

Supplementary Materials and Methods

Supplementary Table 1. Composition of the Murashige & Skoog medium used for the growth of plant cell cultures.

Ingredients	Amount
Carbon source	g/L
Sucrose	30
Micro Elements	mg/l
CoCl ₂ .6H ₂ O	0.025
CuSO ₄ .5H ₂ O	0.025
FeNaEDTA	36.70
H ₃ BO ₃	6.20
KI	0.83
MnSO ₄ .H ₂ O	16.90
Na ₂ MoO ₄ .2H ₂ O	0.25
ZnSO ₄ .7H ₂ O	8.60
Macro Elements	mg/l
CaCl ₂	332.02
KH ₂ PO ₄	170.00
KNO ₃	1900.00
MgSO ₄	180.54
NH ₄ NO ₃	1650.00
Vitamins	mg/l
Glycine	2.00*
myo-Inositol	100.00
Nicotinic acid	0.50*
Pyridoxine HCl	0.50*
Thiamine HCl	0.10* or 1**
Phytohormones	mg/L
Kinetin	0.1*
Naphtalene acetic acid (NAA)	1*
2,4-Dichlorophenoxyacetic acid	0.2**

*only added for the culture of rowan PCCs

**only added for the culture of scurvy grass PCCs

Supplementary Table 2. Histopathology grading parameters of tissue specimens.

Brain								
Parameters	Score							
	0	1	2	3	4	5	6	7
Necrosis (area)	None	≤10%	~20%	~30%	~45%	~60%	~75%	90-100%
Haemorrhage	Absent	Mild	Moderate	Severe				
Heart								
Parameters	Score							
	0	1	2	3	4	5	6	7
Necrosis (area)	None	≤10%	~20%	~30%	~45%	~60%	~75%	90-100%
Lesions	None	Focal lesions at apex and middle of ventricles	Focal lesions over both ventricles	Confluent lesions at apex, middle of both of ventricles				
Haemorrhage	Minimal	None	Mild & moderate	Diffuse				
Lungs								
Parameters	Score							
	0	1	2	3				

Vascular features	Obstruction	None	Mild RBC obstruction	Mild RBC and vascular obstruction	Moderate RBC and vascular obstruction
	Haemorrhage		Blood leaking into interstitium	Areas of mild and moderate haemorrhage	Diffuse haemorrhage
Extravascular and alveolar features	Inflammatory exudate	Minimal	Mild inflammatory exudate; areas of patchy oedema with some disordered structure	Moderate	Moderate-severe
	Alveolar thickening			~25-50%	>50%, loss of structure with amorphous material
	Alveolar obstruction	Minimal	Mild	Moderate	Moderate-severe
Bronchiole features		Mild infiltration of inflammatory cells	Moderate infiltration of inflammatory cells, detachment of lining in some bronchioles	Complete loss of structure, detachment of lining, cellular debris, inflammatory cell exudate	
Liver					
Parameters		Score			
		0	1	2	3
Steatosis		<5%	5-33%	33-66%	>66%
Fibrosis		None	Perisinusoidal / periportal	Perisinusoidal & portal/periportal	Bridging fibrosis
Lobular inflammation (/200x field)		None	<2 foci	2-4 foci	>4 foci
Ballooning		None	Few	Many/prominent	
Kidneys					
Parameters		score			
		0	1	2	3
Glomerular cell proliferation (of glomeruli)		None	<25%	25-50%	>50%
Leukocyte exudation (polymorphonuclear leukocyte per glomerulus)		0-2	Mild	Moderate	Extensive
Cellular crescents		None	<25%	25-50%	>50%
Hyaline deposits		None	Few	Moderate	Extensive
Glomerular sclerosis		None	Mild	Moderate	Extensive
Tubular atrophy		None	Mild	Moderate	Extensive
Spleen & Thymus					
Parameters	Score				
	0	1	2	3	
Lymphoid depletion	0-10%	10-30%	30-70%	>70%	
Ovaries					
Parameters	Score				
	0	1	2	3	
Haemorrhage	Normal	Moderate	Severe		
Congestion	Normal	Moderate	Severe		
Follicular degeneration	Normal	Moderate	Severe		
Inflammation	Normal	Moderate	Severe		
Testes					
Score	Criteria				
0	Complete spermatogenesis with many spermatozoa. Germinal epithelium of regular thickness leaving an open lumen.				
1	Many spermatozoa but germinal epithelium disorganized with marked sloughing or obliteration of lumen.				
2	Only a few spermatozoa are present in the section.				
3	No spermatozoa but many spermatids present.				
4	No spermatozoa, only a few spermatids are present.				
5	No spermatozoa, spermatids but several or many spermatocytes present.				
6	Only few spermatocytes and no spermatids or spermatozoa are present.				
7	Spermatogonia are the only germ cells present.				
8	No germ cells but Sertoli cells present.				
9	No cells in the tubular section				

