Supplementary Materials: A Novel Cyclic Pentadepsipeptide, *N*-methylsansalvamide, Suppresses Angiogenic Responses and Exhibits Antitumor Efficacy against Bladder Cancer

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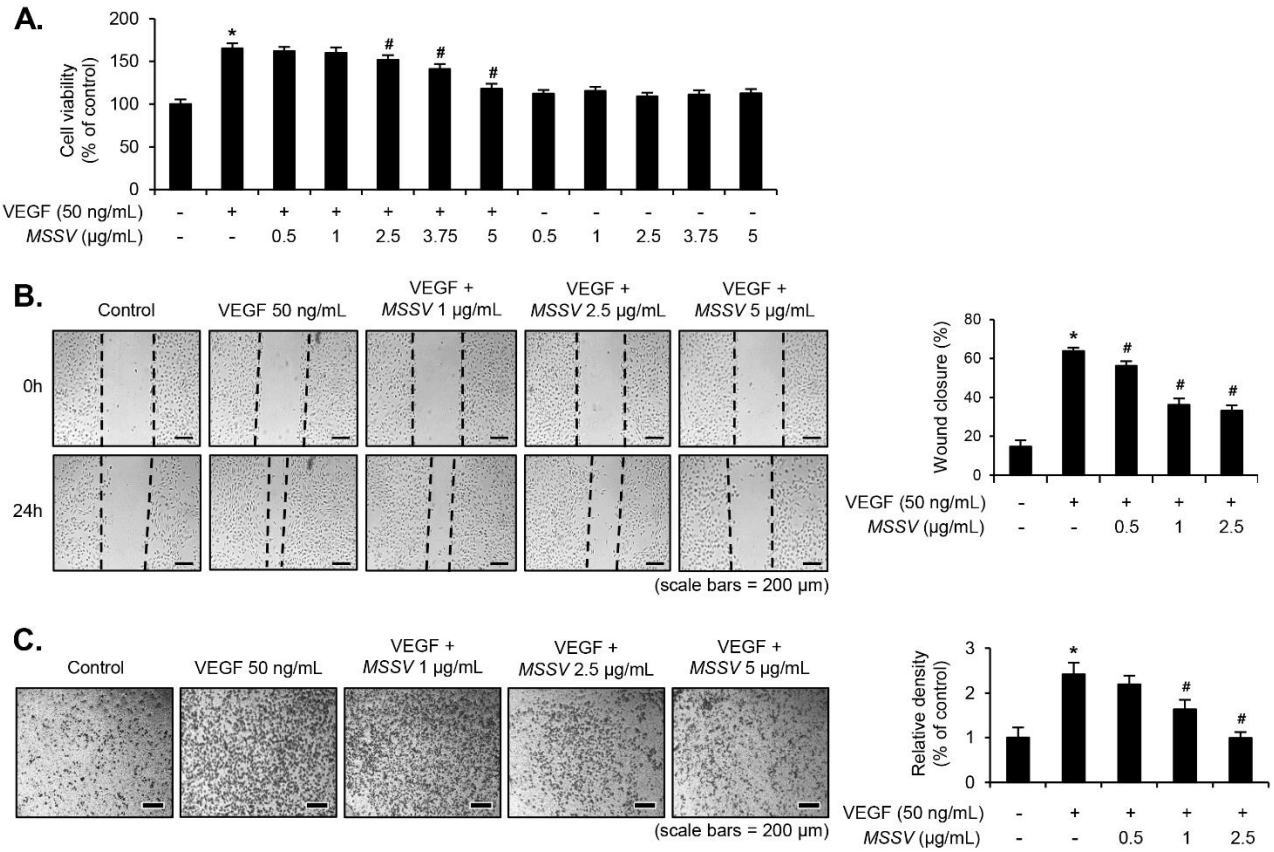


Figure S1. MSSV inhibits VEGF-induced proliferation, migration, and invasion of HUVECs. HUVECs were treated with VEGF (50 ng/mL) for 1 h, followed by the indicated concentrations of MSSV for 24 h. (**A**) MTT assay was performed after treatment with MSSV in the presence or absence of VEGF (50 ng/mL). (**B,C**) The migration and invasion abilities of HUVECs were analyzed by wound healing migration and Boyden chamber invasion assays. All data are reported as the means ± SE from three independent experiments. $^{*}p < 0.05$ compared with control and $^{\#}p < 0.05$ compared with VEGF treatment.

