

SUPPLEMENTARY INFORMATION

ARTICLE TITLE: TRPC5 ion channel permeation promotes weight gain in hypercholesterolaemic mice

AUTHORS: Baptiste Rode PhD, Nadira Y Yuldasheva PhD, Paul D Baxter PhD, Alicia Sedo BSc, Justin F Ainscough PhD, Michael Shires BSc, Mark T Kearney MD, Marc A Bailey MD, Stephen B Wheatcroft MD and David J Beech PhD*

* Corresponding author: Professor David J Beech, Leeds Institute of Cardiovascular and Metabolic Medicine, LIGHT Building, Clarendon Way, School of Medicine, University of Leeds, Leeds LS2 9JT, UK. E-mail d.j.beech@leeds.ac.uk. Telephone +44 (0) 113 3434323.

AUTHOR AFFILIATION: School of Medicine, University of Leeds, Leeds, LS2 9JT, UK

Supplementary File 1. Full-length gels from Figure 1C.

Supplementary File 2. Detailed values for each genotype.

Supplementary File 3. Raw data for each parameter analysed.

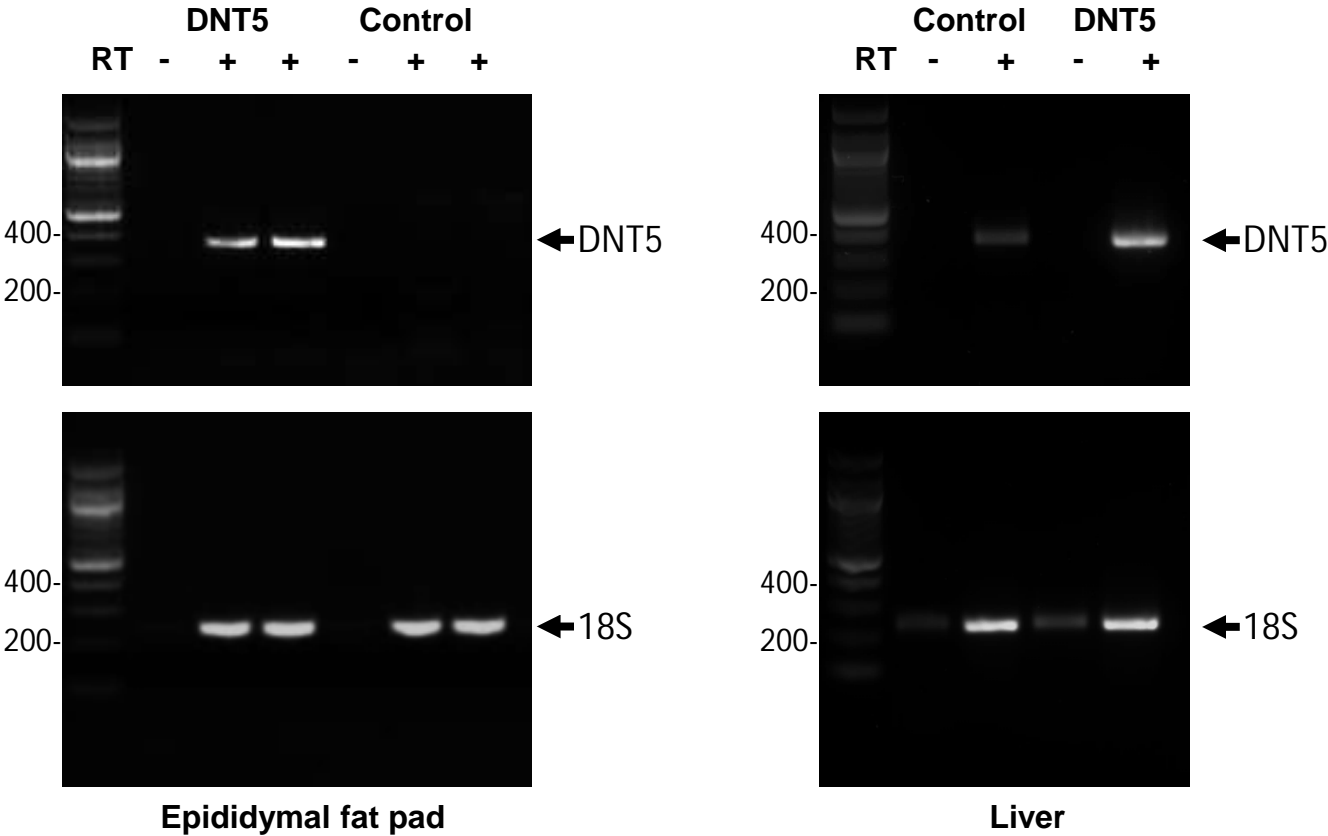
Supplementary File 4. Detailed statistical analysis.

SUPPLEMENTARY INFORMATION LEGEND

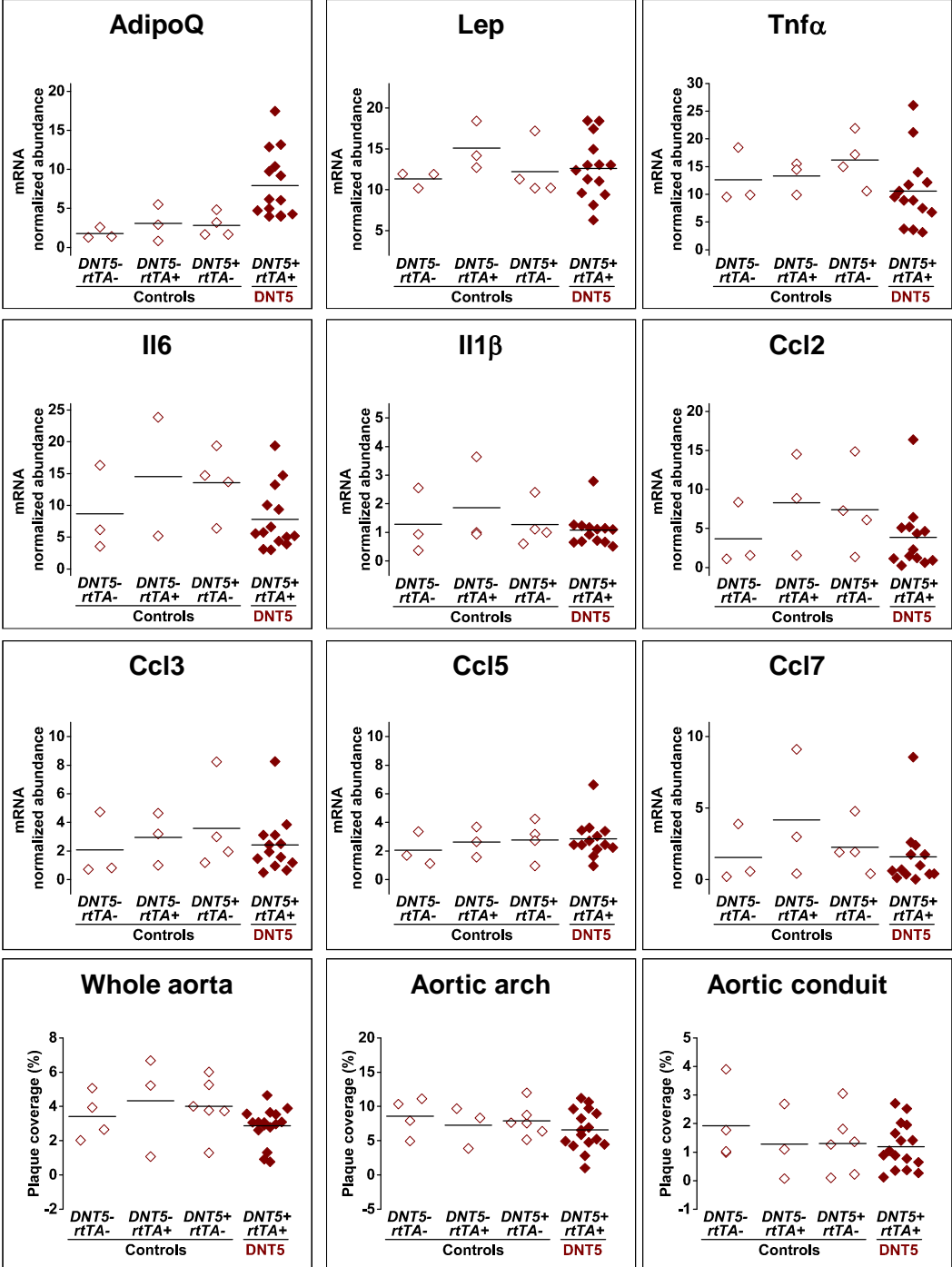
Supplementary File 1. Full-length gels from Figure 1C. Gel-electrophoresis of end-point RT-PCR performed on epididymal fat pad (left) and liver (right). Results are shown without (-) and with (+) reverse transcription (RT). RT+ reactions were loaded in duplicates for epididymal fat pad. The sizes of the DNA markers (New England Biolabs 100bp ladder) are indicated on the left in base-pairs. The arrows point to the expected PCR product sizes. DNT5 mRNA was detected in tissues from DOX-treated double transgenic mice but not Controls.