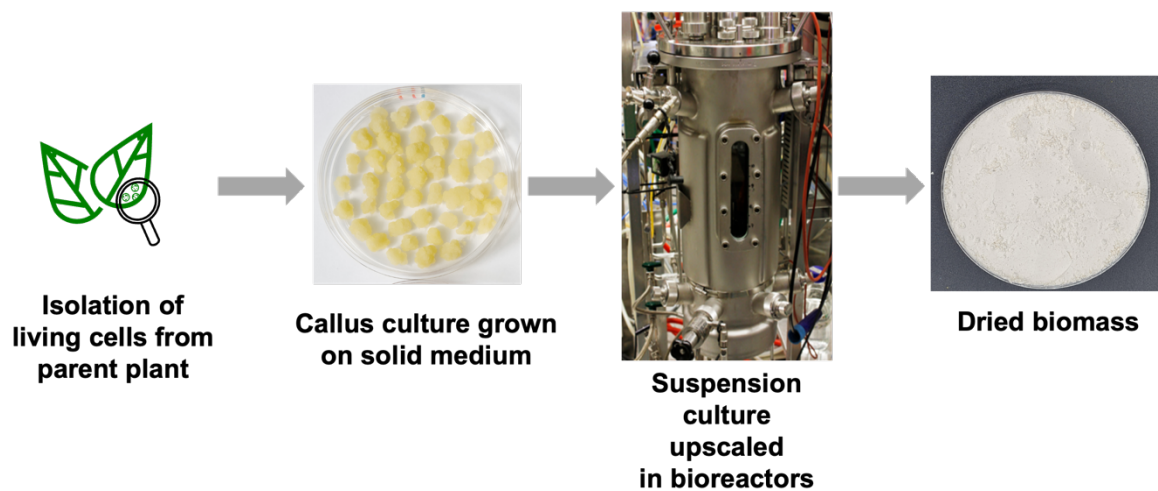
Supplementary Results

Supplementary Table 3. Probable allergens identified in plant cell cultures Scurvy Grass (*Cochlearia danica*). The Protein abundance has been reported as sum of unique peptide reporter ion intensities.