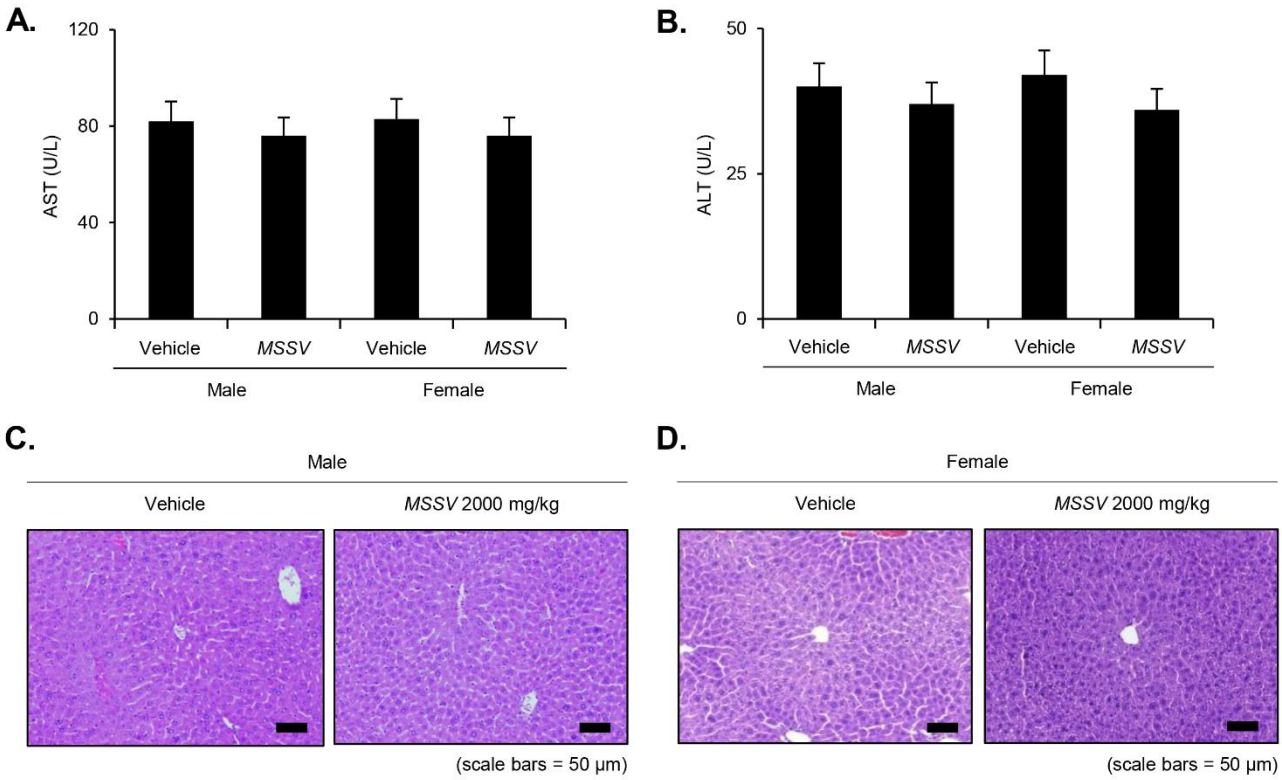


Figure S2. Acute toxicity effects of MSSV on biochemical parameters in male and female mice. After the oral administration of MSSV (2000 mg/kg), the levels of AST (**A**) and ALT (**B**) were measured. (**C,D**) H&E staining of the liver tissues obtained from MSSV-treated mouse. All data are represented as the means ± SE from three independent experiments. *$p < 0.05$ compared with the control.

