Supporting information:

 Table A. Distribution and characteristics of SNP sites in five eIF4E genes and neighboring regions in 61 cassava accessions

Loci [*]	Total	Up/Downstream	Genes	Introns	mRNA	CDS	Synonymous	Non-
	SNP	(+/- 5kb)						synonymous
elF(iso)4E_me1	986	835	151	92	59	22	9	13
nCBP_me2	666	499	167	127	40	15	13	2
nCBP_me1	858	637	221	147	74	22	7	15
elF(iso)4E_me2	992	780	212	175	37	13	9	4
elF4E_me	886	776	110	68	42	30	13	17
Total	4388	3527	861	609	252	102	51	51

Chron Pos	nosome sition	AM560	Albert	AR37- 80	Kibandameno	Kiroba	MCOL22	Mkombozi	Muzege	Namikonga	Nachinyaya	TME3	TME7	TMS30572	TMS60444
Chr03	25396170	G/G	G/G	G/G	G/G	G/G	G/C	G/G	G/G	G/G	./.	G/G	G/G	G/G	G/G
Chr03	25396238	G/G	G/G	G/G	G/G	G/G	G/C	G/G	G/G	G/G	./.	G/G	G/G	G/C	G/C
Chr03	25396987	A/A	A/A	A/A	A/A	A/A	A/T	./.	A/A	A/A	./.	A/A	A/A	./.	A/T
Chr09	25948588	T/T	C/C	T/C	T/C	C/C	T/C	T/C	T/C	T/C	T/C	C/C	C/C	T/T	C/C
Chr15	3318172	C/C	C/T	C/T	./.	C/T	C/T	C/C	C/T	C/C	T/T	./.	C/C	C/C	C/C
Chr17	20187344	A/A	A/A	A/A	A/T	T/T	A/A	A/A	A/A	A/T	A/A	A/A	A/A	A/T	A/A
Chr17	20187753	T/T	T/T	T/T	т/т	T/C	т/т	т/т	T/T	т/т	T/T	т/т	T/T	T/T	т/т
Chr17	20188018	A/A	A/A	A/A	A/A	A/C	A/A	A/A	A/A	A/A	./.	A/A	A/A	A/A	A/A
Chr17	20188026	C/C	C/C	C/C	C/C	C/G	C/C	C/C	C/C	C/C	./.	C/C	C/C	C/C	C/C
Chr17	20189389	C/C	C/C	C/C	C/C	C/T	C/C	C/C	C/C	C/C	C/C	C/C	C/C	C/C	./.
Chr17	20189407	T/T	T/T	T/T	т/т	T/A	т/т	T/T	T/T	T/T	T/T	т/т	T/T	T/T	./.
Chr17	20189457	T/T	./.	т/т	т/т	T/C	т/т	т/т	T/T	т/т	т/т	./.	T/T	T/T	т/т
Chr17	20189544	T/T	./.	T/T	т/т	T/A	т/т	T/A	т/т	T/A	т/т	T/A	T/A	A/A	T/T

Table B. Non-synonymous single nucleotide polymorphisms in five cassava eIF4E genes of 14 cassava accessions

*Accessions in red display tolerant responses to CBSD while accessions in white display susceptible response to CBSD. Missing data are represented by "./."

Table C. Association of SNPs upstream of *eIF4E_me* gene with CBSD disease responses.

CHR	BP	SNP	P value	Regions	Substitutions
Chr17	20190229	ss.94871927	0.004	Intergenic	G/A
Chr17	20190238	ss.94871930	0.007	Intergenic	TA/T
Chr17	20190251	ss.94871935	0.007	Intergenic	A/G
Chr17	20190252	ss.94871936	0.007	Intergenic	TC/T
Chr17	20192085	ss.94872078	0.007	Manes.17G063200	G/T, T187K

							Reads mapped to genes						
SRA	Cassava	CBSV				Total	%mapped						
accessions	accessions	Infection	Tissues	Phenotypes	Bases	reads ²	to genome	elF4E	elF(iso)4E1	elF(iso)4E2	nCBP1	nCBP2	Actin
SRR1213744	Albert	Healthy	leaf	Susceptible	2628M	42768937	0.8765	1676	98	1041	351	4136	6497
SRR1213745	Albert	Infected	leaf	Susceptible	2921M	47596602	0.886	2203	154	1558	490	6171	8700
SRR1213746	Kaleso	Healthy	leaf	Tolerant	1894M	30730846	0.8634	1224	77	804	254	2744	3927
SRR1213747	Kaleso	Infected	leaf	Tolerant	2416M	47596602	0.886	2352	191	1536	422	2825	9677

Table D. Information on Illumina RNAseq unpaired reads of two cassava accessions and Bowtie mapping statistics¹.