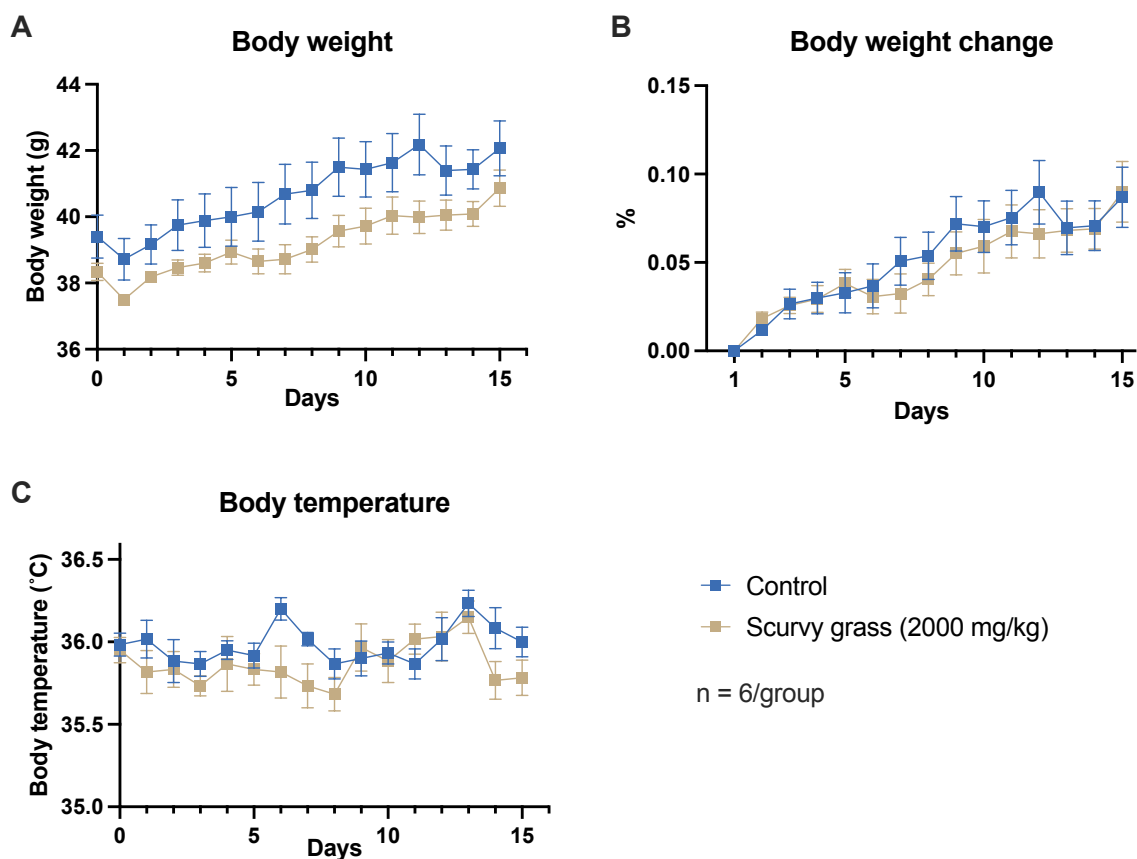
Accession UniProt	Protein	Organism	Coverage [96	Abundance	Reference
Q42418	Profilin-2 (AtPROF2) (AthPRF2)	Arabidopsis thaliana (Mouse-ear cress)	36	1577.8	89% similarity with Q42449,
Q9FUB8	Profilin	Brassica napus (Rape)	24	216	75% similarity with Q42449,
Q38905	Profilin-5 (AtPROF4) (AthPRF4)	Arabidopsis thaliana (Mouse-ear cress)	14	87	75% similarity with Q42449,
P22953	Heat shock 70 kDa protein 1	Arabidopsis thaliana (Mouse-ear cress)	48	12213.8	Reviewed in Shevchenko, Marina et al. Journal of asthma and allergy vol. 13 757-772.5 Jan. 2021, doi:10.2147/JAAS288886
Q9LKR3	Heat shock 70 kDa protein BIP1	Arabidopsis thaliana (Mouse-ear cress)	42	18977.1	
P22954	Heat shock 70 kDa protein 2	Arabidopsis thaliana (Mouse-ear cress)	41	1618.3	
O65719	Heat shock 70 kDa protein 3	Arabidopsis thaliana (Mouse-ear cress)	40	1393.1	
Q9LHA8	Heat shock 70 kDa protein 4	Arabidopsis thaliana (Mouse-ear cress)	34	224.1	
Q9LTX9	Heat shock 70 kDa protein 7	Arabidopsis thaliana (Mouse-ear cress)	24	1892.1	
Q9LDZO	Heat shock 70 kDa protein 10	Arabidopsis thaliana (Mouse-ear cress)	23	6071.8	
Q8HIB3	Heat shock 70 kDa protein BIP3	Arabidopsis thaliana (Mouse-ear cress)	10	406.3	
Q9S9N1	Heat shock 70 kDa protein 5	Arabidopsis thaliana (Mouse-ear cress)	19	571.7	
Q8GUM2	Heat shock 70 kDa protein 9	Arabidopsis thaliana (Mouse-ear cress)	16	1425.2	
Q9S7CO	Heat shock 70 kDa protein 14	Arabidopsis thaliana (Mouse-ear cress)	13	4638.3	
F4JM1	Heat shock 70 kDa protein 17	Arabidopsis thaliana (Mouse-ear cress)	10	4530.6	
P93819	Malate dehydrogenase 1	Arabidopsis thaliana (Mouse-ear cress)	28	992.1	Pastor, Coriosef. International
Q9SN86	Malate dehydrogenase	Arabidopsis thaliana (Mouse-ear cress)	16	2938	archives of allergy and immunology
Q9ZP06	Malate dehydrogenase 1	Arabidopsis thaliana (Mouse-ear cress)	16	3309.5	vol. 149,4 (2009):291-8.
O82399	Malate dehydrogenase 1	Arabidopsis thaliana (Mouse-ear cress)	16	192.1	doi:10.1159/000205574
Q9SKP6	Triosephosphate isomerase	Arabidopsis thaliana (Mouse-ear cress)	31	2009.6	Pastor, Coriosef. International
P48491	Triosephosphate isomerase	Arabidopsis thaliana (Mouse-ear cress)	29	8963.3	archives of allergy and immunology vol. 149,4 (2009):291-8. doi:10.1159/000205574; Yang, Yang et al. Molecular immunology vol. 85 (2017): 35-46. doi:10.1016/j.molimm.2017.02.004
O65570	VilHn4	Arabidopsis thaliana (Mouse-ear cress)	11	2046.3	Mittermann, Irene et al. FEBS letters vol. 579,17 (2005):3807-13. doi:10.1016/j.febslet.2005.05.066
O81644	VilHn.2	Arabidopsis thaliana (Mouse-ear cress)	18	3650.2	Ventura, Anne K R et al. The World Allergy Organization journal vol. 16,12 100845.22 Nov. 2023. doi:10.1016/j.woojou.2023.100845
Q9ZU52	Fructose-bisphosphate aldolase 3	Arabidopsis thaliana (Mouse-ear cress)	22	2181.1	Ogino, Ryohei et al. Allergy International official journal of the Japanese Society of Allergy vol. 70,2 (2021):215-222. doi:10.1016/j.alit.2020.09.005
Q9SR37	Beta-glucosidase 23 (AtBGLU23)	Arabidopsis thaliana (Mouse-ear cress)	15	1442.4	Sander, I et al. The journal of allergy and clinical immunology vol. 102,2 (1998):256-64. doi:10.1016/s0091-6749(98)70109-5
Q9FGY1	Beta-D-xylosidase 1 (AtBXL1)	Arabidopsis thaliana (Mouse-ear cress)	11	119.4	Annals of allergy, asthma & immunology: official publication of the American College of Allergy, Asthma, & Immunology vol. 106,6 (2011):545-7. doi:10.1016/j.anai.2011.03.008
P04796	Glyceraldehyde-3-phosphate dehydrogenase	Sinapis alba (White mustard) (Brassica hirta)	48	587.7	Gómez-Esquivel, Mónica Luz et al. Molecular immunology vol. 132 (2021):150-156, doi:10.1016/j.molimm.2021.01.031
PODH95	Calmodulin-I (CaM-I)	Arabidopsis thaliana (Mouse-ear cress)	54	6696.3	Nikolić, Jasna et al. Journal of proteomics vol. 175 (2018):87-94. doi:10.0016/j.jprot.2018.01.007
Q96528	Catalase-I (EC 1.11.1.6)	Arabidopsis thaliana (Mouse-ear cress)	17	624.4	Experimental allergy journal of the British Society for Allergy and Clinical Immunology vol. 34,11
Q42592	L-ascorbate peroxidase S	Arabidopsis thaliana (Mouse-ear cress)	15	385.4	