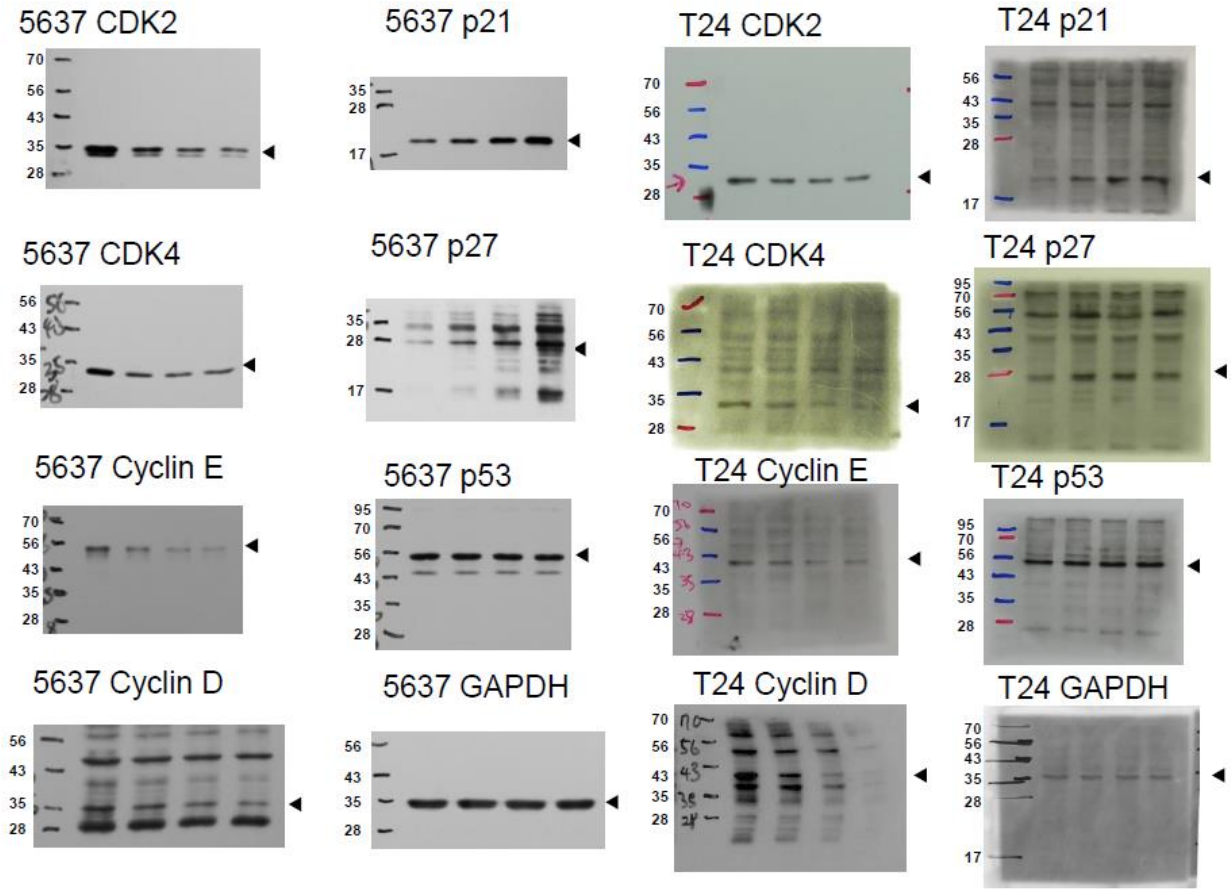


Figure S3. Uncropped Western Blot Images for Figure 2D.

