

A

Mco	MKLFCICILALAATAMA-EESNVVD-LIASSSNEVPTANMEQEVVKAKPGENVVLSAFSVLSPLAQISSLASGESHDEILKAIGLPNDNVTKEVFTDVSQQLRSVKGVELRLANKVYVROGAVLNDEFAAV	126
Bmo	MRTICLICIFTIAAMA-AVTNLNSVLRNGNDNTARMPTEVVNPNGPSVVLSAFSVTPPLAQISSLASGETHEELLKAIIFPDDAIRTEFASKSRDLSSIKGVBLKMANXVYVHDGGKIDENFAAV	127
Mse	MKIIMCIFGLAALAMA-GETDILQKLLRESNDQTAAOMSFSEVVKANPVSQCONVVLSAFSVTPPLGQISSLASGETHEELLKAIIFPDDAIRTEFASKSRDLSSIKGVBLKMANXVYVHDGGKIDENFAAV	127
Pxy	MKLLICLICVLSVVATMADIEIPFKQOINNGNDCTTANMFSEVLPQONGKNNVVLAFSVPITPVQAQISLASGETHEELLKTIIGLPSDEAIKSVEPLVNAKLELIGKVILKQASRIVYLPLTAEIADDFAAV	128
Cpo	MNTLICLICVLSVVATMADIEIPFKQOINNGNDCTTANMFSEVLPQONGKNNVVLAFSVPITPVQAQISLASGETHEELLKTIIGLPSDEAIKSVEPLVNAKLELIGKVILKQASRIVYLPLTAEIADDFAAV	125