Supplementary table 4. Putative and probable allergens identified in plant cell cultures Rowan (*Sorbus aucuparia*). The protein abundance has been reported as sum of unique peptide reporter ion intensities.

Accession UniProt	Protein	Organism	Coverage [%]	Abundance	Reference
Q8GSL5	Profilin	Pyrus communis	28	283.9	UniProt
Q9XF38	Profilin	Prunus armeniaca	23	2420	
Q8GT39	Profilin	Prunus persica	18	1438.5	
Q38905	Major strawberry allergen Fra a 1.06	Fragaria ananassa	54	2629.7	UniProt
P22953	Major strawberry allergen Fra a 1-3	Fragaria ananassa	52	11020.8	
Q9LKR3	Major strawberry allergen Fra a 1.07	Fragaria ananassa	51	2883.4	
Q38906	Major strawberry allergen Fra a 1.06	Fragaria ananassa	49.33	5765	
O50001	Major allergen Pru ar 1	Prunus armeniaca	15	2026.6	UniProt
O81355	Phenylcoumaran benzylic ether reductase Pyrc5	Pyrus communis	10	2286.9	UniProt
					Gómez-Esquivel, Mónica Luz et al. <i>Molecular immunology</i> vol. 132 (2021): 150-156. doi:10.1016/j.molimm.2021.01.031
P48976	Calmodulin	Malus domestica	50	2288	Yang, Yang et al. <i>Molecular immunology</i> vol. 85 (2017): 35-46. doi:10.1016/j.molimm.2017.02.004
Q9M4S8	Triosephosphate isomerase, chloroplastic	Fragaria ananassa	31	1809.5	Pastor, Carlos et al. <i>International archives of allergy and immunology</i> vol. 149,4 (2009): 291-8. doi:10.1159/000205574
P83373	Malate dehydrogenase, mitochondrial	Fragaria ananassa	13	2439.9	



Supplementary Figure 1. Workflow of Plant Cell Culture Production from Parent Plant to Dried Biomass



Supplementary Figure 2. Results of the scurvy grass 14-day acute toxicity study. (A) Body weight of mice. (B) Relative change in body weight from the start of study. (C) Body temperature of mice. No statistical differences were noted between the two groups.