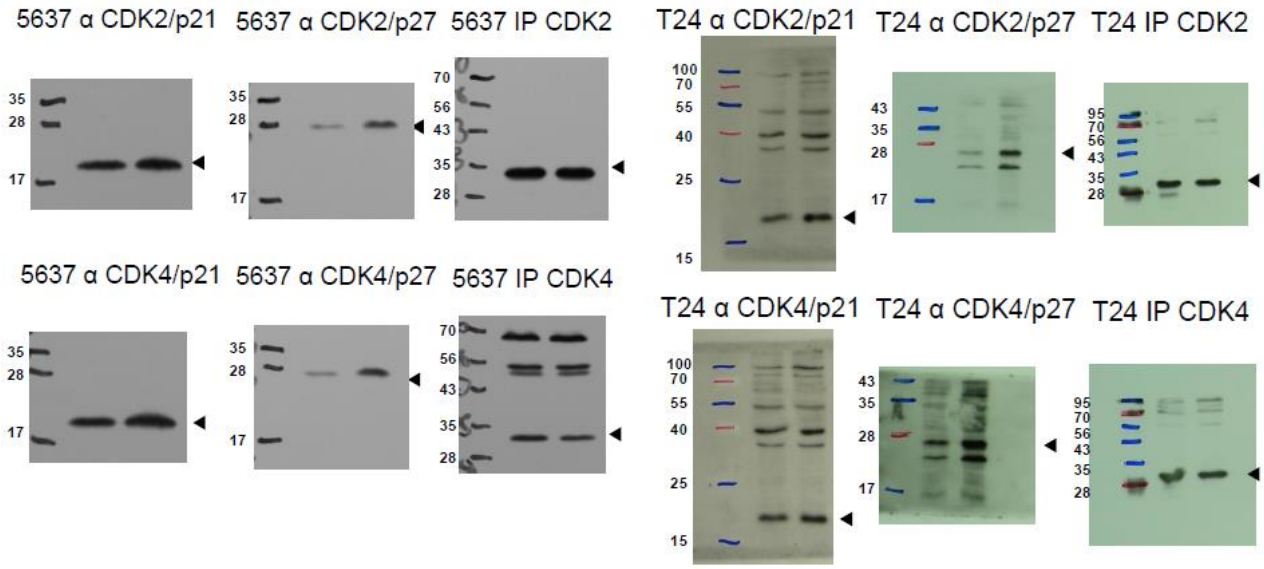


Figure S4. Uncropped Western Blot Images for Figure 2E.