¹Raw sequencing reads were generated by Maruthi et al. in 2014 [1] and retrieved from <u>http://www.ncbi.nlm.nih.gov/sra</u>. ²Average read lengths are 51 bp.

Table E. Differential expression analysis of five cassava eIF4E genes in healthy and CBSV infected Albert and Kaleso accessions.

Genes	Cassava lines	Healthy (FPKM)	Infected (FPKM)	log2	test_stat	p¹	q²
elF4E	Albert	67.4627	80.5457	0.255717	0.447822	0.6613	0.998506
	Kaleso	68.9774	87.6125	0.345012	0.596169	0.54455	0.998506
elF(iso)4E1	Albert	6.18663	8.75624	0.501157	0.599087	0.5463	0.998506
	Kaleso	6.74423	11.0195	0.708338	0.878567	0.36865	0.998506
elF(iso)4E2	Albert	39.6478	53.6243	0.435647	0.752701	0.44975	0.998506
	Kaleso	42.7417	53.4709	0.32311	0.560435	0.5804	0.998506
nCBP1	Albert	9.56893	11.9476	0.320294	0.474463	0.6357	0.998506
	Kaleso	9.70205	10.443	0.106175	0.157073	0.87315	0.998506
nCBP2	Albert	135.035	181.42	0.426	0.720313	0.4565	0.998506
	Kaleso	125.056	83.9671	-0.574679	-0.990581	0.32255	0.998506

¹The uncorrected p-value of the test statistic.

²The FDR-adjusted p-value of the test statistic. FDR is set at 0.05.

Table F. Information on 454 GS FLX Titanium RNAseq reads of six cassava accessions and Bowtie mapping statistics¹.

							Reads mapped to genes					
SRA	Cassava					%mapped to						
accessions	accessions	Tissues	Phenotypes	Bases	Total reads ²	genome	elF4E	eIF(iso)4E1	eIF(iso)4E2	nCBP1	nCBP2	Actin
SRR955450	AR37-80	leaf, stem	Susceptible	88M	190325	70.84%	19	2	5	8	4	32
SRR955446	AR40-6	leaf, stem	Susceptible	101M	208365	64.21%	17	2	15	7	0	68
SRR955445	Mkombozi	leaf, stem	Susceptible	168M	346000	66.89%	22	3	19	10	3	85
SRR955449	Kiroba	leaf, stem	Tolerant	123M	291436	60.83%	7	2	6	N/A	2	54
SRR955447	Nachinyaya	leaf, stem	Tolerant	141M	261139	68.06%	5	4	13	17	1	46
SRR955444	Namikonga	leaf, stem	Tolerant	99M	205405	61.29%	14	1	5	2	3	49

¹Raw sequencing reads were generated by Ferguson et al in 2015 [2] and retrieved from <u>http://www.ncbi.nlm.nih.gov/sra</u>. ²Average read lengths are 500 bp.

	4 2,332 bp →	
	1,168,400 bp 1,168,600 bp 1,168,800 bp 1,170,000 bp 1,170,200 bp 1,170,400 bp 1,170,500 bp 1,170,000 bp 1,171,000 bp 1,170,000 bp 1,170	Ŧ
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Gene		* *

A. elF(iso)4E_me1 (Locus: Manes.03G160000)

B. nCBP_me2 (Locus: Manes.08G145200)

	414,000 bp	415,000 bp I	3,794 bp	417,000 bp
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C. nCBP_me1 (Locus: Manes.09G140300)

D. elF(iso)4E_me2 (Locus: Manes.15G044900)

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Gene			A

E. elF4E_me (Locus: Manes.17G063100)



Figure A. Distribution of nucleotide polymorphisms in five cassava eIF4E genes across 10 landraces. High quality Illumina reads of 10 cassava genomes are anchor-aligned to the cassava genome and viewed in IGV [3]. Vertical colored lines indicate nucleotide polymorphisms. Each color represents a different nucleotide substitution. Black dots indicate deletions or missing data. Biallelic variations are clearly visible in certain genes of cassava lines.