Supplementary Table 5. Haematological parameters of male ICR-CD1 mice in the 14-days acute toxicity study of Scurvy Grass (SG). Values are expressed as mean $\pm$ SD

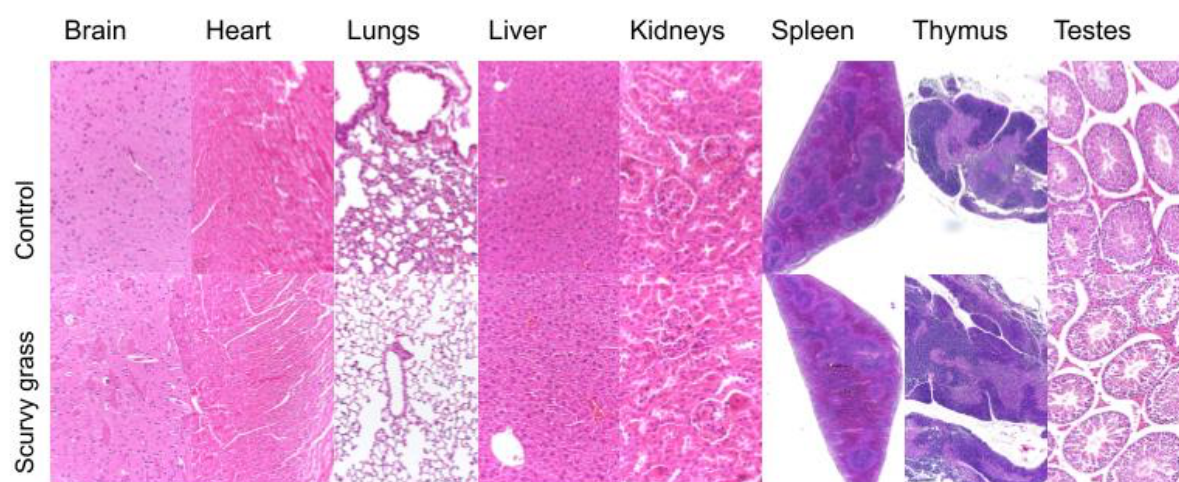
	Control (n = 6)	SG (2000 mg/kg) (n = 6)
WBCs (10^9 g/L)	2.14 $\pm$ 0.56	2.71 $\pm$ 0.86
LYM (10^9 g/L)	1.12 $\pm$ 0.34	1.17 $\pm$ 0.31
MID (10^9 g/L)	0.16 $\pm$ 0.08	0.14 $\pm$ 0.06
GRA (10^9 g/L)	0.86 $\pm$ 0.22	1.40 $\pm$ 0.63
RBCs (10^{12} g/L)	7.60 $\pm$ 0.28	7.40 $\pm$ 0.78
HGB (g/L)	127.17 $\pm$ 6.68	127.25 $\pm$ 12.51
HCT (%)	36.40 $\pm$ 11.85	39.97 $\pm$ 3.48
MCH (pg)	16.73 $\pm$ 0.36	17.24 $\pm$ 0.56
MCHC (g/L)	307.83 $\pm$ 8.93	318.42 $\pm$ 14.57*
MCV (fL)	54.35 $\pm$ 1.66	54.14 $\pm$ 2.09
Platelets (10^9 g/L)	868.83 $\pm$ 47.66	863.67 $\pm$ 145.89