Mco	SKDVENSDVKNVDFTKNV <span style="background-color: green;">EAAKEINENVE</span> NTNHKIKDLVSSLDAS <span style="background-color: green;">TA</span> AVLVNAINYFKGKWKPFDESATR <span style="background-color: green;">IDFF</span> VTKDQP <span style="background-color: blue;">PIK</span> KPT <span style="background-color: green;">HKSGDF</span> FKYAESKELDAKLLELPYEGDOSSL <span style="background-color: green;">LIVLPN</span> EI	254
Bmo	SRDVENS <span style="background-color: green;">D</span> VQNIDFSKNTV <span style="background-color: green;">AA</span> NVE <span style="background-color: green;">NT</span> NR <span style="background-color: green;">I</span> KDLVNPDSLSSA <span style="background-color: green;">TAA</span> VLVNAIYFKGKANSSKFDERLTS <span style="background-color: green;">S</span> DFYVSKDKT <span style="background-color: green;">I</span> KVPM <span style="background-color: green;">Y</span> KRGDYKY <span style="background-color: green;">G</span> ESAA <span style="background-color: green;">L</span> IN <span style="background-color: green;">Q</span> LIEIPY <span style="background-color: green;">K</span> G <span style="background-color: green;">D</span> OSSL <span style="background-color: green;">LIVLP</span> KDK	255
Mse	SRDVENS <span style="background-color: green;">C</span> E <span style="background-color: green;">K</span> EVONVDFVKNVE <span style="background-color: green;">AA</span> GAN <span style="background-color: green;">K</span> N <span style="background-color: green;">D</span> QTN <span style="background-color: green;">T</span> NR <span style="background-color: green;">I</span> KLN <span style="background-color: green;">L</span> V <span style="background-color: green;">P</span> D <span style="background-color: green;">A</span> I <span style="background-color: green;">DET</span> TRSV <span style="background-color: green;">L</span> VLVNAIYFKGK <span style="background-color: green;">W</span> DKFVKERT <span style="background-color: green;">M</span> DFH <span style="background-color: green;">V</span> SKDKT <span style="background-color: green;">I</span> KVPT <span style="background-color: green;">M</span> IGKD <span style="background-color: green;">V</span> R <span style="background-color: green;">Y</span> ADVE <span style="background-color: green;">ELDAKMIEN</span> S <span style="background-color: green;">Y</span> EG <span style="background-color: green;">G</span> DA <span style="background-color: green;">S</span> MLILP <span style="background-color: green;">N</span> O	255
Pxy	SRDVENS <span style="background-color: green;">E</span> AKRNLDFT <span style="background-color: green;">K</span> T <span style="background-color: green;">N</span> V <span style="background-color: green;">E</span> SA <span style="background-color: green;">K</span> E <span style="background-color: green;">I</span> ND <span style="background-color: green;">V</span> NE <span style="background-color: green;">E</span> Q <span style="background-color: green;">T</span> NR <span style="background-color: green;">I</span> Q <span style="background-color: green;">L</span> V <span style="background-color: green;">D</span> PD <span style="background-color: green;">S</span> I <span style="background-color: green;">K</span> PD <span style="background-color: green;">A</span> SL <span style="background-color: green;">L</span> VLVNAIYFKG <span style="background-color: green;">T</span> WS <span style="background-color: green;">K</span> FD <span style="background-color: green;">N</span> AST <span style="background-color: green;">T</span> E <span style="background-color: green;">D</span> F <span style="background-color: green;">H</span> V <span style="background-color: green;">S</span> K <span style="background-color: green;">D</span> K <span style="background-color: green;">T</span> K <span style="background-color: green;">V</span> K <span style="background-color: green;">P</span> TM <span style="background-color: green;">Y</span> Q <span style="background-color: green;">K</span> AS <span style="background-color: green;">F</span> Y <span style="background-color: green;">G</span> E <span style="background-color: green;">S</span> A <span style="background-color: green;">E</span> ELDAKL <span style="background-color: green;">L</span> LELPY <span style="background-color: green;">E</span> GE <span style="background-color: green;">K</span> AS <span style="background-color: green;">F</span> L <span style="background-color: green;">L</span> ILP <span style="background-color: green;">N</span> I	256
Cpo	SKS <span style="background-color: green;">V</span> FC <span style="background-color: green;">S</span> E <span style="background-color: green;">K</span> RNIDFT <span style="background-color: green;">K</span> T <span style="background-color: green;">N</span> V <span style="background-color: green;">P</span> A <span style="background-color: green;">Q</span> E <span style="background-color: green;">I</span> N <span style="background-color: green;">S</span> N <span style="background-color: green;">V</span> ED <span style="background-color: green;">H</span> T <span style="background-color: green;">N</span> H <span style="background-color: green;">K</span> I <span style="background-color: green;">K</span> D <span style="background-color: green;">L</span> V <span style="background-color: green;">D</span> P <span style="background-color: green;">N</span> SL <span style="background-color: green;">G</span> AD <span style="background-color: green;">T</span> RA <span style="background-color: green;">L</span> VLVNAIYFKG <span style="background-color: green;">S</span> WE <span style="background-color: green;">K</span> FN <span style="background-color: green;">D</span> KAT <span style="background-color: green;">T</span> DS <span style="background-color: green;">D</span> F <span style="background-color: green;">H</span> V <span style="background-color: green;">T</span> K <span style="background-color: green;">D</span> K <span style="background-color: green;">T</span> K <span style="background-color: green;">V</span> K <span style="background-color: green;">P</span> TM <span style="background-color: green;">Y</span> KK <span style="background-color: green;">D</span> FKYAESKELDAKL <span style="background-color: green;">L</span> LELPY <span style="background-color: green;">E</span> GF <span style="background-color: green;">K</span> AS <span style="background-color: green;">F</span> L <span style="background-color: green;">L</span> ILP <span style="background-color: green;">N</span> I	253