Supplementary File 2. Detailed values for each genotype. Individual values for each genotype is shown for each parameter of the present study. Mean value is shown with a horizontal bar. The genotype of each group is indicated in italic on the bottom of each scatter plot: “*DNT5*” refers to the transgene carrying the DNT5 coding sequence under the control of the Tetracycline Response Element, “*rtTA*” refers to the transgene allowing the expression of the rtTA transactivator from the ROSA26 locus. Absence and presence of each transgene are denoted by “-” and “+”, respectively. The corresponding groups are indicated underneath.

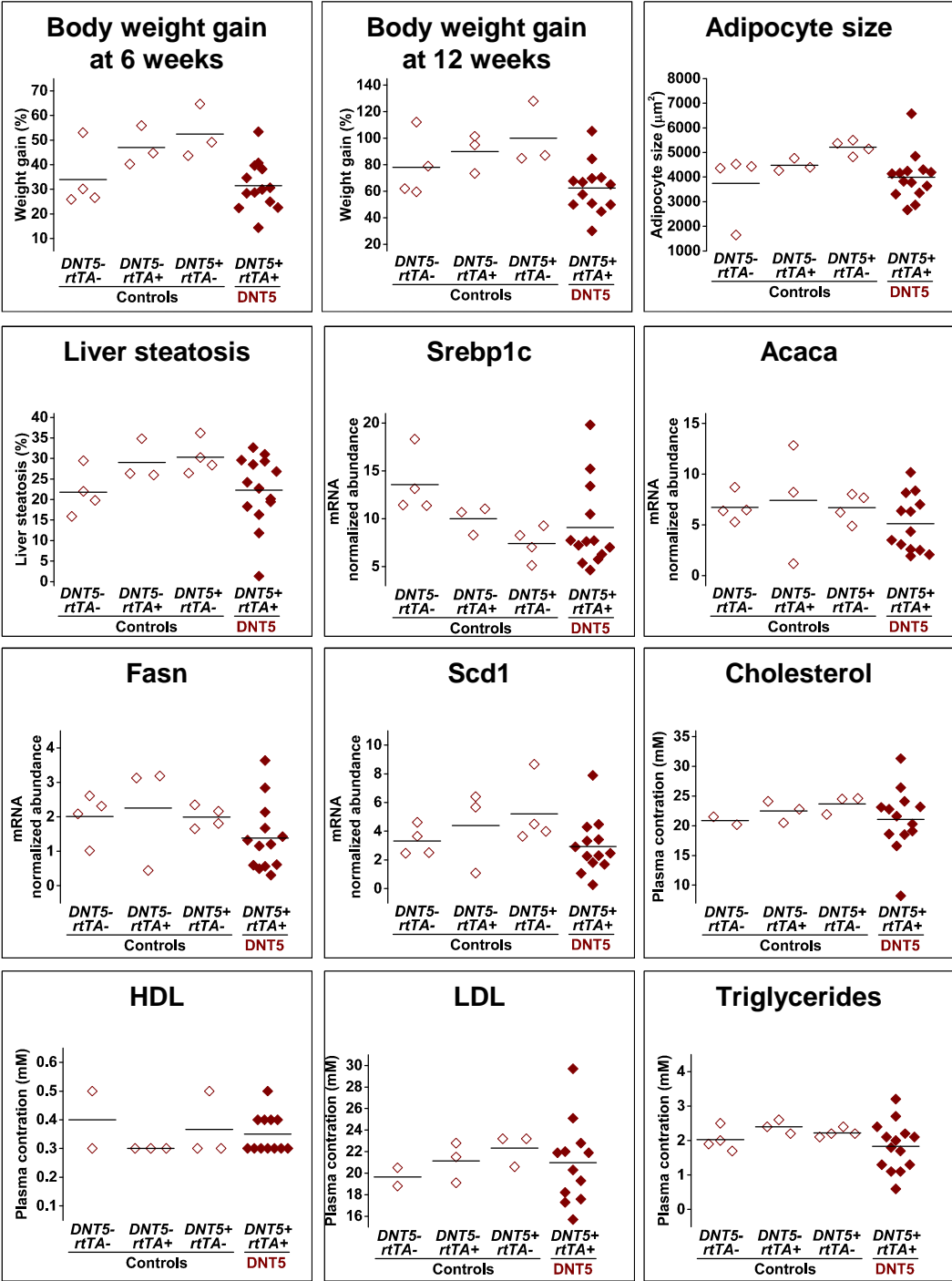
Supplementary File 1



Supplementary File 2 (Part1/2)



Supplementary File 2 (Part2/2)



Supplementary File 3

ID	GENOTYPE	GROUP	CAGE	Adipose tissue Adiponectin mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	4.0
113	DNT5+/rtTA-	CT	4	1.7
118	DNT5+/rtTA+	DN	5	4.1
119	DNT5-/rtTA+	CT	5	2.9
120	DNT5+/rtTA+	DN	5	5.0
121	DNT5+/rtTA+	DN	6	4.0
122	DNT5-/rtTA-	CT	6	2.6
123	DNT5+/rtTA+	DN	6	4.3
124	DNT5+/rtTA+	DN	7	6.0
125	DNT5+/rtTA-	CT	7	3.2
126	DNT5+/rtTA-	CT	7	4.8
127	DNT5+/rtTA+	DN	7	13.2
128	DNT5+/rtTA+	DN	8	4.7
129	DNT5+/rtTA+	DN	8	12.9
130	DNT5-/rtTA+	CT	8	5.5
134	DNT5+/rtTA-	CT	10	1.7
135	DNT5+/rtTA+	DN	10	10.4
136	DNT5+/rtTA+	DN	10	17.5
137	DNT5+/rtTA+	DN	11	9.2
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	1.4
140	DNT5+/rtTA+	DN	12	6.2
141	DNT5-/rtTA+	CT	12	0.8
142	DNT5-/rtTA-	CT	12	1.3
143	DNT5+/rtTA+	DN	12	9.7

ID	GENOTYPE	GROUP	CAGE	Adipose tissue Leptin mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	11.0
113	DNT5+/rtTA-	CT	4	10.2
118	DNT5+/rtTA+	DN	5	11.3
119	DNT5-/rtTA+	CT	5	14.2
120	DNT5+/rtTA+	DN	5	18.4
121	DNT5+/rtTA+	DN	6	18.4
122	DNT5-/rtTA-	CT	6	10.2
123	DNT5+/rtTA+	DN	6	8.1
124	DNT5+/rtTA+	DN	7	9.6
125	DNT5+/rtTA-	CT	7	17.2
126	DNT5+/rtTA-	CT	7	11.3
127	DNT5+/rtTA+	DN	7	17.4
128	DNT5+/rtTA+	DN	8	13.1
129	DNT5+/rtTA+	DN	8	15.0
130	DNT5-/rtTA+	CT	8	12.7
134	DNT5+/rtTA-	CT	10	10.2
135	DNT5+/rtTA+	DN	10	9.4
136	DNT5+/rtTA+	DN	10	6.3
137	DNT5+/rtTA+	DN	11	13.0
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	11.9
140	DNT5+/rtTA+	DN	12	13.0
141	DNT5-/rtTA+	CT	12	18.4
142	DNT5-/rtTA-	CT	12	11.9
143	DNT5+/rtTA+	DN	12	12.4

ID	GENOTYPE	GROUP	CAGE	Adipose tissue TNF α mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	8.9
113	DNT5+/rtTA-	CT	4	10.6
118	DNT5+/rtTA+	DN	5	8.9
119	DNT5-/rtTA+	CT	5	15.5
120	DNT5+/rtTA+	DN	5	21.2
121	DNT5+/rtTA+	DN	6	14.0
122	DNT5-/rtTA-	CT	6	18.4
123	DNT5+/rtTA+	DN	6	7.5
124	DNT5+/rtTA+	DN	7	3.6
125	DNT5+/rtTA-	CT	7	21.9
126	DNT5+/rtTA-	CT	7	15.0
127	DNT5+/rtTA+	DN	7	11.7
128	DNT5+/rtTA+	DN	8	26.1
129	DNT5+/rtTA+	DN	8	12.2
130	DNT5-/rtTA+	CT	8	14.5
134	DNT5+/rtTA-	CT	10	17.2
135	DNT5+/rtTA+	DN	10	9.5
136	DNT5+/rtTA+	DN	10	6.7
137	DNT5+/rtTA+	DN	11	3.7
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	9.9
140	DNT5+/rtTA+	DN	12	10.6
141	DNT5-/rtTA+	CT	12	9.9
142	DNT5-/rtTA-	CT	12	9.5
143	DNT5+/rtTA+	DN	12	3.1