* $p < 0.05$ Control vs SG

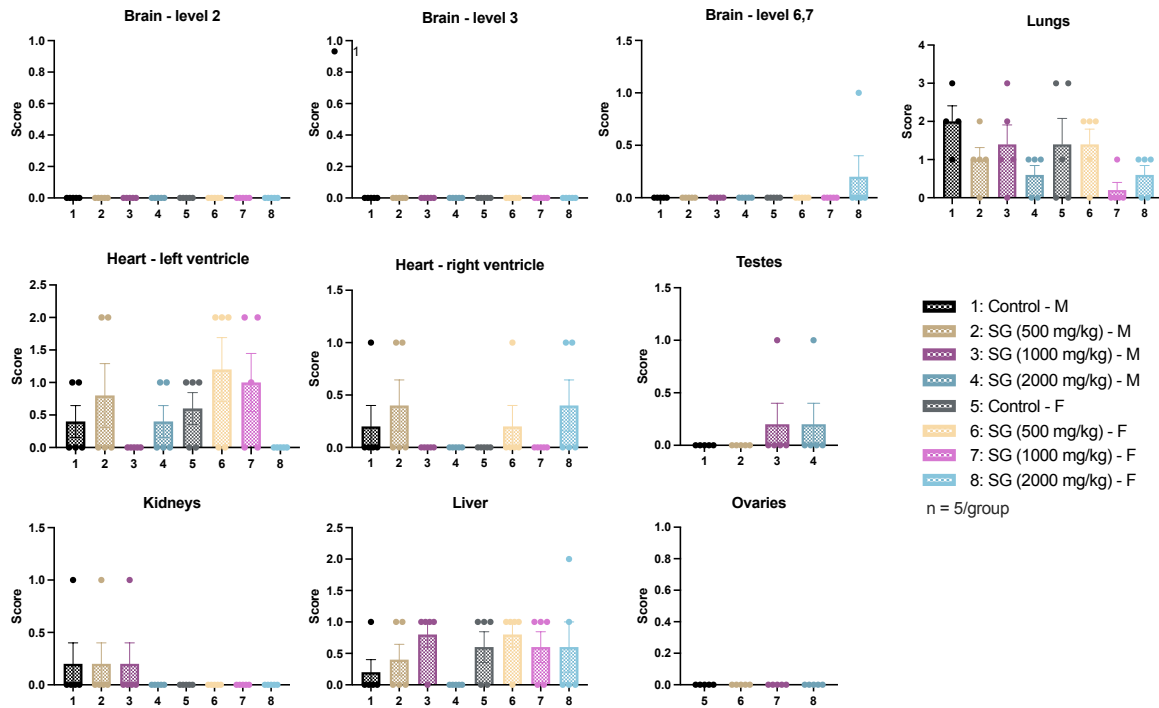
Supplementary Table 6. Organ-body weight ratio for male ICR-CD1 mice in the acute toxicity study of Scurvy Grass (SG). Values are expressed as mean $\pm$ SD

	Control (n = 6)	SG (2000 mg/kg) (n = 6)
Brain	1.13 $\pm$ 0.11	1.15 $\pm$ 0.12
Heart	0.44 $\pm$ 0.03	0.49 $\pm$ 0.05
Lungs	0.52 $\pm$ 0.06	0.59 $\pm$ 0.04*
Liver	6.04 $\pm$ 0.19	6.29 $\pm$ 0.35
Kidneys	1.60 $\pm$ 0.14	1.55 $\pm$ 0.11
Spleen	0.31 $\pm$ 0.04	0.34 $\pm$ 0.08
Thymus	0.10 $\pm$ 0.03	0.09 $\pm$ 0.02
Testes	0.67 $\pm$ 0.12	0.71 $\pm$ 0.06