100 110 120 140 RKWTFWFDN-OSRPKQCAAWGTSIRKVYTFDTVEEFWCLYDOIFKPSKLPANADFHLFKTGVEPKWEDPE RKWTFWFDN-OSRPKQCAAWGTSIRKVYTFDTVEEFWCLYDOIFKPSKLPANADFHLFKTGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKVYTFDTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKVYTFDTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKVYTFDTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKVYTFDTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKAYTFETVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKAYTFETVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKAYTFETVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKAYTFETVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKAYTFETVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLONADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWCLYDOIFKPSKLPANADFHLFKAGVEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWULHDOIFKPSKLPANADFHLFKAGTEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWULHDOIFKPSKLPANADFHLFKAGTEPKWEDPE RKWTFWFDN-OSKPKQCAAWGTSIRKYYFFTTVEEFWULHDOIFKPSKLPANADFHLFKAGTEPKWEDPE RKWTFWFDN-OSKPKQCAAWGSIRKAYFFETVEEFWSLYDOIFKPSKLPANADFHLFKAGTEPKWEDPE RKWTFWFDN-OSKPKQCAAWGSIRKAYFFETVEEFWSLYDOIFKPSKLTVNADFHLFKAGTEPKWEDPE Cm iso ABY56090.1 Cs iso ABY56102.1 eIF(iso)4E mel eIF(iso)4E_me2 Rc EEF34040.1 Jc KDP40383.1 Vv CBI23707.3 Vv XP 002285444.1 Pt EEE89965.1 Pt EEF01754.1 Gm ACU23400.1 Pv_iso ABU54804.1 Ps iso BAM28884.1 St iso CBJ34337.1 St_iso_CCM43797.1