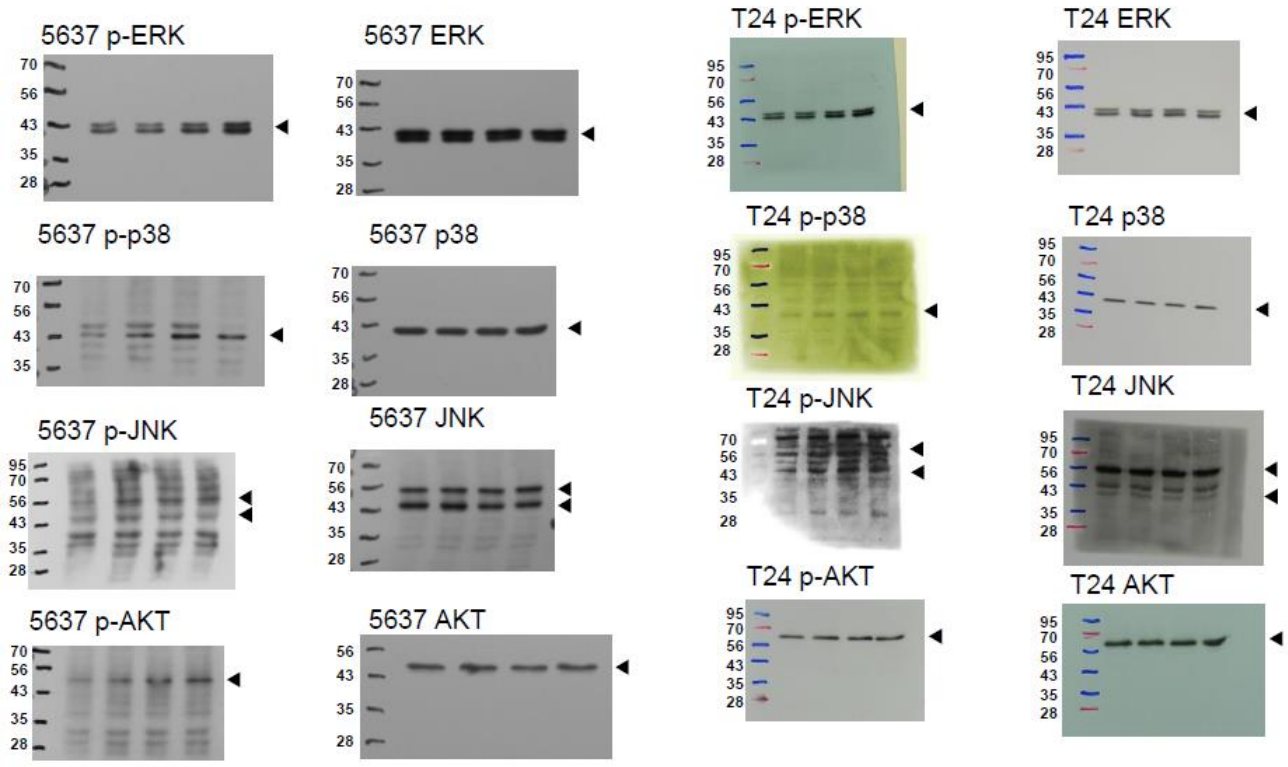


Figure S5. Uncropped Western Blot Images for Figure 3A.

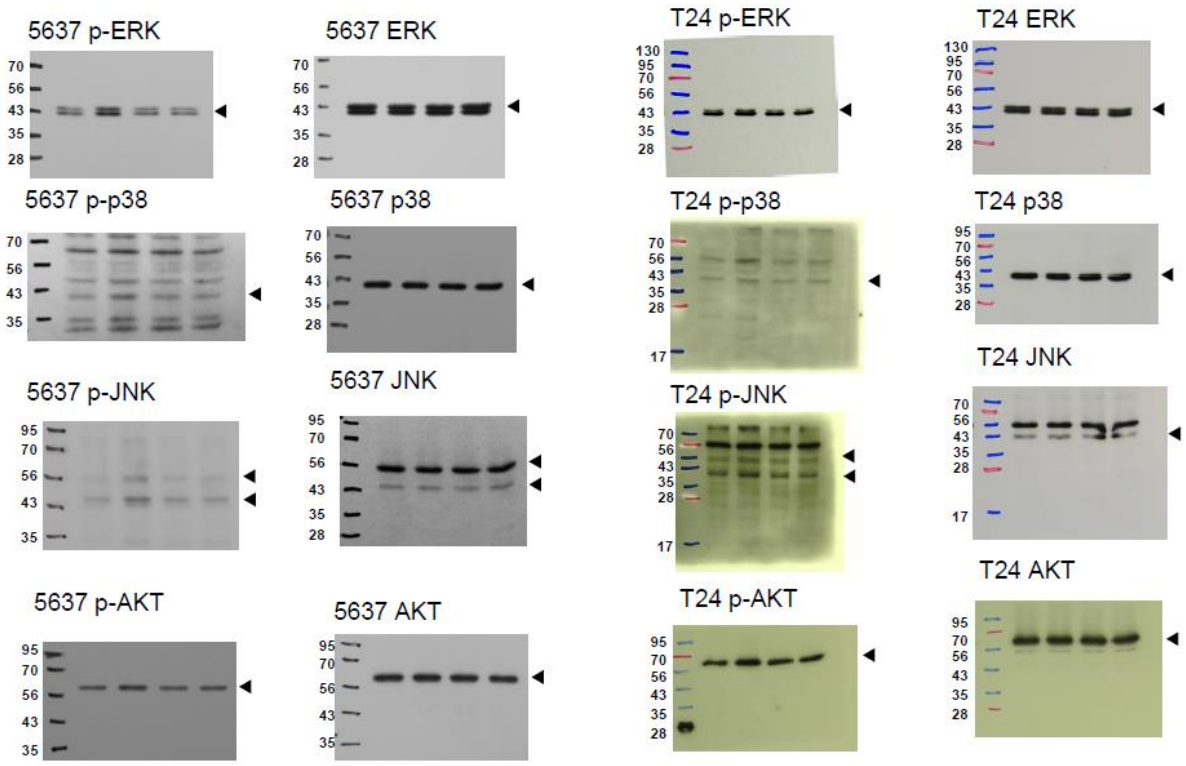


Figure S6. Uncropped Western Blot Images for Figure 3B.