Mco	DGIGS <span style="background-color: green;">T</span> VE <span style="background-color: green;">K</span> KL <span style="background-color: green;">D</span> PTA <span style="background-color: green;">S</span> KA <span style="background-color: green;">V</span> ENMFY <span style="background-color: green;">N</span> V <span style="background-color: green;">D</span> LP <span style="background-color: green;">K</span> FK <span style="background-color: green;">I</span> E <span style="background-color: green;">T</span> T <span style="background-color: green;">D</span> DL <span style="background-color: green;">K</span> AV <span style="background-color: green;">L</span> KK <span style="background-color: green;">N</span> I <span style="background-color: green;">V</span> L <span style="background-color: green;">F</span> T <span style="background-color: green;">G</span> EAR <span style="background-color: green;">R</span> NN <span style="background-color: green;">L</span> IK <span style="background-color: green;">G</span> ES <span style="background-color: green;">D</span> LF <span style="background-color: green;">I</span> FT <span style="background-color: green;">T</span> DA <span style="background-color: green;">I</span> Q <span style="background-color: green;">K</span> AF <span style="background-color: green;">I</span> TD <span style="background-color: green;">V</span> NE <span style="background-color: green;">E</span> GA <span style="background-color: green;">A</span> AAA <span style="background-color: green;">A</span> AN	351
Bmo	DG <span style="background-color: green;">T</span> Q <span style="background-color: green;">I</span> PE <span style="background-color: green;">A</span> EL <span style="background-color: green;">K</span> D <span style="background-color: green;">P</span> K <span style="background-color: green;">T</span> E <span style="background-color: green;">T</span> E <span style="background-color: green;">A</span> Q <span style="background-color: green;">D</span> SMY <span style="background-color: green;">S</span> Y <span style="background-color: green;">D</span> V <span style="background-color: green;">L</span> Y <span style="background-color: green;">L</span> PK <span style="background-color: green;">K</span> FI <span style="background-color: green;">E</span> T <span style="background-color: green;">T</span> ET <span style="background-color: green;">D</span> LN <span style="background-color: green;">K</span> D <span style="background-color: green;">V</span> L <span style="background-color: green;">S</span> N <span style="background-color: green;">M</span> V <span style="background-color: green;">N</span> I <span style="background-color: green;">F</span> NN <span style="background-color: green;">D</span> -DAQ <span style="background-color: green;">I</span> TR <span style="background-color: green;">L</span> LK <span style="background-color: green;">G</span> ES <span style="background-color: green;">S</span> -L <span style="background-color: green;">V</span> SE <span style="background-color: green;">A</span> I <span style="background-color: green;">Q</span> KAF <span style="background-color: green;">I</span> E <span style="background-color: green;">T</span> NE <span style="background-color: green;">E</span> GA <span style="background-color: green;">A</span> AAA <span style="background-color: green;">A</span> AN	350
Mse	D <span style="background-color: green;">G</span> I <span style="background-color: green;">T</span> A <span style="background-color: green;">P</span> E <span style="background-color: green;">A</span> EL <span style="background-color: green;">K</span> D <span style="background-color: green;">P</span> K <span style="background-color: green;">T</span> E <span style="background-color: green;">T</span> E <span style="background-color: green;">A</span> Q <span style="background-color: green;">D</span> SMY <span style="background-color: green;">S</span> Y <span style="background-color: green;">D</span> V <span style="background-color: green;">L</span> Y <span style="background-color: green;">L</span> PK <span style="background-color: green;">K</span> FI <span style="background-color: green;">E</span> T <span style="background-color: green;">T</span> ET <span style="background-color: green;">D</span> LN <span style="background-color: green;">K</span> D <span style="background-color: green;">V</span> L <span style="background-color: green;">S</span> N <span style="background-color: green;">M</span> V <span style="background-color: green;">N</span> I <span style="background-color: green;">F</span> NN <span style="background-color: green;">D</span> -DAQ <span style="background-color: green;">I</span> TR <span style="background-color: green;">L</span> LK <span style="background-color: green;">G</span> ES <span style="background-color: green;">S</span> -L <span style="background-color: green;">V</span> SE <span style="background-color: green;">A</span> I <span style="background-color: green;">Q</span> KAF <span style="background-color: green;">I</span> E <span style="background-color: green;">T</span> NE <span style="background-color: green;">E</span> GA <span style="background-color: green;">A</span> AAA <span style="background-color: green;">A</span> AN	352
Pxy	E <span style="background-color: green;">G</span> LP <span style="background-color: green;">A</span> TE <span style="background-color: green;">K</span> E <span style="background-color: green;">K</span> LP <span style="background-color: green;">E</span> P <span style="background-color: green;">Q</span> H <span style="background-color: green;">N</span> K <span style="background-color: green;">A</span> LS <span style="background-color: green;">M</span> SE <span style="background-color: green;">Y</span> K <span style="background-color: green;">G</span> E <span style="background-color: green;">I</span> Q <span style="background-color: green;">V</span> Y <span style="background-color: green;">L</span> PK <span style="background-color: green;">K</span> FI <span style="background-color: green;">E</span> T <span style="background-color: green;">T</span> ET <span style="background-color: green;">D</span> LN <span style="background-color: green;">K</span> E <span style="background-color: green;">V</span> M <span style="background-color: green;">S</span> G <span style="background-color: green;">L</span> SS <span style="background-color: green;">F</span> EP <span style="background-color: green;">K</span> AT <span style="background-color: green;">D</span> K <span style="background-color: green;">L</span> KK <span style="background-color: green;">S</span> K <span style="background-color: green;">A</span> PL <span style="background-color: green;">Y</span> I <span style="background-color: green;">G</span> DA <span style="background-color: green;">K</span> OK <span style="background-color: green;">A</span> K <span style="background-color: green;">I</span> E <span style="background-color: green;">V</span> NE <span style="background-color: green;">E</span> CS <span style="background-color: green;">E</span> AAA <span style="background-color: green;">A</span> AN	353
Cpo	D <span style="background-color: green;">G</span> LP <span style="background-color: green;">A</span> TE <span style="background-color: green;">K</span> E <span style="background-color: green;">K</span> LP <span style="background-color: green;">E</span> P <span style="background-color: green;">Q</span> H <span style="background-color: green;">N</span> K <span style="background-color: green;">A</span> LS <span style="background-color: green;">M</span> SE <span style="background-color: green;">Y</span> K <span style="background-color: green;">G</span> E <span style="background-color: green;">I</span> Q <span style="background-color: green;">V</span> Y <span style="background-color: green;">L</span> PK <span style="background-color: green;">K</span> FI <span style="background-color: green;">E</span> T <span style="background-color: green;">T</span> ET <span style="background-color: green;">D</span> LN <span style="background-color: green;">K</span> E <span style="background-color: green;">V</span> M <span style="background-color: green;">S</span> G <span style="background-color: green;">L</span> SS <span style="background-color: green;">F</span> EP <span style="background-color: green;">K</span> AT <span style="background-color: green;">D</span> K <span style="background-color: green;">L</span> KK <span style="background-color: green;">S</span> K <span style="background-color: green;">A</span> PL <span style="background-color: green;">Y</span> I <span style="background-color: green;">G</span> DA <span style="background-color: green;">K</span> OK <span style="background-color: green;">A</span> K <span style="background-color: green;">I</span> E <span style="background-color: green;">V</span> NE <span style="background-color: green;">E</span> CS <span style="background-color: green;">E</span> AAA <span style="background-color: green;">A</span> AN	350