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Consensus

eIF4E me	CONTENEDN-PRAKSKO	ATWESSMRSITTE	ATVDEFWSIYNNI	PSKLAVGADEH	CEKYKLERKWEDRV
Jc KDP21178.1	HOWTFWFDN-PSSKSKC	AUWGSSMRSVYTF	ATVEEFWSVYNNIHE	PSKLAVGADEH	CEKYKIEPKWEDPV
Rc EEF42744.1	HOWTEWEDN-PSAKSKO	ATTWGSSMRPIYTE	ATVEEFWSIYNNIH	PSKLAVGADEH	CEKHKIEPKWEDPV
Pt_EEE97358.1	HOWTEWEDN-PSAKSKC	ASWGSSIRSIFTE	STIEEFWSVYNNIH	PSKLAVGADEH	CEKDKIEPKWEDPI
St_AER62155.1	HSWTEWEDN-PSGKSKC	AAWGSSTRPTYTE	SAAEDEWSVYNNTHE	PSKLAVGADEH	CEKNKTEPKWEDPV
C1_ACN51301_1	HSWTEWEDN-PSAKSKO	AUWGASTRPTYTE	STVEEFWSVYNNTHE	PSKLALBADLY	CEKHKTEPKWEDPV
C1 AEK98516 1	HSWTEWEGN-PSAKSKC	AUWCASTRPTYTE	STVEEFWSVYNNTHE	PSKLALPADLY	CEKHKTEPKWEDPV
C1 = ACN51302 = 1	HSWTEWEDN-PSAKSKC	APWCASTRPTYTE	STVEFFWSVYNNTHE	PSKLALRADLY	CEKHKTEPKWEDPV
Cm ABD57969 1	HSWTEWEDN-PSAKSKC		STVEREWSVINNTHE		
Cm_ABO53636_1	HSWTEWEDN-PSAKSKC	AUWCASTRPTYTE	STVEFFWSVYNNTHE		
C_{s} ABY 56085 1	HSWTEWEDN-PSAKSKC		STVEREWSVYNNTHE	PSKLALPADLY	CEKHKTEPKWEDPV
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NP 001959 1					
$ns_{NF_{001959.1}}$					
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FV_AB034823.1					
$DC_{RDF39093.1}$					
$RC_{XP} = 002323418.1$				SSALPSPIDLA DOLLDONDIU	
Pt_XP_002319689.1			STVEGEWVCYCHLAF	PSLIPSPTDLH	
NCBP_mei	HKFVFLYTRRTPGVRT(PSSLPIPIDLH	
VV_XP_002281697.1	HKFVFWY IRRIPGVRIC		STVEGEWICY CHLAF	PSALPSPIDLH	
Cs_XP_004141135.1	FRETPGVRIC		SSVEGEWISYCHLAP	PASEPSPTULH	
At_AACI/220.1	TRRIPGVRN			SSLLPSPTDLH	
Consensus	• •		<mark></mark>	_	
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	150	160 17	0 180	190	200 210
	150 ±. ••.	160 17	0 1 80 •••••	190	200 210
Cm iso ABY56090.1	150 ★.★. CANGGKWTVTSSR	160 17 	0 180 	190 • * • * * •• * *• • VVASVR-OROD	200 210 ★ ↓ KLALWIKTATNDAA
Cm_iso_ABY56090.1 Cs_iso_ABY56102.1		160 17 	0 180 LICEOFE-ESDEICC	190 (•★•★ ★••★★• VVASVR-OROD	200 210
Cm_iso_ABY56090.1 Cs_iso_ABY56102.1 eIF(iso)4E_me1	150 ★ .★ . CANGGKWTVTSSR CANGGKWTVTSSR CANGGKWSVTCGR	160 17 	0 180 LIGEOFE-ESDEICO LIGEOFE-ESDEICO LIGEOFD-EADEICO	190 VVASVR-QRQD VVASVR-QRQD VVASVR-QRQD	200 210
Cm_iso_ABY56090.1 Cs_iso_ABY56102.1 eIF(iso)4E_me1 eIF(iso)4E_me2	150 ★.★. . CANGGKWTVTSSR CANGGKWTVTSSR CANGGKWSVTCGR CACGGKWSVTCGR	160 17 	0 180 $LIGEQFE-ESDEICO$ $LIGEQFE-ESDEICO$ $LIGEQFD-EADEICO$ $LIGEQFD-EADEICO$	190 ••••••••••••••••••••••••••••••••••••	200 210
Cm_iso_ABY56090.1 Cs_iso_ABY56102.1 eIF(iso)4E_me1 eIF(iso)4E_me2 Rc EEF34040.1	150 CANGGKWTVTSSR CANGGKWTVTSSR CANGGKWSVTCGR CAGGKWSVTCGR CAGGKWSVTCGR	160 17 	0 180 $LIGEQFE-ESDEICO$ $LIGEQFE-ESDEICO$ $LIGEQFD-EADEICO$ $LIGEQFD-EADEICO$ $LIGEQFD-EADEICO$	190 • • • • • • • • • • • • • • • • • • •	200 210 KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA
Cm_iso_ABY56090.1 Cs_iso_ABY56102.1 eIF(iso)4E_me1 eIF(iso)4E_me2 Rc_EEF34040.1 Jc KDP40383.1	150 	160 17 	0 180 $LIGEQFE-ESDEICO$ $LIGEQFE-ESDEICO$ $LIGEQFD-EADEICO$ $LIGEQFD-EADEICO$ $LIGEQFD-SDEICO$ $LIGEQFD-SDEICO$	190 • • • • • • • • • • • • • • • • • • •	200 210 KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA
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Cm_iso_ABY56090.1 Cs_iso_ABY56102.1 eIF(iso)4E_me1 eIF(iso)4E_me2 Rc_EEF34040.1 Jc_KDP40383.1 Vv_CBI23707.3 Vv_XP_002285444.1 Pt_EEE89965.1 Pt_EEF01754.1 Gm_ACU23400.1 Pv_iso_ABU54804.1 Ps_iso_CBJ34337.1 St_iso_CBJ34337.1 St_iso_CBJ34337.1 St_iso_CCM43797.1 eIF4E_me Jc_KDF21178.1 Rc_EEF42744.1 Pt_EEE97358.1 St_AER62155.1 Cl_ACN51301.1 Cl_ACN51302.1 Cm_AB05769.1 Cm_AB053636.1	150 	160 17 ANIDNMWLETIMA ANIDNMWLETIMA AILD TIWLETIMA AILD TIWLETIMA AILD TIWLETIMA BILB TIWLETIMA GNLD TIWLETIMA GNLD TIWLETIMA GNLD TIWLETIMA ANID TIWLETIMA ANID TIWLETIMA ANID TIWLETIMA ANID TIWLETIMA ANID TIWLETIMA GNLD TIWLETIMA GNLD TIWLETIMA GNLD TIWLETIMA GNLD TIWLETIMA GNLD TIWLETIMA GNLD TIWLETIMA GNLD TIWLETIMA GNSD TSWLYTLLA GKSD TSWLYTLLA GKSD NGWLYTLLA GKSD NGWLYTLLA GKSD NGWLYTLLA	0 180 	190	200 210 KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTAANEAA KLSLWTKTAANEAA KLSLWTKTAANEAA KLSLWTKTAANEAA KLSLWTKTATNEAA KLSLWTKTATNEAA KLSLWTKTATNEAA KLSLWTKNASNEAA KLALWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA KLSLWTKNASNEAA
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Cm_iso_ABY56090.1 Cs_iso_ABY56102.1 eIF(iso)4E_me1 eIF(iso)4E_me2 Rc_EEF34040.1 Jc_KDP40383.1 Vv_CB123707.3 Vv_XP_002285444.1 Pt_EEE89965.1 Pt_EEF01754.1 Gm_ACU23400.1 Pv_iso_ABU54804.1 Ps_iso_BAM28884.1 St_iso_CBJ34337.1 St_iso_CCM43797.1 eIF4E_me Jc_KDP21178.1 Rc_EEF42744.1 Pt_EEE97358.1 St_AER62155.1 Cl_ACN51301.1 Cl_AEK98516.1 Cl_ACN51302.1 Cm_ABD57969.1 Cm_ABD57969.1 Cm_ABQ53636.1 Cs_APY56085.1 Cs_XP_004161134.1 Ps_AAT44121.1_sbm1	150 I * * I I I CANGGKWTVTSS RK CANGGKWSVTCG RK CANGGKWSVTCG RK CANGGKWSVTCG RK CANGGKWSVACG RK CANGGKWSVACG RK CANGGKWSVACG RK CANGGKWSVTSS RK CANGGKWTVASS RK CANGGKWTVASS RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSNIG-RK CANGGKWTVTSIG-RK CANGGKWTVTSIG-RK CANGGKWTVTSRK CANGGKWTVTFGF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF CANGGKWTVNFSF	160 17 ANIDNMWLETIMA ANIDNMWLETIMA AIID TIWLETIMA AIID TIWLETIMA AIID TIWLETIMA AIID TIWLETIMA GNID TIWLETIMA GNID TIWWLETIMA GNID TIWWLETIMA ANID TIWWLETIMA ANID TIWWLETIMA ANID TIWWLETIMA ANID TIWWLETIMA ANID TIWWLETIMA ANID TIWWLETIMA ANID TIWWLETIMA GNID TI	0 180 LIGEQFE -ESDEICG LIGEQFD -EADEICG LIGEQFD -ESEDICG MIGEQFD -GDEICG MIGEQFD -CGDEICG MIGEQFD -CGDEICG	190	200 210 KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTAANBAA KLALWTKTAANBAA KLSLWTKTAANBAA KLSLWTKTAANBAA KLSLWTKTAANBAA KLSLWTKTAANBAA KLSLWTKTAANBAA KLSLWTKNASNBAA KIALWTKNASNBAA KIALWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA KISLWTKNASNBAA
Cm_iso_ABY56090.1 Cs_iso_ABY56102.1 eIF(iso)4E_me1 eIF(iso)4E_me2 Rc_EEF34040.1 Jc_KDP40383.1 Vv_CB123707.3 Vv_XP_002285444.1 Pt_EEE89965.1 Pt_EEF01754.1 Gm_ACU23400.1 Pv_iso_ABU54804.1 Ps_iso_BAM28884.1 St_iso_CBJ34337.1 St_iso_CCM43797.1 eIF4E_me Jc_KDF21178.1 Rc_EEF42744.1 Pt_EEE97358.1 St_AER62155.1 C1_ACN51301.1 C1_AEK98516.1 C1_ACN51302.1 Cm_ABD57969.1 Cm_ABD57969.1 Cm_ABQ53636.1 Cs_XP_004161134.1 Ps_AAT44121.1_sbm1 Ps_ABG35118.1	150 *.