ID	GENOTYPE	GROUP	CAGE	Adipose tissue Il6 mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	9.4
113	DNT5+/rtTA-	CT	4	13.7
118	DNT5+/rtTA+	DN	5	13.2
119	DNT5-/rtTA+	CT	5	23.8
120	DNT5+/rtTA+	DN	5	19.4
121	DNT5+/rtTA+	DN	6	10.0
122	DNT5-/rtTA-	CT	6	16.3
123	DNT5+/rtTA+	DN	6	4.4
124	DNT5+/rtTA+	DN	7	3.0
125	DNT5+/rtTA-	CT	7	19.4
126	DNT5+/rtTA-	CT	7	14.7
127	DNT5+/rtTA+	DN	7	6.6
128	DNT5+/rtTA+	DN	8	14.7
129	DNT5+/rtTA+	DN	8	5.0
130	DNT5-/rtTA+	CT	8	5.2
134	DNT5+/rtTA-	CT	10	6.4
135	DNT5+/rtTA+	DN	10	5.8
136	DNT5+/rtTA+	DN	10	5.2
137	DNT5+/rtTA+	DN	11	3.9
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	6.2
140	DNT5+/rtTA+	DN	12	5.6
141	DNT5-/rtTA+	CT	12	
142	DNT5-/rtTA-	CT	12	3.5
143	DNT5+/rtTA+	DN	12	3.1

ID	GENOTYPE	GROUP	CAGE	Adipose tissue Il1b mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	1.11
113	DNT5+/rtTA-	CT	4	1.11
118	DNT5+/rtTA+	DN	5	0.71
119	DNT5-/rtTA+	CT	5	3.64
120	DNT5+/rtTA+	DN	5	1.17
121	DNT5+/rtTA+	DN	6	0.92
122	DNT5-/rtTA-	CT	6	2.55
123	DNT5+/rtTA+	DN	6	1.15
124	DNT5+/rtTA+	DN	7	0.67
125	DNT5+/rtTA-	CT	7	2.40
126	DNT5+/rtTA-	CT	7	1.00
127	DNT5+/rtTA+	DN	7	1.23
128	DNT5+/rtTA+	DN	8	2.79
129	DNT5+/rtTA+	DN	8	1.10
130	DNT5-/rtTA+	CT	8	1.00
134	DNT5+/rtTA-	CT	10	0.60
135	DNT5+/rtTA+	DN	10	0.68
136	DNT5+/rtTA+	DN	10	1.27
137	DNT5+/rtTA+	DN	11	0.51
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	0.37
140	DNT5+/rtTA+	DN	12	0.66
141	DNT5-/rtTA+	CT	12	0.94
142	DNT5-/rtTA-	CT	12	0.93
143	DNT5+/rtTA+	DN	12	

ID	GENOTYPE	GROUP	CAGE	Adipose tissue Ccl2 mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	4.35
113	DNT5+/rtTA-	CT	4	6.10
118	DNT5+/rtTA+	DN	5	5.21
119	DNT5-/rtTA+	CT	5	14.50
120	DNT5+/rtTA+	DN	5	6.43
121	DNT5+/rtTA+	DN	6	4.65
122	DNT5-/rtTA-	CT	6	8.37
123	DNT5+/rtTA+	DN	6	2.29
124	DNT5+/rtTA+	DN	7	1.21
125	DNT5+/rtTA-	CT	7	14.87
126	DNT5+/rtTA-	CT	7	7.29
127	DNT5+/rtTA+	DN	7	5.10
128	DNT5+/rtTA+	DN	8	16.37
129	DNT5+/rtTA+	DN	8	1.46
130	DNT5-/rtTA+	CT	8	8.85
134	DNT5+/rtTA-	CT	10	1.35
135	DNT5+/rtTA+	DN	10	0.61
136	DNT5+/rtTA+	DN	10	0.26
137	DNT5+/rtTA+	DN	11	0.91
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	1.54
140	DNT5+/rtTA+	DN	12	1.15
141	DNT5-/rtTA+	CT	12	1.54
142	DNT5-/rtTA-	CT	12	1.09
143	DNT5+/rtTA+	DN	12	

ID	GENOTYPE	GROUP	CAGE	Adipose tissue Ccl3 mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	2.50
113	DNT5+/rtTA-	CT	4	1.95
118	DNT5+/rtTA+	DN	5	1.57
119	DNT5-/rtTA+	CT	5	4.64
120	DNT5+/rtTA+	DN	5	3.11
121	DNT5+/rtTA+	DN	6	3.11
122	DNT5-/rtTA-	CT	6	4.73
123	DNT5+/rtTA+	DN	6	1.93
124	DNT5+/rtTA+	DN	7	1.18
125	DNT5+/rtTA-	CT	7	8.23
126	DNT5+/rtTA-	CT	7	2.98
127	DNT5+/rtTA+	DN	7	3.84
128	DNT5+/rtTA+	DN	8	8.25
129	DNT5+/rtTA+	DN	8	2.42
130	DNT5-/rtTA+	CT	8	3.19
134	DNT5+/rtTA-	CT	10	1.18
135	DNT5+/rtTA+	DN	10	0.96
136	DNT5+/rtTA+	DN	10	0.49
137	DNT5+/rtTA+	DN	11	0.66
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	0.81
140	DNT5+/rtTA+	DN	12	1.46
141	DNT5-/rtTA+	CT	12	1.00
142	DNT5-/rtTA-	CT	12	0.71
143	DNT5+/rtTA+	DN	12	

ID	GENOTYPE	GROUP	CAGE	Adipose tissue Ccl5 mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	3.03
113	DNT5+/rtTA-	CT	4	3.18
118	DNT5+/rtTA+	DN	5	2.11
119	DNT5-/rtTA+	CT	5	3.67
120	DNT5+/rtTA+	DN	5	3.62
121	DNT5+/rtTA+	DN	6	3.39
122	DNT5-/rtTA-	CT	6	3.35
123	DNT5+/rtTA+	DN	6	2.71
124	DNT5+/rtTA+	DN	7	2.44
125	DNT5+/rtTA-	CT	7	4.24
126	DNT5+/rtTA-	CT	7	2.73
127	DNT5+/rtTA+	DN	7	2.43
128	DNT5+/rtTA+	DN	8	6.63
129	DNT5+/rtTA+	DN	8	2.24
130	DNT5-/rtTA+	CT	8	2.64
134	DNT5+/rtTA-	CT	10	0.96
135	DNT5+/rtTA+	DN	10	2.43
136	DNT5+/rtTA+	DN	10	0.97
137	DNT5+/rtTA+	DN	11	1.63
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	1.12
140	DNT5+/rtTA+	DN	12	3.45
141	DNT5-/rtTA+	CT	12	1.57
142	DNT5-/rtTA-	CT	12	1.69
143	DNT5+/rtTA+	DN	12	

ID	GENOTYPE	GROUP	CAGE	Adipose tissue Ccl7 mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	2.41
113	DNT5+/rtTA-	CT	4	1.94
118	DNT5+/rtTA+	DN	5	0.04
119	DNT5-/rtTA+	CT	5	9.11
120	DNT5+/rtTA+	DN	5	2.60
121	DNT5+/rtTA+	DN	6	1.76
122	DNT5-/rtTA-	CT	6	3.89
123	DNT5+/rtTA+	DN	6	0.99
124	DNT5+/rtTA+	DN	7	0.38
125	DNT5+/rtTA-	CT	7	4.79
126	DNT5+/rtTA-	CT	7	1.91
127	DNT5+/rtTA+	DN	7	1.76
128	DNT5+/rtTA+	DN	8	8.56
129	DNT5+/rtTA+	DN	8	0.70
130	DNT5-/rtTA+	CT	8	2.99
134	DNT5+/rtTA-	CT	10	0.42
135	DNT5+/rtTA+	DN	10	0.39
136	DNT5+/rtTA+	DN	10	0.12
137	DNT5+/rtTA+	DN	11	0.41
138	DNT5-/rtTA-	CT	11	
139	DNT5-/rtTA-	CT	11	0.57
140	DNT5+/rtTA+	DN	12	0.61
141	DNT5-/rtTA+	CT	12	0.42
142	DNT5-/rtTA-	CT	12	0.21
143	DNT5+/rtTA+	DN	12	

