRPRVTSGGVSE-SPSGFSKHIQSNL	299
Chk1 (HYDRA)	297 RSVSTSVNSKLVSNFSSSQPIPSC-----TSSDYELQEIREEQQGIWYS	340
Chk1 (HUMAN)	300 FSPVNSASSEENVKYSSSQPEPRTGLSLWDTSPSY----IDKLVQGISFS	345
Chk1 (HYDRA)	341 QPVNIEDMLL-SQISLTPGSSQNPMALAKRMTRFNLSLTLEAVKKLSS	389
Chk1 (HUMAN)	346 QPTCPDHMLLNSQLLGTGSSQNPWQRLVKRMTRFFTKLDAKSYQCLKE	395
Chk1 (HYDRA)	390 TLKELSFQYKIVSLNQIRITS HDRRKHTLTYL TNLIEINQRPLLVDFRLS	439
Chk1 (HUMAN)	396 TCEKLGYQWKKSCMNQVTISTDRRNNKLIFKVNLLEMDDK-ILVDFRLS	444
Chk1 (HYDRA)	440 KGDGLEFKRQFKTIKGLLCQYVV*-----	463
Chk1 (HUMAN)	445 KGDGLEFKRHFLKIKGKLIDIVSSQKVWLPAT	476

Fig. S1. Global pairwise alignment of Chk1 (*Hydra*) and Chk1 (Human) amino acid sequences. A global pairwise alignment of Chk1 (*H. vulgaris*) and Chk1 (Human) amino acid sequences was performed using EMBOSS Needle. The catalytic domain and critical residues of the ATP binding pocket are represented by orange and yellow, respectively.

TABLE S1

ORGANIZATION OF THE 80 KINASE INHIBITORS BY THEIR TARGET SIGNAL TRANSDUCTION PATHWAY

Signal Transduction Pathway	Target Kinase	Cat. No.	Product Name	Signal Transduction Pathway	Target Kinase	Cat. No.	Product Name
Cell Cycle Regulation	CDK	1937	NSC 693868	DDR	Chk1	2560	SB 218078
	CDK	2072	Aminopurvalanol A		Chk1	2694	PD 407824
	CDK	2457	Arcyriafavin A		ATR/ATM	2639	CGK 733
	CDK	1284	Olomoucine		ATM	3544	KU 55933
	CDK	1580	Purvalanol A		DNA-PK	2828	NU 7026
	CDK	1581	Purvalanol B		DNA-PK	3271	Compound 401
	CDK	2609	Ryuvidine		EGFR	0414	AG 490
	Aurora	2458	ZM 447439		EGFR	1110	Genistein
	PLK	2977	GW 843682X		EGFR	2239	GW 583340 dihydrochloride
					EGFR	2416	BIBX 1382 dihydrochloride
MAPK/ERK	P38 MAPK	1962	SB 239063		EGFR	3000	Iressa
	P38 MAPK	1264	SB 202190	PI3K-AKT/PKB	PKB	2926	FPA 124
	P38 MAPK	1402	SB 203580 hydrochloride		PKB	2151	API-2
	P38 MAPK	2908	EO 1428		PKB	2558	10-DEBC hydrochloride
	MEK	1213	PD 98059		FLT3	2591	TCS 359
	MEK	1969	SL 327		PI3K	2814	PI 828
	MEK	1144	U0126		PI3K	1130	LY 294002 hydrochloride
	MEK	2605	PD 198306		SGK	3572	GSK 650394
	MEK	1777	Arctigenin	VEGF	VEGFR	1459	SU 4312
	Mnk1	2731	CGP 57380		VEGFR	2475	ZM 323881 hydrochloride
DAG-PKC	Raf	1321	ZM 336372		VEGFR	2499	ZM 306416 hydrochloride
	Raf	1381	GW 5074		VEGFR	2542	KI 8751
	PKC	0741	GF 109203X		VEGFR	3037	SU 5416
JAK/STAT	PKC	2002	Ro 31-8220 mesylate	Wnt	GSK-3	1616	SB 216763
	PKC	2442	CGP 53353		GSK-3	1617	SB 415286
	JAK3	1366	ZM 449829		GSK-3	3194	BIO
NF-κB	JAK3	1367	ZM 39923 hydrochloride		CK1	2902	D 4476
	JAK2	2291	1,2,3,4,5,6-Hexabromocyclohexane		CK2	3194	TBB
	IKK	2539	IKK 16	BCR	BKT	1300	LFM-A13
	IKK	2559	TPCA-1		BKT	1405	(-)Terreic acid
ROCK/MLCK	IKK	2611	IMD 0354	cAMP	PKA	2910	H 89 dihydrochloride
	IKK	3318	SC 514		CaM Kinase III	3439	NH 125
	MLCK	0431	ML 9 hydrochloride		FceRI	2417	ER 27319 maleate
	ROCK	0541	Fasudil hydrochloride		HGF/cMET	2693	PHA 665752
TGF-β	ROCK	1254	Fasudil hydrochloride		IGF-1R	2768	PQ 401
	ROCK	2415	HA 1100 hydrochloride	JNK/SAPK	JNK	1496	SP 600125
	TGFbR1	1614	SB 431542		JNK	3314	BI 78D3
SRF	TGFbR1	2718	LY 364947	NGF	TrkA	2238	GW 441756
	TGFbR1	3269	SD 208		TrkA	2272	Ro 08-2750
	Src	1397	PP 1				
	Src	1407	PP 2				
	Src	3063	1-Naphthyl PP1				