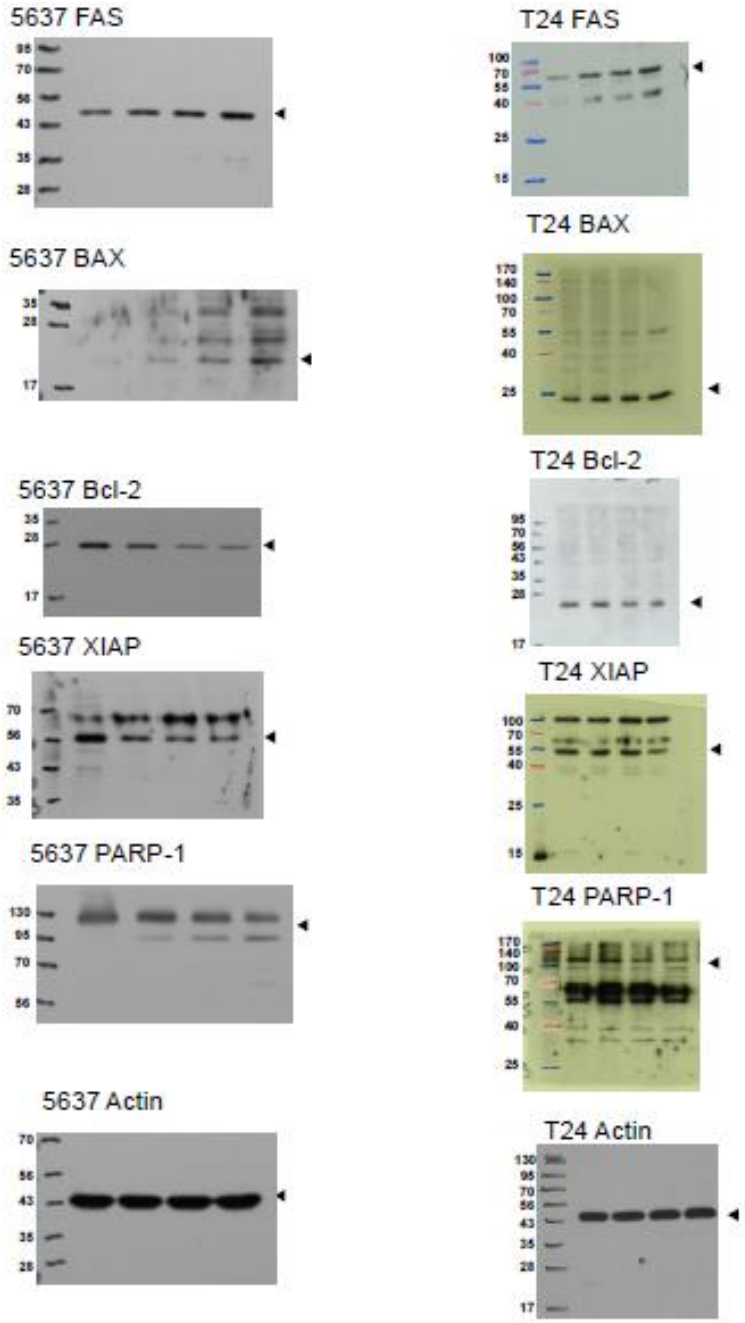


Figure S7. Uncropped Western Blot Images for Figure 5B.