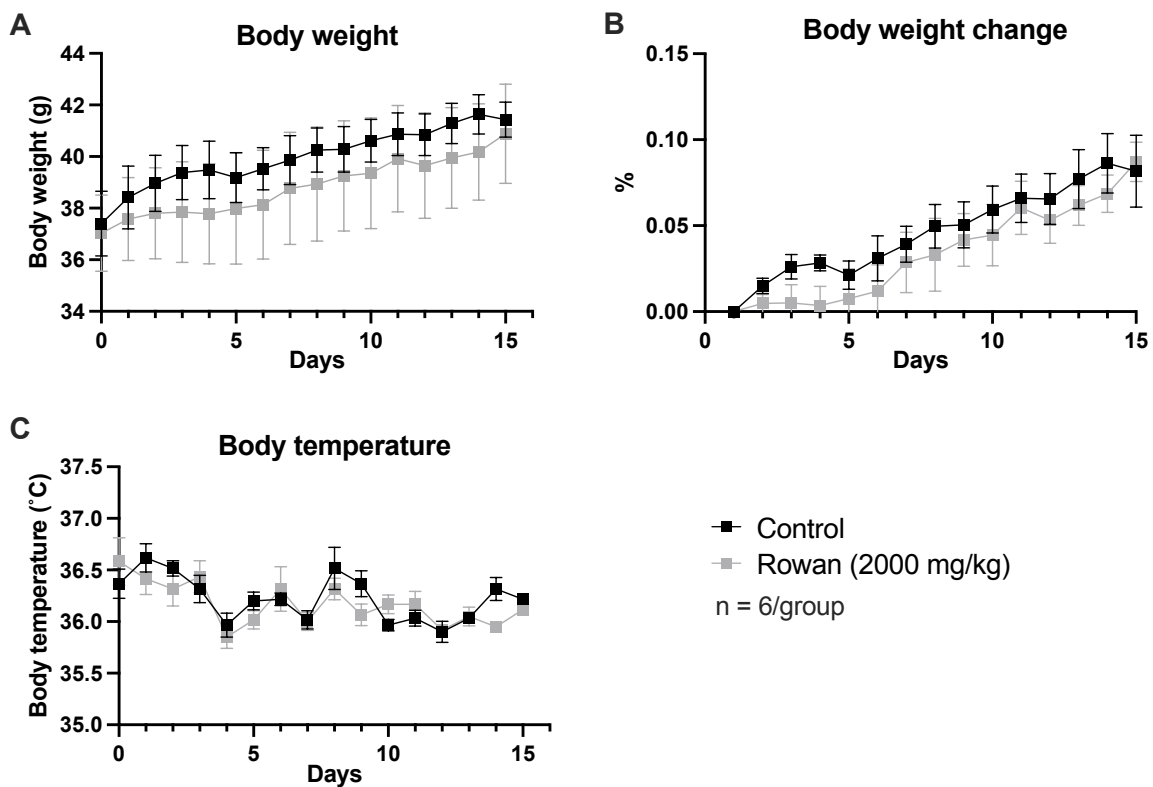
* $p < 0.05$ Control vs SG



Supplementary Figure 3. H&E-stained sections of brain (2000X), heart (2000X), lungs (200X), liver (2000X), kidneys (2000X), spleen (400X), thymus (400X) and testes (2000X) of control and SG-treated mice in the 14-day acute oral toxicity study at 2000 mg/kg.



Supplementary Figure 4. Histopathological analysis scoring of organs and tissues for the 28-day subacute oral toxicity study of Scurvy Grass (SG) cell cultures.



Supplementary Figure 5. Results of the rowanberry 14-day acute toxicity study. (A) Body weight of mice. (B) Relative change in body weight from the start of study. (C) Body temperature of mice. No statistical differences were noted between the two groups.

Supplementary Table 7. Organ-body weight ratio for male ICR-CD1 mice in the acute toxicity study of Rowan (RW). Values are expressed as mean $\pm$ SD

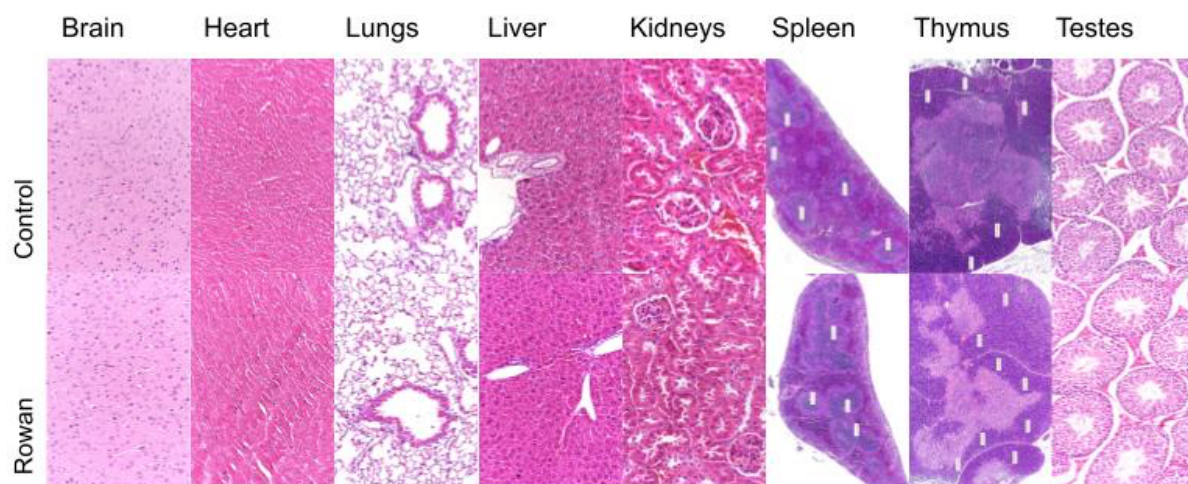
	Control (n = 6)	RW (2000 mg/kg) (n = 6)
Brain	1.09 $\pm$ 0.19	1.12 $\pm$ 0.14
Heart	0.44 $\pm$ 0.01	0.47 $\pm$ 0.05
Lungs	0.64 $\pm$ 0.04	0.68 $\pm$ 0.08
Liver	5.99 $\pm$ 0.22	5.03 $\pm$ 2.48
Kidneys	1.63 $\pm$ 0.07	1.61 $\pm$ 0.12
Spleen	0.27 $\pm$ 0.04	0.38 $\pm$ 0.17
Thymus	0.14 $\pm$ 0.03	0.12 $\pm$ 0.04
Testes	0.66 $\pm$ 0.04	0.74 $\pm$ 0.10

No statistical significance between the control group and treatment group was observed