Column 1 is the signal transduction pathways targeted by the kinase inhibitors; column 2 is the main target kinase within the signaling pathway; column 3 is the catalogue number (Cat. No.); column 4 is the product name of the kinase inhibitor. Abbreviations for signal transduction pathways: MAPK/ERK = mitogen-activated protein kinase/extracellular signal-regulated kinase, DAG-PKC = diacylglycerol-protein kinase C, JAK/STAT = Janus kinase/signal transducers and activators of transcription, NF-κB = nuclear factor kappa-light-chain-enhancer of activated B cells, ROCK/MLCK = Rho-associated kinase/myosin light chain kinase, TGF-β = transforming growth factor-β, SRF = serum response factor, DDR = DNA damage response, EGFR = epidermal growth factor receptor, PI3K-PKB = phosphoinositide-3-kinase-protein kinase B, VEGF = vascular endothelial growth factor, BCR = B cell antigen receptor, FcεRI = high affinity IgE receptor, HGF/cMET = hepatocyte growth factor, IGF-1R = insulin-like growth factor 1, JNK/SAPK = c-Jun NH2-terminal kinase/stress-activated protein kinase, NGF = nerve growth factor. Abbreviations for target kinases: Cdk = cyclin-dependent kinase, PLK = polo-like kinase, P38 MAPK = P38 mitogen-activated protein kinase, MEK = mitogen-activated protein kinase kinase, Mnk1 = MAP kinase-interacting serine/threonine-protein kinase 1, JAK3 = Janus kinase 3, JAK2 = Janus kinase 2, IKK = IκB kinase, TGFbR1 = transforming growth factor-beta receptor type 1, ATM = ataxia-telangiectasia, mutated, ATR = ATM and Rad3-related, DNA-PK = DNA-dependent protein kinase, FLT3 = fms like tyrosine kinase 3, SGK = serum- and glucocorticoid-inducible protein kinase, VEGFR = vascular endothelial growth factor receptor, GSK-3 = glycogen synthase kinase 3, CK1 = casein kinase 1, CK2 = casein kinase 2, BKT = Bruton's tyrosine kinase, PKA = protein kinase A, Syk = spleen tyrosine kinase, TrkA = tropomyosin receptor kinase A.