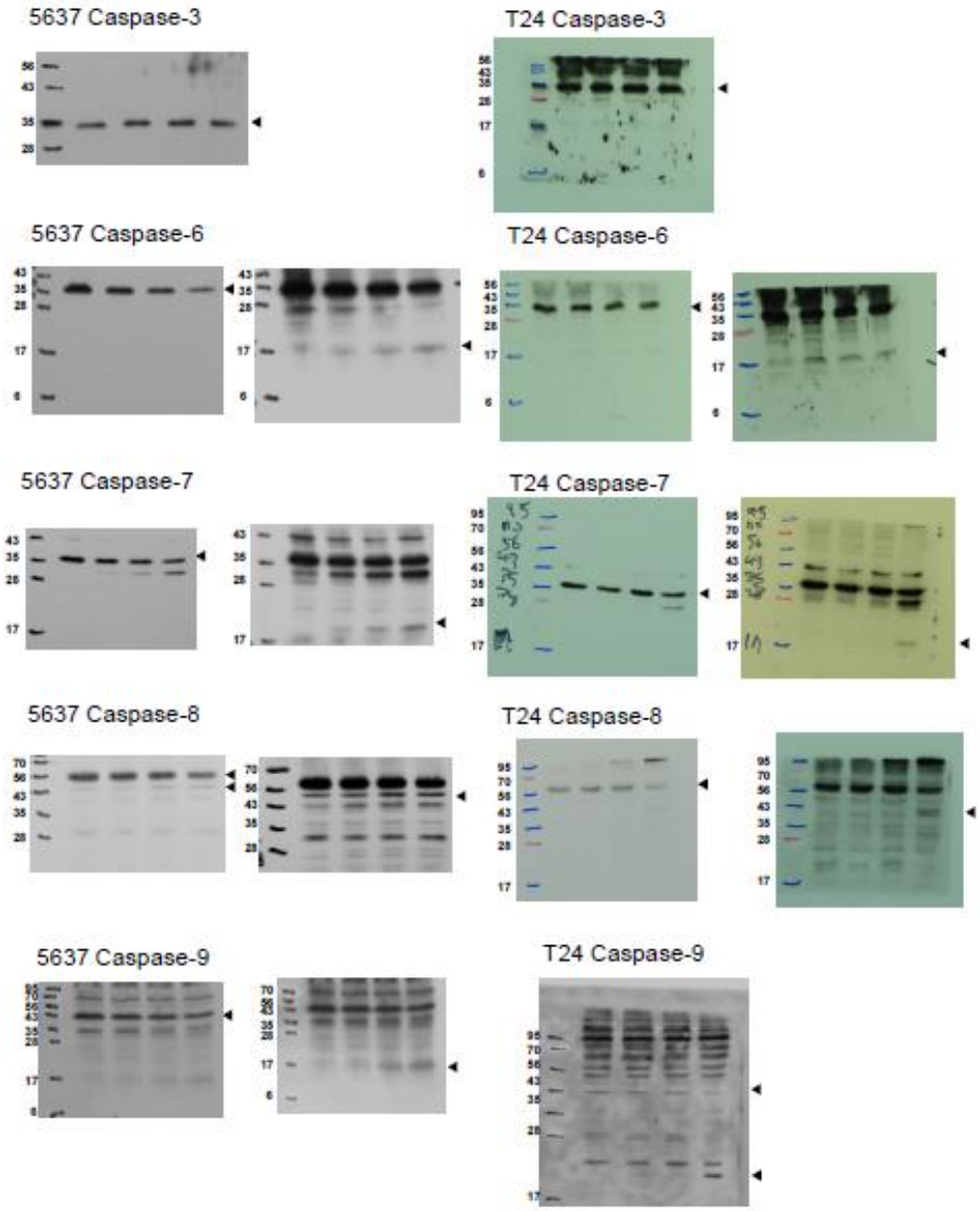


Figure S8. Uncropped Western Blot Images for Figure 5C.