ID	GENOTYPE	GROUP	CAGE	Whole aorta atherosclerosis area coverage (%)
49	DNT5+/rtTA-	CT	1	1.30
50	DNT5+/rtTA+	DN	1	1.31
51	DNT5+/rtTA-	CT	2	6.01
52	DNT5+/rtTA+	DN	2	0.92
112	DNT5+/rtTA+	DN	4	2.79
113	DNT5+/rtTA-	CT	4	3.76
118	DNT5+/rtTA+	DN	5	3.65
119	DNT5-/rtTA+	CT	5	5.22
120	DNT5+/rtTA+	DN	5	3.06
121	DNT5+/rtTA+	DN	6	2.91
122	DNT5-/rtTA-	CT	6	3.93
123	DNT5+/rtTA+	DN	6	3.54
124	DNT5+/rtTA+	DN	7	3.07
125	DNT5+/rtTA-	CT	7	4.00
126	DNT5+/rtTA-	CT	7	3.74
127	DNT5+/rtTA+	DN	7	3.10
128	DNT5+/rtTA+	DN	8	3.06
129	DNT5+/rtTA+	DN	8	3.89
130	DNT5-/rtTA+	CT	8	1.07
134	DNT5+/rtTA-	CT	10	5.26
135	DNT5+/rtTA+	DN	10	4.64
136	DNT5+/rtTA+	DN	10	0.76
137	DNT5+/rtTA+	DN	11	2.97
138	DNT5-/rtTA-	CT	11	2.65
139	DNT5-/rtTA-	CT	11	2.02
140	DNT5+/rtTA+	DN	12	3.57
141	DNT5-/rtTA+	CT	12	6.68
142	DNT5-/rtTA-	CT	12	5.07
143	DNT5+/rtTA+	DN	12	2.62

ID	GENOTYPE	GROUP	CAGE	Aortic arch atherosclerosis area coverage (%)
49	DNT5+/rtTA-	CT	1	5.14
50	DNT5+/rtTA+	DN	1	4.91
51	DNT5+/rtTA-	CT	2	11.99
52	DNT5+/rtTA+	DN	2	2.82
112	DNT5+/rtTA+	DN	4	9.72
113	DNT5+/rtTA-	CT	4	7.58
118	DNT5+/rtTA+	DN	5	4.77
119	DNT5-/rtTA+	CT	5	8.30
120	DNT5+/rtTA+	DN	5	5.85
121	DNT5+/rtTA+	DN	6	6.93
122	DNT5-/rtTA-	CT	6	7.92
123	DNT5+/rtTA+	DN	6	8.25
124	DNT5+/rtTA+	DN	7	10.64
125	DNT5+/rtTA-	CT	7	7.63
126	DNT5+/rtTA-	CT	7	8.73
127	DNT5+/rtTA+	DN	7	6.55
128	DNT5+/rtTA+	DN	8	5.22
129	DNT5+/rtTA+	DN	8	4.26
130	DNT5-/rtTA+	CT	8	3.87
134	DNT5+/rtTA-	CT	10	6.38
135	DNT5+/rtTA+	DN	10	4.45
136	DNT5+/rtTA+	DN	10	1.00
137	DNT5+/rtTA+	DN	11	8.95
138	DNT5-/rtTA-	CT	11	11.13
139	DNT5-/rtTA-	CT	11	4.94
140	DNT5+/rtTA+	DN	12	9.65
141	DNT5-/rtTA+	CT	12	9.68
142	DNT5-/rtTA-	CT	12	10.34
143	DNT5+/rtTA+	DN	12	11.19

ID	GENOTYPE	GROUP	CAGE	Aortic conduit atherosclerosis area
49	DNT5+/rtTA-	CT	1	0.10
50	DNT5+/rtTA+	DN	1	0.27
51	DNT5+/rtTA-	CT	2	1.81
52	DNT5+/rtTA+	DN	2	0.12
112	DNT5+/rtTA+	DN	4	0.77
113	DNT5+/rtTA-	CT	4	1.36
118	DNT5+/rtTA+	DN	5	1.40
119	DNT5-/rtTA+	CT	5	1.10
120	DNT5+/rtTA+	DN	5	1.96
121	DNT5+/rtTA+	DN	6	2.02
122	DNT5-/rtTA-	CT	6	1.77
123	DNT5+/rtTA+	DN	6	1.66
124	DNT5+/rtTA+	DN	7	0.89
125	DNT5+/rtTA-	CT	7	0.22
126	DNT5+/rtTA-	CT	7	1.27
127	DNT5+/rtTA+	DN	7	0.65
128	DNT5+/rtTA+	DN	8	2.52
129	DNT5+/rtTA+	DN	8	1.05
130	DNT5-/rtTA+	CT	8	0.07
134	DNT5+/rtTA-	CT	10	3.05
135	DNT5+/rtTA+	DN	10	2.70
136	DNT5+/rtTA+	DN	10	0.37
137	DNT5+/rtTA+	DN	11	0.90
138	DNT5-/rtTA-	CT	11	0.98
139	DNT5-/rtTA-	CT	11	1.03
140	DNT5+/rtTA+	DN	12	1.41
141	DNT5-/rtTA+	CT	12	2.69
142	DNT5-/rtTA-	CT	12	3.90
143	DNT5+/rtTA+	DN	12	0.35

ID	GENOTYPE	GROUP	CAGE	Weight gain 6 weeks (%)
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	
113	DNT5+/rtTA-	CT	4	
118	DNT5+/rtTA+	DN	5	30.0
119	DNT5-/rtTA+	CT	5	55.9
120	DNT5+/rtTA+	DN	5	38.2
121	DNT5+/rtTA+	DN	6	40.7
122	DNT5-/rtTA-	CT	6	53.0
123	DNT5+/rtTA+	DN	6	39.7
124	DNT5+/rtTA+	DN	7	28.7
125	DNT5+/rtTA-	CT	7	64.7
126	DNT5+/rtTA-	CT	7	49.1
127	DNT5+/rtTA+	DN	7	30.7
128	DNT5+/rtTA+	DN	8	53.4
129	DNT5+/rtTA+	DN	8	34.7
130	DNT5-/rtTA+	CT	8	44.8
134	DNT5+/rtTA-	CT	10	43.7
135	DNT5+/rtTA+	DN	10	25.0
136	DNT5+/rtTA+	DN	10	14.4
137	DNT5+/rtTA+	DN	11	28.4
138	DNT5-/rtTA-	CT	11	26.6
139	DNT5-/rtTA-	CT	11	25.9
140	DNT5+/rtTA+	DN	12	22.6
141	DNT5-/rtTA+	CT	12	40.2
142	DNT5-/rtTA-	CT	12	30.2
143	DNT5+/rtTA+	DN	12	22.4

ID	GENOTYPE	GROUP	CAGE	Weight gain 12 weeks (%)
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	
113	DNT5+/rtTA-	CT	4	
118	DNT5+/rtTA+	DN	5	69.6
119	DNT5-/rtTA+	CT	5	94.9
120	DNT5+/rtTA+	DN	5	66.9
121	DNT5+/rtTA+	DN	6	84.5
122	DNT5-/rtTA-	CT	6	112.1
123	DNT5+/rtTA+	DN	6	70.3
124	DNT5+/rtTA+	DN	7	50.9
125	DNT5+/rtTA-	CT	7	127.9
126	DNT5+/rtTA-	CT	7	87.2
127	DNT5+/rtTA+	DN	7	67.6
128	DNT5+/rtTA+	DN	8	105.3
129	DNT5+/rtTA+	DN	8	57.5
130	DNT5-/rtTA+	CT	8	101.5
134	DNT5+/rtTA-	CT	10	84.9
135	DNT5+/rtTA+	DN	10	44.7
136	DNT5+/rtTA+	DN	10	30.0
137	DNT5+/rtTA+	DN	11	65.1
138	DNT5-/rtTA-	CT	11	59.6
139	DNT5-/rtTA-	CT	11	78.9
140	DNT5+/rtTA+	DN	12	50.0
141	DNT5-/rtTA+	CT	12	73.4
142	DNT5-/rtTA-	CT	12	62.0
143	DNT5+/rtTA+	DN	12	49.8