*. CANGGKWTVTSSR CANGGKWTVTSSR CANGGKWSVTCGR CANGGKWSVTCGR CANGGKWSVACGR CANGGKWSVACGR CANGGKWTVASSR CANGGKWTVASSR CANGGKWTVASSR CANGGKWTVASSR CANGGKWTVTSSR CANGGKWTVTSSR CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSR CANGGKWTVTFGF CANGGKWTVFSF CANGGKWTVNFSF	160 17 ANIDNMWLETIMA ANIDNMWLETIMA ANID TWLETIMA AILD TWLETIMA ALLD TWWLETIMA ALLD TWWLETIMA CHID TWWLETIMA CHID TWWLETIMA CHID TWWLETIMA CHID TWWLETIMA CHID TWWLETIMA ANID TWWLETIMA ANID TWWLETIMA ANID TWWLETIMA ANID TWWLETIMA CHID TWWLE	0 180 	190	200 210 KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTATNBAA KLALWTKTAANBAA KLALWTKTAANBAA KLSLWTKTAANBAA KLSLWTKTAANBAA KLSLWTKTAANBAA KLSLWTKTAANBAA KLSLWTKTAANBAA KLSLWTKNASNBAA KIALWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA KISIWTKNASNBAA
Cm_iso_ABY56090.1 Cs_iso_ABY56102.1 eIF(iso)4E_me1 eIF(iso)4E_me2 Rc_EEF34040.1 Jc_KDP40383.1 Vv_CB123707.3 Vv_XP_002285444.1 Pt_EEE89965.1 Pt_EEF01754.1 Gm_ACU23400.1 Pv_iso_ABU54804.1 Ps_iso_BAM28884.1 St_iso_CCM43797.1 eIF4E_me Jc_KDF21178.1 Rc_EEF42744.1 Pt_EEE97358.1 St_AER62155.1 C1_ACN51301.1 C1_AEK98516.1 C1_ACN51302.1 Cm_ABD57969.1 Cm_ABQ53636.1 Cs_ABY56085.1 Cs_XP_004161134.1 Ps_AAT44121.1_sbm1 Ps_ABG35118.1 Gm_ACU23738.1	150 *.*. CANGGKWTVTSSR CANGGKWTVTSSR CANGGKWSVTCGR CANGGKWSVTCGR CANGGKWSVTCGR CANGGKWTVASSR CANGGKWTVASSR CANGGKWTVASSR CANGGKWTVASSR CANGGKWTVSSR CANGGKWTVTSSR CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSNIG-R CANGGKWTVTSR CANGGKWTVTFG	160 17 ANLDNMWLETLMA ANLDNMWLETLMA ALLD TIWLETLMA ALLD TIWLETLMA ALLD TIWLETLMA ALLD TIWLETLMA CALLD	0 180 	190	200 210 KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTATNEAA KLALWTKTAANEAA KLALWTKTAANEAA KLSLWTKTAANEAA KLSLWTKTAANEAA KLSLWTKTATNEAA KLSLWTKTATNEAA KLSLWTKTAANEAA KLSLWTKNASNEAA KLALWTKNASNEAA KLSLWTKNASNEAA
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Pv ABU54812.1		-RGKSDTSWI	THAMICPOFF	-YCDDICCAVVNVR	-NRODKISIWIKNASNDAA
Pv ABU54817.1	CANGGKWTMTFO	-RGKSDTSWL	TLLAMIGEOFD	-YGDEICGAVVNVR	-NRODKISIWTKNASNEAA
Vy XP 002267488.1	CANGGKWTVIIFA		TLLAMIGEOFD	-HGDEICGAVINVR	AROEKIALWTKNASNEAA
Hs NP 001959.1	NKRGGRWLITLNKOO	RRSDLDRFWL	TLLCLICESFD	DYSDDVCGAVVNVR-	-AKCOKIAIWTTECENREA
Gm ACU15859.1	NCNGGKWIIRFK	- <mark>K</mark> VVS <mark>GR</mark> FWEI		-YGDNICGAVLSIR	-FNEDILSVWNRNASDHOA
Pv ABU54825.1	NCNGGKWIIRFK	- <mark>K</mark> VVS <mark>GRF</mark> WEI		-YGDNICGAVLSIR	-FNEDILSVWNRNASDHOA
nCBP me2	NSNGGKWII <mark>RFK</mark>	- <mark>K</mark> VVS <mark>GRFWE</mark> I	MVLALVGDOLD	-YGDNICGAVLSIR	-FNEDILSVWNRNSSDHOA
Jc KDP39693.1	NSNGGKWII <mark>RFK</mark>	- <mark>K</mark> VVS <mark>GRY</mark> W <mark>E</mark> I	UVLALVGDQLD	-YGDNICGAVLSIR	-FNEDILSVWNRNASDHOA
Rc XP 002525418.1	NSNGGKWII <mark>RFK</mark>	- <mark>K</mark> AVSARFWEI	UVLALVGDOLE	-YGDNICGAVLSIR	-FNEDIVSVWNRNASDHOA
Pt XP 002319689.1	NCNGGKWIIRFK	- <mark>K</mark> VVS <mark>GRFWE</mark> I	DIVLALVGDQID	-YSDDICGAVLSIR	-YNEDILSVWNRNASDROG
nCBP me1	NSNGGKWIIRFK	- <mark>K</mark> AVS <mark>GRF</mark> W <mark>E</mark> I	DIVLALVGDQLG	-YSDNVCGAVLSIR	-FNEDILSIWNRNASDHOA
Vv XP 002281697.1	NCNGGKWIIRFK	- <mark>K</mark> VVS <mark>GRF</mark> W <mark>E</mark> I	DIVLALVGDQLD	-YGDNICGAVLSIR	-FNEDILSVWNRNASDHOA