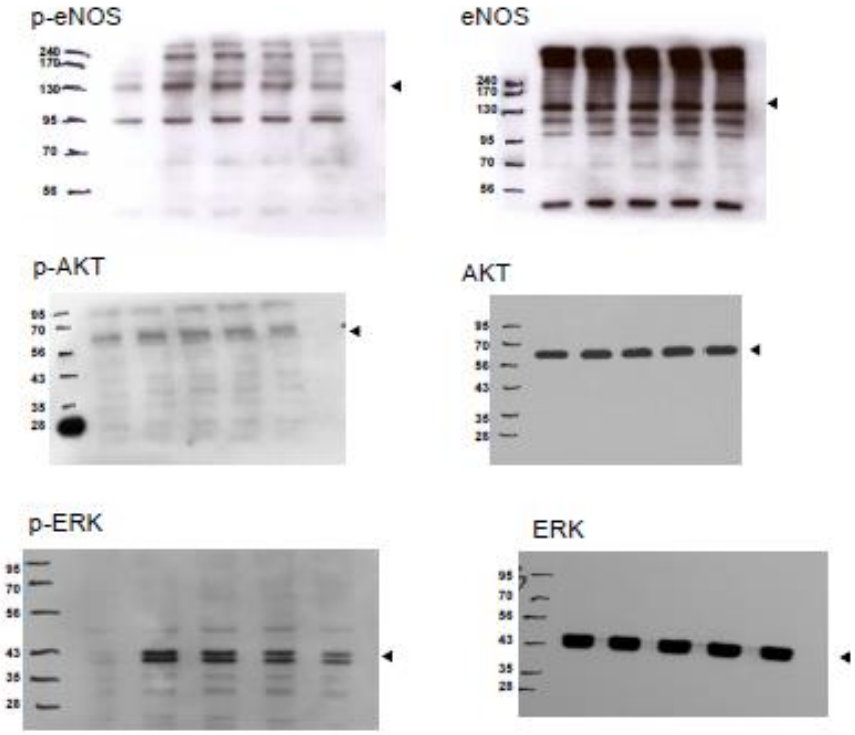


Figure S9. Uncropped Western Blot Images for Figure 7B.

Table S1. ^{1}H (400 MHz) and ^{13}C (100 MHz) NMR data for N-methylsansalvamide in pyridine-d_5.

Position		δ_C, mult.	δ_H, (*J* in Hz)
Valine	NH		7.76 (d. 8.9)
	α	55.5, CH	4.97 (dd. 8.9, 6.3)
	β	31.1, CH	2.31 (m)
	γ	20.6, CH_3	1.01 (d, 6.8)
	γ'	17.7, CH_3	1.11 (d, 6.6)
	CO	173.2, C	
N-methyl Leu	N-CH_3	39.9	3.18 (s)
	α	68.4, CH	3.71 (dd. 9.0, 6.6)
	β	39.7, CH_2	2.31[a], 2.06[a]
	γ	26.1, CH	1.60 (m)
	δ	24.0, CH_3	0.83 (d, 6.3)
	δ'	23.6, CH_3	0.92 (d, 6.6)
	CO	172.2	
Phenylalanine	NH		8.68 (d, 9.0)
	α	56.5, CH	5.10 (td. 9.1, 6.4)
	β	39.7, CH_2	3.54 (dd, 13.8, 6.4), 3.39 (dd, 13.8, 9.2)
	C1	138.9, C	
	C2	129.4, CH	7.30[a]
	C3	129.3, CH	7.32[a]
	C4	127.4, CH	7.28[a]
	CO	172.9, C	
Leucine	NH		9.45 (d, 8.6)
	α	52.6, CH	4.87 (ddd, 10.9, 8.5, 4.6)
	β	38.3, CH_2	2.22[a], 1.93[a]
	γ	25.0, CH	1.50[a]
	δ	22.4, CH_3	0.94 (d, 6.3)
	δ'	23.3, CH_3	0.88 (d, 6.4)
	CO	172.0[a]	
2-hydroxy-4-methyl pentanoic acid	C2	76.3, CH	5.35 (dd, 9.6, 3.7)
	C3	41.5, CH_2	1.91[a]
	C4	25.6, CH	2.05[a]
	C5	21.7[a], CH_3	0.94[a]
	C6	22.1[a], CH_3	0.92[a]
	CO	171.4[a]	

[a]Overlapped signals.

Table S2. Assessment of acute toxicity on oral consumption of *MSSV* in male and female mice.

Group	Dose (mg/kg)	Volume (mL/kg)	Number of Animals		Symptom	Autopsy
			Male	Female		
Control	0	4	5	5	Normal	Normal
MSSV	2000	4	5	5	Normal	Normal

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