ID	GENOTYPE	GROUP	CAGE	Adipocyte Area (μm^2)
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	4847.8
113	DNT5+/rtTA-	CT	4	4819.1
118	DNT5+/rtTA+	DN	5	4243.9
119	DNT5-/rtTA+	CT	5	4764.1
120	DNT5+/rtTA+	DN	5	4304.2
121	DNT5+/rtTA+	DN	6	4156.1
122	DNT5-/rtTA-	CT	6	4528.6
123	DNT5+/rtTA+	DN	6	3780.8
124	DNT5+/rtTA+	DN	7	2862.6
125	DNT5+/rtTA-	CT	7	5498.6
126	DNT5+/rtTA-	CT	7	5367.7
127	DNT5+/rtTA+	DN	7	3822.6
128	DNT5+/rtTA+	DN	8	6574.7
129	DNT5+/rtTA+	DN	8	3349.3
130	DNT5-/rtTA+	CT	8	4262.4
134	DNT5+/rtTA-	CT	10	5143.1
135	DNT5+/rtTA+	DN	10	3305.9
136	DNT5+/rtTA+	DN	10	2666.2
137	DNT5+/rtTA+	DN	11	3634.2
138	DNT5-/rtTA-	CT	11	4357.5
139	DNT5-/rtTA-	CT	11	4433.5
140	DNT5+/rtTA+	DN	12	4192.4
141	DNT5-/rtTA+	CT	12	4396.4
142	DNT5-/rtTA-	CT	12	1645.3
143	DNT5+/rtTA+	DN	12	4131.8

ID	GENOTYPE	GROUP	CAGE	Liver steatosis area coverage (%)
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	29.4
113	DNT5+/rtTA-	CT	4	36.2
118	DNT5+/rtTA+	DN	5	28.5
119	DNT5-/rtTA+	CT	5	26.0
120	DNT5+/rtTA+	DN	5	31.0
121	DNT5+/rtTA+	DN	6	22.7
122	DNT5-/rtTA-	CT	6	29.4
123	DNT5+/rtTA+	DN	6	16.3
124	DNT5+/rtTA+	DN	7	11.8
125	DNT5+/rtTA-	CT	7	28.4
126	DNT5+/rtTA-	CT	7	30.2
127	DNT5+/rtTA+	DN	7	26.8
128	DNT5+/rtTA+	DN	8	32.6
129	DNT5+/rtTA+	DN	8	24.2
130	DNT5-/rtTA+	CT	8	34.8
134	DNT5+/rtTA-	CT	10	26.4
135	DNT5+/rtTA+	DN	10	20.2
136	DNT5+/rtTA+	DN	10	1.4
137	DNT5+/rtTA+	DN	11	18.3
138	DNT5-/rtTA-	CT	11	19.8
139	DNT5-/rtTA-	CT	11	22.0
140	DNT5+/rtTA+	DN	12	19.4
141	DNT5-/rtTA+	CT	12	26.3
142	DNT5-/rtTA-	CT	12	15.9
143	DNT5+/rtTA+	DN	12	29.6

ID	GENOTYPE	GROUP	CAGE	Liver Srebp1c mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	4.6
113	DNT5+/rtTA-	CT	4	5.1
118	DNT5+/rtTA+	DN	5	7.7
119	DNT5-/rtTA+	CT	5	11.0
120	DNT5+/rtTA+	DN	5	7.6
121	DNT5+/rtTA+	DN	6	6.3
122	DNT5-/rtTA-	CT	6	11.4
123	DNT5+/rtTA+	DN	6	5.4
124	DNT5+/rtTA+	DN	7	7.2
125	DNT5+/rtTA-	CT	7	9.3
126	DNT5+/rtTA-	CT	7	8.2
127	DNT5+/rtTA+	DN	7	7.0
128	DNT5+/rtTA+	DN	8	13.4
129	DNT5+/rtTA+	DN	8	10.5
130	DNT5-/rtTA+	CT	8	10.7
134	DNT5+/rtTA-	CT	10	7.0
135	DNT5+/rtTA+	DN	10	7.7
136	DNT5+/rtTA+	DN	10	
137	DNT5+/rtTA+	DN	11	19.8
138	DNT5-/rtTA-	CT	11	18.3
139	DNT5-/rtTA-	CT	11	13.1
140	DNT5+/rtTA+	DN	12	15.2
141	DNT5-/rtTA+	CT	12	8.3
142	DNT5-/rtTA-	CT	12	11.4
143	DNT5+/rtTA+	DN	12	5.7

ID	GENOTYPE	GROUP	CAGE	Liver Acaca mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	2.58
113	DNT5+/rtTA-	CT	4	4.90
118	DNT5+/rtTA+	DN	5	3.06
119	DNT5-/rtTA+	CT	5	8.22
120	DNT5+/rtTA+	DN	5	6.31
121	DNT5+/rtTA+	DN	6	4.34
122	DNT5-/rtTA-	CT	6	6.48
123	DNT5+/rtTA+	DN	6	1.93
124	DNT5+/rtTA+	DN	7	2.50
125	DNT5+/rtTA-	CT	7	7.67
126	DNT5+/rtTA-	CT	7	8.02
127	DNT5+/rtTA+	DN	7	8.36
128	DNT5+/rtTA+	DN	8	10.18
129	DNT5+/rtTA+	DN	8	6.37
130	DNT5-/rtTA+	CT	8	12.84
134	DNT5+/rtTA-	CT	10	6.24
135	DNT5+/rtTA+	DN	10	8.15
136	DNT5+/rtTA+	DN	10	
137	DNT5+/rtTA+	DN	11	7.02
138	DNT5-/rtTA-	CT	11	8.71
139	DNT5-/rtTA-	CT	11	6.39
140	DNT5+/rtTA+	DN	12	3.49
141	DNT5-/rtTA+	CT	12	1.18
142	DNT5-/rtTA-	CT	12	5.30
143	DNT5+/rtTA+	DN	12	2.07

ID	GENOTYPE	GROUP	CAGE	Liver Fasn mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	0.56
113	DNT5+/rtTA-	CT	4	2.17
118	DNT5+/rtTA+	DN	5	0.59
119	DNT5-/rtTA+	CT	5	3.19
120	DNT5+/rtTA+	DN	5	1.20
121	DNT5+/rtTA+	DN	6	1.67
122	DNT5-/rtTA-	CT	6	2.31
123	DNT5+/rtTA+	DN	6	0.31
124	DNT5+/rtTA+	DN	7	0.50
125	DNT5+/rtTA-	CT	7	2.35
126	DNT5+/rtTA-	CT	7	1.80
127	DNT5+/rtTA+	DN	7	2.14
128	DNT5+/rtTA+	DN	8	2.84
129	DNT5+/rtTA+	DN	8	1.16
130	DNT5-/rtTA+	CT	8	3.13
134	DNT5+/rtTA-	CT	10	1.66
135	DNT5+/rtTA+	DN	10	1.42
136	DNT5+/rtTA+	DN	10	
137	DNT5+/rtTA+	DN	11	3.63
138	DNT5-/rtTA-	CT	11	2.61
139	DNT5-/rtTA-	CT	11	2.08
140	DNT5+/rtTA+	DN	12	1.32
141	DNT5-/rtTA+	CT	12	0.45
142	DNT5-/rtTA-	CT	12	1.02
143	DNT5+/rtTA+	DN	12	0.61

ID	GENOTYPE	GROUP	CAGE	Liver Scd1 mRNA relative abundance
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	1.80
113	DNT5+/rtTA-	CT	4	3.98
118	DNT5+/rtTA+	DN	5	2.29
119	DNT5-/rtTA+	CT	5	5.68
120	DNT5+/rtTA+	DN	5	2.26
121	DNT5+/rtTA+	DN	6	3.42
122	DNT5-/rtTA-	CT	6	4.61
123	DNT5+/rtTA+	DN	6	0.27
124	DNT5+/rtTA+	DN	7	1.06
125	DNT5+/rtTA-	CT	7	8.67
126	DNT5+/rtTA-	CT	7	4.50
127	DNT5+/rtTA+	DN	7	7.89
128	DNT5+/rtTA+	DN	8	4.48
129	DNT5+/rtTA+	DN	8	2.46
130	DNT5-/rtTA+	CT	8	6.40
134	DNT5+/rtTA-	CT	10	3.65
135	DNT5+/rtTA+	DN	10	3.33
136	DNT5+/rtTA+	DN	10	
137	DNT5+/rtTA+	DN	11	4.30
138	DNT5-/rtTA-	CT	11	3.65
139	DNT5-/rtTA-	CT	11	2.51
140	DNT5+/rtTA+	DN	12	2.92
141	DNT5-/rtTA+	CT	12	1.08
142	DNT5-/rtTA-	CT	12	2.47
143	DNT5+/rtTA+	DN	12	1.69