Cs XP 004141135.1	NCNGGKWIIRFK	- <mark>K</mark> VVS <mark>GREWE</mark> I	D <mark>IVL<mark>A</mark>LV<mark>GDQI</mark>D</mark>	-FGDNICGAVLSIR	-FNEDILSVWNRNASDHOA
At AAC17220.1	NCNGGKWIIRFS	- <mark>K</mark> VVS <mark>ARFWE</mark> I	LLL <mark>A</mark> LV <mark>GDQL</mark> D	-DADNICGAVLSVR-	-FNEDIISVWNRNASDHQA
Consensus	<mark>GG</mark> - <mark>W</mark>	- - <mark>w</mark>	<mark>G</mark>	<mark>CG</mark> -VR	<mark></mark> w
		1	_		1
	220	230	240	250	
		•••••	· · · · · ** * •		
Cm_iso_ABY56090.1	Q-MSI <mark>GRKWKE</mark> IIDV	NDKISESEH	<mark>EDLRR</mark> -EKSA	KARYSV-	
Cs_iso_ABY56102.1	Q-MSI <mark>GRKWKE</mark> IIDV	T <mark>DKI</mark> SFSFH	<mark>EDLRR</mark> -EKSA	KARYSV-	
eIF(iso)4E_me1	Q-MSI <mark>GKKWKE</mark> IIDV	TEKITYSFH	<mark>DDSRR</mark> -ERSV	K <mark>G</mark> RYNV-	
eIF(iso)4E_me2	Q-M <mark>GIGKKWKE</mark> IIDV	T <mark>DKI</mark> TYSFH	<mark>DDSRR</mark> -ERS <mark>A</mark>	K <mark>G</mark> RYNV-	
Rc_EEF34040.1	Q-M <mark>GVGKKWKE</mark> IIDV	TDKITYSFH	DDSRR-ERSV	KSRYNV-	
Jc_KDP40383.1	Q-MSI <mark>GKKWKE</mark> IIDV	TDKITYSFH	DDSRR-ERSV	KSRYSV-	
Vv_CBI23707.3	Q-MSI <mark>GRKWKE</mark> VIDV	TDKITYSFH	DDSRR-ERSV	KVRYNV-	
Vv_XP_002285444.1	Q-MSI <mark>GRKWKE</mark> VIDV	I <mark>DK</mark> ITYSFH	- – <mark>DDSRR</mark> –ERSV	KVRYNV-	
Pt_EEE89965.1	Q-MSI <mark>GRKWKE</mark> IIDV	N <mark>D</mark> KITYSFH	DDSKR-ERNA	KSRYNV-	
Pt_EEF01754.1	QMMSI <mark>GRKWKE</mark> VIDV	INKITYSFH	DDSKR-ERNA	KSRYNV-	
Gm_ACU23400.1	Q-MSI <mark>GRKWKE</mark> IIDV	NDKITYNFH	DDSRT-KGAS	K <mark>GRY</mark> TV-	
Pv_iso_ABU54804.1	Q-MSI <mark>GRKWKE</mark> IIDVI	NDKITYNEH	DDSRT-KGAT		
Ps_iso_BAM28884.1	Q-MSI <mark>GRKWKE</mark> IIDV	SDKMTYNFH	EDAKT-RGA-	KARYIV-	
St_iso_CBJ34337.1	Q-MGIGRKWKEIIDT	- <mark>EKI</mark> SYSEH	DDSKR-ERSA	KSRYIV-	
St_iso_CCM43797.1	Q-M <mark>GIGRKWKGIID</mark> T	-EKISYSEH	DDSKR-ERSA	KSRYIV-	
eIF4E_me	Q-LSI <mark>CKQWKE</mark> BLDY	NDTIGEIEH	EDAKKIDRGA	KNRYTI –	
Jc_KDP21178.1	Q-LSI <mark>GKQWKE</mark> ILDY		EDAKKHDRAA	KNRYTT-	
Rc_EEF42/44.1			DDAKKHERSA	KNRYTI –	
Pt_EEE9/358.1			DDAKK-DRNA	KNRYSV-	
St_AER62155.1				KNRYSV-	
CI_ACN51301.1				KNKYSV-	
CI_AEK98516.1				KNKYSV-	
CI_ACN51302.1				KNKYSV-	
$Cm_ABD57969.1$					
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De ABC35110 1				KNKYW	
Gm ACII23738 1				KNKYWV-	
Gm XP 003546060 1				KNKYWZ-	
Gm ACM45958 1			EDAKKLDRCA	KNKYWV-	
Pv ABU54812 1			EDAKKHEB		
Pv ABU54817 1		NEPTGETEH	EDAKKHER		
Vv XP 002267488 1	0-LSIGKOWKEFLDY	SESTGETEH	EDAKKLDRAA	KNRYTT-	
Hs NP 001959 1	V-THIGRVYKERI CT	PPKIVIGYOSE	ADTATKSGSTT	KNRFVV-	
Gm ACU15859 1	V-MATEDSIKEHLKT	PHSYVMEYKP-	HDASLEDNSSY		
Pv ABU54825.1	V-MATRDSIKRHIKI	PHSYVMEYKP-	HDASLRDNSSY	RNHWLIRG	
nCBP me2	V-MALEDSIKEHLKI	PHSYVMEYKP-	HDASLRDNSSY	RNUWLRG	
Jc KDP39693.1	V-MALEDSIKEHLKI	PHSYVMEYKP-	HDASLRDNSSY	RNUWLRG	
Rc XP 002525418.1	V-MATEDSIKEHTKT	PHGYVMEYKP-	HDASLRDNSSY	RNHWLIRG	
Pt XP 002319689.1	V-MALRDSIKRHIKI	PHSYVMDYKP-	-HDASLRDNSSY		
nCBP me1	V-MALRDSLKRHLKL	PHNYVMEYKP-	-HDASLRDNSSY		
Vv XP 002281697.1	V-MALRDAIKRHIKI	PHGYVMEYKA-	HDASLRDNSSY		
Cs XP 004141135.1	V-MGLRDSIKRHLKL	PHGYVMEYKP-	HDASLRDNSSY		
At AAC17220.1	V-MGLRDSIKRHLKT	PHAYVMEYKP-	-HDASLRDNSSY		
Consensus	<mark>K</mark>				
	-				