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B

CpSPN1C	---VFT <span style="background-color: green;">IDR</span> TM <span style="background-color: green;">I</span> I <span style="background-color: green;">P</span> E <span style="background-color: green;">T</span> Y <span style="background-color: green;">L</span> E <span style="background-color: green;">M</span> A <span style="background-color: green;">D</span> R <span style="background-color: green;">P</span> F <span style="background-color: green;">I</span> Y <span style="background-color: green;">L</span> N <span style="background-color: green;">V</span> G <span style="background-color: green;">S</span> S <span style="background-color: green;">I</span> L <span style="background-color: green;">F</span> N <span style="background-color: green;">G</span> S <span style="background-color: green;">Q</span> S	40
CpSPN1D	---D <span style="background-color: green;">I</span> G <span style="background-color: green;">Y</span> V <span style="background-color: green;">M</span> MLK <span style="background-color: green;">P</span> E <span style="background-color: green;">I</span> W <span style="background-color: green;">-</span> F <span style="background-color: green;">K</span> A <span style="background-color: green;">D</span> R <span style="background-color: green;">P</span> F <span style="background-color: green;">I</span> E <span style="background-color: green;">F</span> I <span style="background-color: green;">L</span> N <span style="background-color: green;">A</span> G <span style="background-color: green;">S</span> T <span style="background-color: green;">T</span> L <span style="background-color: green;">F</span> N <span style="background-color: green;">G</span> S <span style="background-color: green;">Q</span> S	39
CpSPN1B	AFQVV <span style="background-color: green;">S</span> RSGKFYYV <span style="background-color: green;">E</span> P <span style="background-color: green;">K</span> F <span style="background-color: green;">N</span> A <span style="background-color: green;">D</span> H <span style="background-color: green;">P</span> F <span style="background-color: green;">I</span> E <span style="background-color: green;">F</span> I <span style="background-color: green;">L</span> N <span style="background-color: green;">S</span> H <span style="background-color: green;">S</span> N <span style="background-color: green;">E</span> L <span style="background-color: green;">F</span> N <span style="background-color: green;">G</span> A <span style="background-color: green;">Y</span> I <span style="background-color: green;">H</span>	43
CpSPN1E	--AF <span style="background-color: green;">I</span> LL <span style="background-color: green;">V</span> E <span style="background-color: green;">E</span> E <span style="background-color: green;">S</span> L <span style="background-color: green;">A</span> LV <span style="background-color: green;">E</span> E <span style="background-color: green;">Y</span> F <span style="background-color: green;">D</span> A <span style="background-color: green;">D</span> H <span style="background-color: green;">P</span> F <span style="background-color: green;">I</span> E <span style="background-color: green;">F</span> I <span style="background-color: green;">L</span> N <span style="background-color: green;">R</span> Q <span style="background-color: green;">G</span> A <span style="background-color: green;">T</span> R <span style="background-color: green;">I</span> L <span style="background-color: green;">F</span> N <span style="background-color: green;">G</span> V <span style="background-color: green;">V</span> N <span style="background-color: green;">F</span>	41
CpSPN1A	-DFG <span style="background-color: green;">V</span> TY <span style="background-color: green;">I</span> L <span style="background-color: green;">S</span> A <span style="background-color: green;">Y</span> V <span style="background-color: green;">K</span> P <span style="background-color: green;">E</span> T <span style="background-color: green;">-</span> Y <span style="background-color: green;">F</span> N <span style="background-color: green;">A</span> D <span style="background-color: green;">D</span> H <span style="background-color: green;">P</span> F <span style="background-color: green;">I</span> E <span style="background-color: green;">F</span> I <span style="background-color: green;">L</span> K <span style="background-color: green;">S</span> G <span style="background-color: green;">E</span> A <span style="background-color: green;">L</span> F <span style="background-color: green;">G</span> V <span style="background-color: green;">Y</span> N <span style="background-color: green;">N</span>	41
CpSPN1F	-VFP <span style="background-color: green;">P</span> V <span style="background-color: green;">M</span> A <span style="background-color: green;">S</span> W <span style="background-color: green;">P</span> P <span style="background-color: green;">R</span> V <span style="background-color: green;">V</span> E <span style="background-color: green;">F</span> I <span style="background-color: green;">A</span> D <span style="background-color: green;">A</span> H <span style="background-color: green;">F</span> I <span style="background-color: green;">I</span> L <span style="background-color: green;">K</span> A <span style="background-color: green;">G</span> D <span style="background-color: green;">S</span> A <span style="background-color: green;">L</span> F <span style="background-color: green;">G</span> G <span style="background-color: green;">V</span> P <span style="background-color: green;">A</span> N	42

PF

Reactive Centre Loop

**Figure S3** Comparison of protein sequences encoded by *serpin1* in several lepidopteran insects. **(A)** Sequence alignment of conserved region of serpin1 proteins in *Mamestra configurata*, *Bombyx mori*, *Manduca sexta*, *Plutella xylostella*, and *Cydia pomonella*. The hinge region preceding the reactive center loop (RCL) showed almost identical residues (in blue background) and similar residues (in green or purple background) were also indicated. **(B)** Alignment of *C. pomonella* serpin1 carboxyl-terminal inhibitor domain variants following the hinge region with the RCL and P1 positions are highlighted in red.