ID	GENOTYPE	GROUP	CAGE	Cholesterol plasma concentration (mmol/L)
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	24.1
113	DNT5+/rtTA-	CT	4	
118	DNT5+/rtTA+	DN	5	26.4
119	DNT5-/rtTA+	CT	5	20.5
120	DNT5+/rtTA+	DN	5	31.3
121	DNT5+/rtTA+	DN	6	
122	DNT5-/rtTA-	CT	6	20.2
123	DNT5+/rtTA+	DN	6	18.5
124	DNT5+/rtTA+	DN	7	21.6
125	DNT5+/rtTA-	CT	7	24.5
126	DNT5+/rtTA-	CT	7	21.9
127	DNT5+/rtTA+	DN	7	20.3
128	DNT5+/rtTA+	DN	8	22.8
129	DNT5+/rtTA+	DN	8	23.2
130	DNT5-/rtTA+	CT	8	24.1
134	DNT5+/rtTA-	CT	10	24.6
135	DNT5+/rtTA+	DN	10	23.1
136	DNT5+/rtTA+	DN	10	8.2
137	DNT5+/rtTA+	DN	11	16.6
138	DNT5-/rtTA-	CT	11	21.5
139	DNT5-/rtTA-	CT	11	
140	DNT5+/rtTA+	DN	12	19.1
141	DNT5-/rtTA+	CT	12	22.8
142	DNT5-/rtTA-	CT	12	
143	DNT5+/rtTA+	DN	12	18.6

ID	GENOTYPE	GROUP	CAGE	HDL plasma concentration (mmol/L)
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	0.4
113	DNT5+/rtTA-	CT	4	
118	DNT5+/rtTA+	DN	5	0.4
119	DNT5-/rtTA+	CT	5	0.3
120	DNT5+/rtTA+	DN	5	0.4
121	DNT5+/rtTA+	DN	6	
122	DNT5-/rtTA-	CT	6	0.5
123	DNT5+/rtTA+	DN	6	0.4
124	DNT5+/rtTA+	DN	7	0.3
125	DNT5+/rtTA-	CT	7	0.3
126	DNT5+/rtTA-	CT	7	0.3
127	DNT5+/rtTA+	DN	7	0.3
128	DNT5+/rtTA+	DN	8	0.3
129	DNT5+/rtTA+	DN	8	0.3
130	DNT5-/rtTA+	CT	8	0.3
134	DNT5+/rtTA-	CT	10	0.5
135	DNT5+/rtTA+	DN	10	0.3
136	DNT5+/rtTA+	DN	10	
137	DNT5+/rtTA+	DN	11	0.3
138	DNT5-/rtTA-	CT	11	0.3
139	DNT5-/rtTA-	CT	11	
140	DNT5+/rtTA+	DN	12	0.3
141	DNT5-/rtTA+	CT	12	0.3
142	DNT5-/rtTA-	CT	12	
143	DNT5+/rtTA+	DN	12	0.5

ID	GENOTYPE	GROUP	CAGE	LDL plasma concentration (mmol/L)
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	22.8
113	DNT5+/rtTA-	CT	4	
118	DNT5+/rtTA+	DN	5	25.1
119	DNT5-/rtTA+	CT	5	19.1
120	DNT5+/rtTA+	DN	5	29.7
121	DNT5+/rtTA+	DN	6	
122	DNT5-/rtTA-	CT	6	18.8
123	DNT5+/rtTA+	DN	6	17.6
124	DNT5+/rtTA+	DN	7	20.3
125	DNT5+/rtTA-	CT	7	23.2
126	DNT5+/rtTA-	CT	7	20.6
127	DNT5+/rtTA+	DN	7	19.3
128	DNT5+/rtTA+	DN	8	22
129	DNT5+/rtTA+	DN	8	21.9
130	DNT5-/rtTA+	CT	8	22.8
134	DNT5+/rtTA-	CT	10	23.2
135	DNT5+/rtTA+	DN	10	21.9
136	DNT5+/rtTA+	DN	10	
137	DNT5+/rtTA+	DN	11	15.7
138	DNT5-/rtTA-	CT	11	20.5
139	DNT5-/rtTA-	CT	11	
140	DNT5+/rtTA+	DN	12	18.2
141	DNT5-/rtTA+	CT	12	21.5
142	DNT5-/rtTA-	CT	12	
143	DNT5+/rtTA+	DN	12	17.3

ID	GENOTYPE	GROUP	CAGE	Triglycerides plasma concentration (mmol/L)
49	DNT5+/rtTA-	CT	1	
50	DNT5+/rtTA+	DN	1	
51	DNT5+/rtTA-	CT	2	
52	DNT5+/rtTA+	DN	2	
112	DNT5+/rtTA+	DN	4	2
113	DNT5+/rtTA-	CT	4	2.4
118	DNT5+/rtTA+	DN	5	2.1
119	DNT5-/rtTA+	CT	5	2.6
120	DNT5+/rtTA+	DN	5	2.7
121	DNT5+/rtTA+	DN	6	3.2
122	DNT5-/rtTA-	CT	6	2
123	DNT5+/rtTA+	DN	6	1.1
124	DNT5+/rtTA+	DN	7	2.2
125	DNT5+/rtTA-	CT	7	2.2
126	DNT5+/rtTA-	CT	7	2.2
127	DNT5+/rtTA+	DN	7	1.7
128	DNT5+/rtTA+	DN	8	1.1
129	DNT5+/rtTA+	DN	8	2.4
130	DNT5-/rtTA+	CT	8	2.2
134	DNT5+/rtTA-	CT	10	2.1
135	DNT5+/rtTA+	DN	10	2.1
136	DNT5+/rtTA+	DN	10	0.6
137	DNT5+/rtTA+	DN	11	1.3
138	DNT5-/rtTA-	CT	11	1.7
139	DNT5-/rtTA-	CT	11	2.5
140	DNT5+/rtTA+	DN	12	1.3
141	DNT5-/rtTA+	CT	12	2.4
142	DNT5-/rtTA-	CT	12	1.9
143	DNT5+/rtTA+	DN	12	1.8

Supplementary File 4

Adipose tissue Adiponectin mRNA

- Log transformed outcome
- Separate variance for each cage

```
lme(log(MEASUREMENTS)~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE,weights=varIdent(form=1|CAGE))
```

Linear mixed-effects model fit by REML

AIC	BIC	logLik
54.23618	66.23765	-16.11809

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 0.05721544 0.327978

Variance function:

Structure: Different standard deviations per stratum

Formula: ~1 | CAGE

Parameter estimates:

	4	5	6	7	8	10	11	12
	1.0000000	0.3108447	0.1478915	1.9500141	2.2698978	2.9893407	2.1503730	2.5430880

Fixed effects: log(MEASUREMENTS) ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	0.9914514	0.05880142	15	16.861011	0
as.factor(GROUP)DN	0.4837610	0.05271480	15	9.176948	0

Correlation:

	(Intr)
as.factor(GROUP)DN	-0.585

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.4517986	-0.5715181	0.4326466	0.9476340	1.7014354

Number of Observations: 24

Number of Groups: 8

Adipose tissue Leptin mRNA

- No random intercept

```
lm(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude)
```

Call:

```
lm(formula = MEASUREMENTS ~ as.factor(GROUP), data = baptiste,  
    na.action = na.exclude)
```

Residuals:

```
    Min      1Q  Median      3Q     Max  
-6.300 -2.620 -0.560  1.635  5.800
```

Coefficients:

```
                Estimate Std. Error t value Pr(>|t|)  
(Intercept)         12.820      1.081  11.862 4.96e-11 ***  
as.factor(GROUP)DN   -0.220      1.415   -0.155  0.878  
---  
Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
```

Residual standard error: 3.418 on 22 degrees of freedom

(11 observations deleted due to missingness)

Multiple R-squared: 0.001098, Adjusted R-squared: -0.04431

F-statistic: 0.02417 on 1 and 22 DF, p-value: 0.8779

Adipose Tissue Tnf α mRNA

- Log transformed outcome

```
lme(log(MEASUREMENTS)~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)
```

Linear mixed-effects model fit by REML

```
      AIC      BIC    logLik
53.68639 58.39861 -22.8432
```

Random effects:

```
Formula: ~1 | CAGE
```

```
(Intercept) Residual
```

```
StdDev:    0.247604 0.5189493
```

Fixed effects: log(MEASUREMENTS) ~ as.factor(GROUP)

```
              Value Std.Error DF   t-value p-value
(Intercept)    2.7541954 0.1725478 16 15.961926 0.0000
as.factor(GROUP)DN -0.5628675 0.2085491 16 -2.698968 0.0158
```

Correlation:

```
(Intr)
```

```
as.factor(GROUP)DN -0.644
```

Standardized Within-Group Residuals:

```
      Min      Q1      Med      Q3      Max
-1.6725677 -0.4910287 -0.1186797 0.5136341 2.1121970
```

Number of Observations: 26

Number of Groups: 9

Adipose Tissue Il6 mRNA

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
153.7599	158.3019	-72.87997