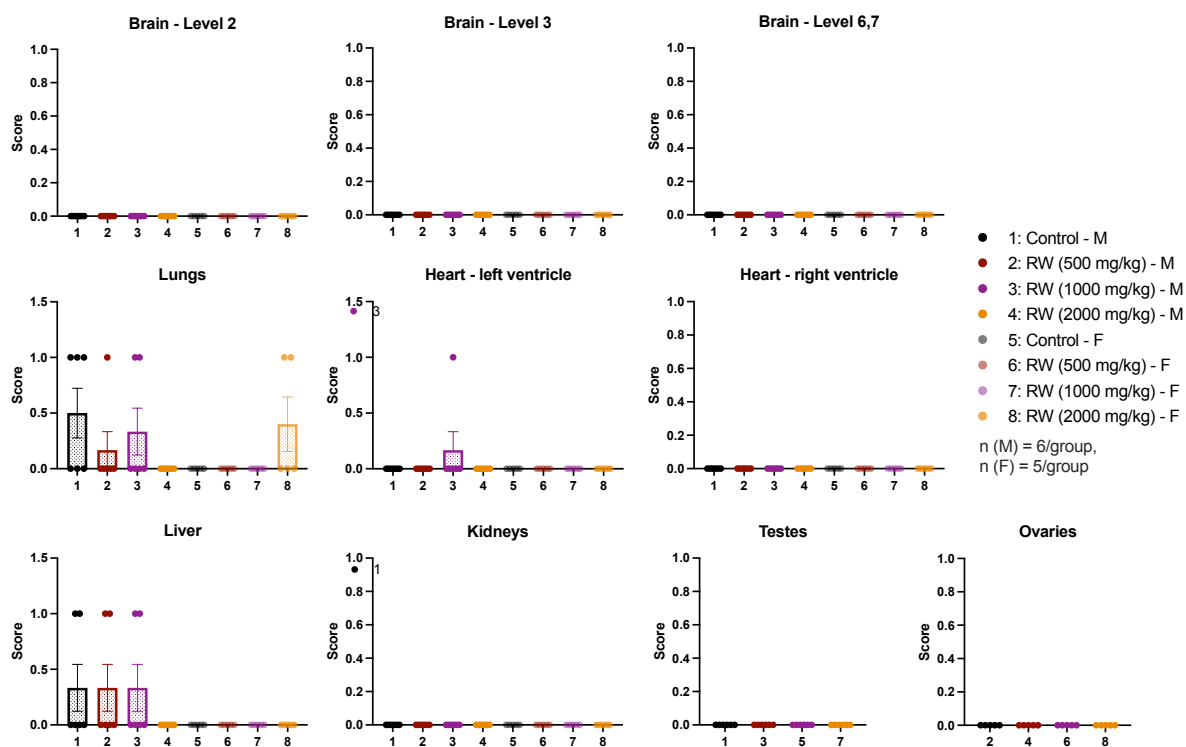
Supplementary Table 8. Haematological parameters of male ICR-CD1 mice in the 14-days acute toxicity study of Rowan (RW). Values are expressed as mean $\pm$ SD

	Control (n = 6)	RW (2000 mg/kg) (n = 6)
WBCs (10^9 g/L)	2.63 $\pm$ 0.94	3.03 $\pm$ 0.71
LYM (10^9 g/L)	1.40 $\pm$ 0.60	1.20 $\pm$ 0.40
MID (10^9 g/L)	0.18 $\pm$ 0.09	0.18 $\pm$ 0.06
GRA (10^9 g/L)	1.06 $\pm$ 0.44	1.65 $\pm$ 0.53
RBCs (10^{12} g/L)	7.59 $\pm$ 0.28	7.69 $\pm$ 0.26
HGB (g/L)	128.50 $\pm$ 5.39	127.50 $\pm$ 5.13
HCT (%)	301.17 $\pm$ 4.45	305.50 $\pm$ 4.55
MCH (pg)	16.95 $\pm$ 0.23	16.62 $\pm$ 0.57
MCHC (g/L)	56.20 $\pm$ 0.93	54.43 $\pm$ 2.15
MCV (fL)	42.67 $\pm$ 1.77	41.82 $\pm$ 1.64
Platelets (10^9 g/L)	802.33 $\pm$ 44.57	765.50 $\pm$ 87.24

No statistical significance between the control group and treatment group was observed



Supplementary Figure 6. Haematoxylin and eosin-stained sections of brain (200X), heart (200X), lungs (200X), liver (200X), kidneys (200X), spleen (40X), thymus (40X) and testes (200X) of control and Rowan-treated mice in 14-day acute oral toxicity study.



Supplementary Figure 7. Histopathological analysis scoring of organs and tissues for the 28-day subacute oral toxicity study of rowan (RW) cell cultures.