Figure B. Alignment of cassava eIF4E proteins with representative plant and human eIF4E proteins.

Proteins were aligned with ClustalX 2.0 [4] and manually inspected and adjusted. Shading of similar amino acids and display were performed with BioEdit 7.09 [5]. Amino acids conserved in all eIF4E proteins are noted in the consensus line. Eight conserved tryptophan (W) amino acids are indicated with arrows and two involved in cap-binding are indicated in red arrows. Positions of eIF4E proteins where amino acid substitutions resulted in recessive resistance to potyviruses in nature or experimentally [6,7] are indicated by red six-point stars. Protein sequences of additional eIF4E family proteins were retrieved from NCBI. Protein sources are identified with two letters representing the initials of genus and species, followed by NCBI accession number. At, *Arabidopsis thaliana*; Ca: *Capsicum annuum*; Cl, *Citrullus lanatus*; Cm, *Cucumis melo*; Cs, *Cucumis sativus*; Gm, *Glycine max*; Hs, *Homo sapiens*; Jc, *Jatropha curcas*; Ps, *Pisum sativum*; Pt, *Populus trichocarpal*; Pv, *Phaseolus vulgaris*; Rc, *Ricinus communis*; St, *Solanum tuberosum*; and Vv, *Vitis vinifera*.





Figure C. Single nucleotide polymorphisms (SNPs) in the five cassava eIF4E genes and the upstream and downstream 5 kb regions. Data were obtained from combined SNPs and indels from 61 cassava accessions curated at Phytozome (phytozome.jgi.doe.gov). Images of SNPs were rendered in the Genome Browser implemented at the same site. Blue diamonds: synonymous nucleotide substitutions in coding regions or substitutions in non-coding regions; yellow diamonds: non-synonymous substitutions in coding regions; red diamonds: missense substitutions; vertical green lines: indels. Only a 2kb region upstream of nCBP2 is available from the genomic sequence.



Figure D. Expression levels of five eIF4E genes in CBSD-susceptible and -tolerant lines. RNAseq long reads (0.5 kb) specific to each gene from each of the lines generated by the Life Sciences 454 sequencing were counted and normalized to total mapped reads from each sample to generate the FPKM counts. AR37-80, AR40-6, and Mkombozi were susceptible to CBSD while Kiroba, Nachinyaya, and Namikonga were tolerant to CBSD. Raw data were retrieved from NCBI SRA (accessions SRR955444-SRR955447, SRR955449, SRR955450, SRR955453, and SRR955456) [2].

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