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 3.762222 4.232612

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	12.248834	1.806212	15	6.781502	0.0000
as.factor(GROUP)DN	-4.481659	1.783115	15	-2.513387	0.0239

Correlation:

(Intr)

as.factor(GROUP)DN -0.539

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.50567381	-0.49408593	-0.03143692	0.39807985	1.79764299

Number of Observations: 25

Number of Groups: 9

Adipose Tissue Il1 β mRNA

- Log transformed outcome
- Separate variance for each cage

```
lme(log(MEASUREMENTS)~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE,weights=varIdent(form
m=~1|CAGE))
```

Linear mixed-effects model fit by REML

```
      AIC      BIC    logLik
51.50191 65.12784 -13.75095
```

Random effects:

```
Formula: ~1 | CAGE
      (Intercept) Residual
StdDev:  0.2203154 0.2179007
```

Variance function:

```
Structure: Different standard deviations per stratum
Formula: ~1 | CAGE
Parameter estimates:
      4      5      6      7      8      9      10      11
12
1.00000000 2.83543986 1.59626891 1.93602280 3.03493341 1.01670964 2.35663960 4.22466494
0.03467693
```

Fixed effects: log(MEASUREMENTS) ~ as.factor(GROUP)

```
      Value Std.Error DF  t-value p-value
(Intercept)      0.2613991 0.10525641 15   2.48345  0.0253
as.factor(GROUP)DN -0.3480921 0.00924146 15  -37.66634  0.0000
```

Correlation:

```
      (Intr)
as.factor(GROUP)DN -0.039
```

Standardized Within-Group Residuals:

```
      Min      Q1      Med      Q3      Max
-1.33165049 -0.56574422 -0.03472342  0.80643271  1.55250694
```

Number of Observations: 25

Number of Groups: 9

Adipose Tissue Ccl2 mRNA

- Log transformed outcome

```
lme(log(MEASUREMENTS)~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)
```

Linear mixed-effects model fit by REML

```
      AIC      BIC    logLik
71.2721 75.81408 -31.63605
```

Random effects:

```
Formula: ~1 | CAGE
```

```
(Intercept) Residual
```

```
StdDev:    0.865878 0.6270326
```

Fixed effects: log(MEASUREMENTS) ~ as.factor(GROUP)

```
              Value Std.Error DF   t-value p-value
(Intercept)    1.5810537 0.3433604 15   4.604647  0.0003
as.factor(GROUP)DN -0.8187237 0.2673886 15  -3.061924  0.0079
```

Correlation:

```
(Intr)
```

```
as.factor(GROUP)DN -0.392
```

Standardized Within-Group Residuals:

```
      Min      Q1      Med      Q3      Max
-1.62607729 -0.35049144  0.02538033  0.35756435  2.22361879
```

Number of Observations: 25

Number of Groups: 9

Adipose tissue Ccl3 mRNA

- Log transformed outcome

```
lme(log(MEASUREMENTS)~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)
```

Linear mixed-effects model fit by REML

AIC	BIC	logLik
61.5593	66.10128	-26.77965

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 0.6421199 0.5256003

Fixed effects: log(MEASUREMENTS) ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	0.9059330	0.2646800	15	3.422749	0.0038
as.factor(GROUP)DN	-0.3028831	0.2234118	15	-1.355716	0.1953

Correlation:

(Intr)

as.factor(GROUP)DN -0.426

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.522076562	-0.536108186	-0.008631966	0.593118802	1.808921007

Number of Observations: 25

Number of Groups: 9

Adipose tissue Ccl5 mRNA

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
89.37191	93.91388	-40.68595

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 0.4870267 1.192009

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	2.7151350	0.3834892	15	7.080082	0.0000
as.factor(GROUP)DN	0.1428942	0.4873623	15	0.293199	0.7734

Correlation:

(Intr)

as.factor(GROUP)DN -0.654

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.20815645	-0.71751485	-0.06598957	0.43911221	2.87451011

Number of Observations: 25

Number of Groups: 9

Adipose tissue Ccl7 mRNA

- Log transformed outcome

```
lme(log(MEASUREMENTS)~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)
```

Linear mixed-effects model fit by REML

```
      AIC      BIC    logLik
89.56963 94.1116 -40.78481
```

Random effects:

```
Formula: ~1 | CAGE
```

```
(Intercept) Residual
```

```
StdDev:    0.5189745 1.188207
```

Fixed effects: log(MEASUREMENTS) ~ as.factor(GROUP)

```
              Value Std.Error DF   t-value p-value
(Intercept)    0.4811997 0.3874108 15   1.242091  0.2333
as.factor(GROUP)DN -0.7951036 0.4868322 15  -1.633219  0.1232
```

Correlation:

```
(Intr)
```

```
as.factor(GROUP)DN -0.646
```

Standardized Within-Group Residuals:

```
      Min      Q1      Med      Q3      Max
-2.45429421 -0.68462015  0.04534429  0.52957325  1.76155561
```

Number of Observations: 25

Number of Groups: 9

Atherosclerosis - Whole aorta

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
137.3486	143.3346	-64.6743

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 0.8535712 1.391928

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	3.429986	0.4018913	21	8.534611	0.0000
as.factor(GROUP)DN	-0.793874	0.4975658	21	-1.595516	0.1255

Correlation:

(Intr)

as.factor(GROUP)DN -0.546

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.6088751	-0.5122920	0.0443579	0.4559438	1.7203455

Number of Observations: 35

Number of Groups: 13

Atherosclerosis - Aortic arch

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
176.4626	182.4486	-84.23128

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 1.84959 2.407985

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	6.882241	0.7625181	21	9.025675	0.0000
as.factor(GROUP)DN	-0.626173	0.8723517	21	-0.717799	0.4808

Correlation:

(Intr)

as.factor(GROUP)DN -0.5

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.72605497	-0.39237228	-0.04790167	0.56550442	1.93328981

Number of Observations: 35

Number of Groups: 13

Atherosclerosis - Aortic Conduit

- Separate variance for each cage

```
lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE,weights=varIdent(form=~1|CAGE))
```

Linear mixed-effects model fit by REML

```
      AIC      BIC    logLik
103.0166 126.9607 -35.50829
```

Random effects:

```
Formula: ~1 | CAGE
```

```
(Intercept) Residual
```

```
StdDev:    0.6754179 0.1922569
```

Variance function:

```
Structure: Different standard deviations per stratum
```

```
Formula: ~1 | CAGE
```

```
Parameter estimates:
```

```
      1      2      4      5      6      7      8      9     10
11      12     13     14
1.0000000 4.8726024 1.7466658 2.4423519 1.0661179 2.2941552 5.9888016 0.7913750 7.2411049
0.1830296 7.7441739 2.3529391 0.8158550
```

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

```
              Value Std.Error DF   t-value p-value
(Intercept)    1.1660331 0.211175552 21   5.506506  0.0000
as.factor(GROUP)DN -0.0936825 0.04138925 21  -2.263451  0.0343
```

Correlation:

```
(Intr)
as.factor(GROUP)DN -0.077
```

Standardized Within-Group Residuals:

```
      Min      Q1      Med      Q3      Max
-1.40383817 -0.76777524 -0.06686094  0.79237245  1.54262569
```

Number of Observations: 35

Number of Groups: 13

Weight Gain Week 6

- Separate variance for each cage

```
lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE,weights=varIdent(form=~1|CAGE))
```

Linear mixed-effects model fit by REML

```
      AIC      BIC    logLik
226.9532 243.7991 -100.4766
```

Random effects:

```
Formula: ~1 | CAGE
```

```
(Intercept) Residual
```

```
StdDev:    10.42275 6.598615
```

Variance function:

```
Structure: Different standard deviations per stratum
```

```
Formula: ~1 | CAGE
```

```
Parameter estimates:
```

```
      5      6      7      8      9      10      11      12      13
14
1.0000000 0.1034838 1.5789848 1.7823585 2.9816233 1.2284970 1.2961055 0.6203762 0.9423738
0.4006431
```

```
Fixed effects: MEASUREMENTS ~ as.factor(GROUP)
```

```
              Value Std.Error DF   t-value p-value
(Intercept)    40.30330   3.661662 18  11.00683     0
as.factor(GROUP)DN -12.82013   0.804876 18 -15.92808     0
```

```
Correlation:
```

```
              (Intr)
as.factor(GROUP)DN -0.092
```

Standardized Within-Group Residuals:

```
      Min      Q1      Med      Q3      Max
-1.281830598 -0.695807847 -0.006646045  0.768642613  1.619162874
```

Number of Observations: 29

Number of Groups: 10

Weight Gain Week 12

- Separate variance for each cage

```
lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE,weights=varIdent(form=~1|CAGE))
```

Linear mixed-effects model fit by REML

```
      AIC      BIC    logLik
262.3421 279.188 -118.1711
```

Random effects:

```
Formula: ~1 | CAGE
```

```
(Intercept) Residual
```

```
StdDev:    19.09569 1.840794
```

Variance function:

```
Structure: Different standard deviations per stratum
```

```
Formula: ~1 | CAGE
```

```
Parameter estimates:
```

```
      5      6      7      8      9      10      11      12      13
14
1.000000 4.844124 11.841168 13.314933 11.952929 7.701824 8.327383 3.637733 15.390009
5.920469
```

```
Fixed effects: MEASUREMENTS ~ as.factor(GROUP)
```

```
              Value Std.Error DF   t-value p-value
(Intercept)    80.74312  6.849613 18  11.78798     0
as.factor(GROUP)DN -25.93712  2.048303 18 -12.66274     0
```

```
Correlation:
```

```
              (Intr)
as.factor(GROUP)DN -0.129
```

```
Standardized Within-Group Residuals:
```

```
      Min      Q1      Med      Q3      Max
-1.4987695 -0.7199205 0.1069572 0.6272421 1.6227194
```

```
Number of Observations: 29
```

```
Number of Groups: 10
```

Adipocyte size

- Separate variance for each cage

```
lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE,weights=varIdent(form=~1|CAGE))
```

Linear mixed-effects model fit by REML

```
      AIC      BIC  logLik
417.096 431.7225 -196.548
```

Random effects:

```
Formula: ~1 | CAGE
```

```
(Intercept) Residual
```

```
StdDev:    205.8004 553.1374
```

Variance function:

```
Structure: Different standard deviations per stratum
```

```
Formula: ~1 | CAGE
```

```
Parameter estimates:
```

```
      4      5      6      7      8      9      10      11
12
1.00000000 0.07704862 0.33401133 1.61261520 2.78493998 0.61984231 1.71281120 0.27927668
2.60268312
```

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

```
              Value Std.Error DF   t-value p-value
(Intercept)    4526.546 109.96148 17  41.16483    0
as.factor(GROUP)DN -508.252  48.61002 17 -10.45571    0
```

Correlation:

```
(Intr)
as.factor(GROUP)DN -0.218
```

Standardized Within-Group Residuals:

```
      Min      Q1      Med      Q3      Max
-1.96562200 -0.58563257 0.07957057 0.64274551 1.64166232
```

Number of Observations: 27

Number of Groups: 9

Liver Steatosis

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
170.5876	175.1296	-81.29379

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 3.769916 6.706968

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	27.225011	2.444700	16	11.13634	0.0000
as.factor(GROUP)DN	-4.896344	2.738187	16	-1.78817	0.0927

Correlation:

(Intr)

as.factor(GROUP)DN -0.629

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-2.5428596	-0.3756805	0.2148794	0.5911253	1.2486900

Number of Observations: 25

Number of Groups: 8

Liver Srebplc mRNA

- Separate variance for each cage

```
lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE,weights=varIdent(form=~1|CAGE))
```

Linear mixed-effects model fit by REML

```
      AIC      BIC    logLik
128.8651 140.8665 -53.43253
```

Random effects:

```
Formula: ~1 | CAGE
```

```
(Intercept) Residual
```

```
StdDev:    0.5253005 3.495393
```

Variance function:

```
Structure: Different standard deviations per stratum
```

```
Formula: ~1 | CAGE
```

```
Parameter estimates:
```

```
      4      5      6      7      8      10      11      12
1.0000000 0.2624571 0.5421273 0.1554060 1.1049000 0.4750143 2.5899755 1.1895210
```

```
Fixed effects: MEASUREMENTS ~ as.factor(GROUP)
```

```
              Value Std.Error DF   t-value p-value
(Intercept)      9.298797 0.4697197 15 19.796480  0.000
as.factor(GROUP)DN -1.874583 0.4590505 15 -4.083609  0.001
```

```
Correlation:
```

```
              (Intr)
as.factor(GROUP)DN -0.554
```

Standardized Within-Group Residuals:

```
      Min      Q1      Med      Q3      Max
-1.3887363 -0.4700683  0.2445098  0.8079815  1.8443333
```

Number of Observations: 24

Number of Groups: 8

Liver Acaca mRNA

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
136.9992	141.8747	-64.49961

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 2.417757 2.254051

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	7.195587	0.9827833	16	7.321641	0.0000
as.factor(GROUP)DN	-1.988653	0.9206586	16	-2.160033	0.0463

Correlation:

(Intr)

as.factor(GROUP)DN -0.423

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.5057243	-0.6889085	0.1725828	0.4741543	1.8815818

Number of Observations: 27

Number of Groups: 10

Liver Fasn mRNA

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
79.85878	84.73428	-35.92939

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 0.3574923 0.858634

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	2.0619022	0.2579059	16	7.994786	0.0000
as.factor(GROUP)DN	-0.6705682	0.3379362	16	-1.984304	0.0646

Correlation:

(Intr)

as.factor(GROUP)DN -0.618

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.4592421	-0.7099803	-0.2408227	0.3804016	2.2347082

Number of Observations: 27

Number of Groups: 10

Liver Scd1 mRNA

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
117.8604	122.7359	-54.93018

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 0.688461 1.857884

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	3.924979	0.5460840	16	7.187501	0.0000
as.factor(GROUP)DN	-1.067380	0.7286659	16	-1.464841	0.1623

Correlation:

(Intr)

as.factor(GROUP)DN -0.631

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-1.4126724	-0.4593962	-0.1350479	0.4058632	2.3005514

Number of Observations: 27

Number of Groups: 10

CHOLESTEROL

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

AIC	BIC	logLik
123.4102	127.188	-57.70511

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 0.6351555 4.421418

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	22.517433	1.583070	12	14.223904	0.000
as.factor(GROUP)DN	-1.466909	1.989032	12	-0.737499	0.475

Correlation:

(Intr)

as.factor(GROUP)DN -0.779

Standardized Within-Group Residuals:

Min	Q1	Med	Q3	Max
-2.86810542	-0.50231786	0.08201051	0.46205784	2.25844724

Number of Observations: 21

Number of Groups: 8

HDL

```
lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)
```

Linear mixed-effects model fit by REML

	AIC	BIC	logLik
	-28.40044	-24.83895	18.20022

Random effects:

Formula: ~1 | CAGE

	(Intercept)	Residual
--	-------------	----------

StdDev: 0.03013001 0.07249113

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	0.3535842	0.02813218	11	12.568674	0.0000
as.factor(GROUP)DN	-0.0010577	0.03332860	11	-0.031736	0.9753

Correlation:

(Intr)

as.factor(GROUP)DN -0.716

Standardized Within-Group Residuals:

	Min	Q1	Med	Q3	Max
	-0.8908851	-0.5403010	-0.4401045	0.5663041	1.9694419

Number of Observations: 20

Number of Groups: 8

LDL

lme(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude,random=~1|CAGE)

Linear mixed-effects model fit by REML

	AIC	BIC	logLik
	104.6889	108.2503	-48.34443

Random effects:

Formula: ~1 | CAGE

(Intercept) Residual

StdDev: 1.638636 2.781935

Fixed effects: MEASUREMENTS ~ as.factor(GROUP)

	Value	Std.Error	DF	t-value	p-value
(Intercept)	21.291422	1.163471	11	18.299918	0.0000
as.factor(GROUP)DN	-0.416924	1.284565	11	-0.324564	0.7516

Correlation:

(Intr)

as.factor(GROUP)DN -0.67

Standardized Within-Group Residuals:

	Min	Q1	Med	Q3	Max
	-1.4513596	-0.5360393	0.1487976	0.4505872	2.5088054

Number of Observations: 20

Number of Groups: 8

TRIGLYCERIDES

- No random intercept

```
lm(MEASUREMENTS~as.factor(GROUP),na.action=na.exclude)
```

```
Call:
```

```
lm(formula = MEASUREMENTS ~ as.factor(GROUP),  
    na.action = na.exclude)
```

```
Residuals:
```

```
      Min       1Q   Median       3Q      Max  
-1.2286 -0.3000  0.0000  0.2714  1.3714
```

```
Coefficients:
```

```
              Estimate Std. Error t value Pr(>|t|)  
(Intercept)         2.2000     0.1681  13.088 3.84e-12 ***  
as.factor(GROUP)DN  -0.3714     0.2246  -1.654   0.112  
---  
Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
```

```
Residual standard error: 0.5575 on 23 degrees of freedom
```

```
(10 observations deleted due to missingness)
```

```
Multiple R-squared:  0.1062,    Adjusted R-squared:  0.06739
```

```
F-statistic: 2.734 on 1 and 23 DF,  p-value: 0.1118
```