# Angiogenin and Osteoprotegerin are type II muscle specific myokines protecting pancreatic beta-cells against proinflammatory cytokines

Sabine Rutti<sup>1</sup>, Rodolphe Dusaulcy<sup>2</sup>, Jakob S. Hansen<sup>3,4</sup>, Cédric Howald<sup>5</sup>, Emmanouil T. Dermitzakis<sup>5</sup>, Bente K. Pedersen<sup>3</sup>, Michel Pinget<sup>1</sup>, Peter Plomgaard<sup>3,4,\*</sup> and Karim Bouzakri<sup>1, \$,\*</sup>

1: UMR DIATHEC, EA 7294, Centre Européen d'Etude du Diabète, Université de Strasbourg, Fédération de Médecine Translationnelle de Strasbourg (FMTS), Bld René Leriche, 67200 Strasbourg, France..

2: Molecular Diabetes Laboratory, Division of Endocrinology-Diabetes-Hypertension and Nutrition, University Hospital/University of Geneva Medical School, 1211 Geneva, Switzerland.

3: Department of Clinical Biochemistry, Rigshospitalet, Copenhagen, Denmark.

4: Centre of Physical Activity Research, Rigshospitalet, Copenhagen, Denmark.

5: Department of Genetic Medicine and Development, University of Geneva Medical School 1211 Geneva, Switzerland; Institute for Genetics and Genomics in Geneva (iGE3), University of Geneva, 1211 Geneva, Switzerland; Swiss Institute of Bioinformatics, Geneva, Switzerland

\* : Equally contribution

\$ Correspondence:

Karim Bouzakri UMR DIATHEC, EA 7294, Centre Européen d'Etude du Diabète, Université de Strasbourg, Fédération de Médecine Translationnelle de Strasbourg (FMTS), Bld René Leriche, 67200 Strasbourg, France. Email : <u>K.Bouzakri@Ceed-diabete.org</u>



**Figure S1**: Hierarchical clustering of the spearman correlation coefficients of RPKM values. Genes having a RPKM value >= 1 in at least 90% of the samples were taken into account. Myotubes are well segregated from biopsies but the type of tissue (soleus, triceps and vastus) doesn't cluster together.



**Figure S2**: principal component analysis of the RPKM values of all genes with a RPKM >= 1 in at least 90% of the samples. Only biopsies were kept. Principal component 2 (PC2 axis) separates the soleus (HM-S) form the triceps (HM-T) whereas the vastus (HM-V) is spread all over PC2 suggesting the heterogeneous nature of this tissue (mix of type I and type II fibers).

# Figure S3



**Figure S3:** principal component analysis of the RPKM values of all genes with an RPKM >= 1 in at least 90% of the samples. Only myotubes were kept.



Table S1: GO terms and KEGG pathways of the the differentialy expressed genes in Cells-S versus Cells-T/Biopsies-S versus Biopsies-T (A and D), Cells-T versus Cells -S (B and E) and Biopsies-S versus Biopsies-T (C and F).

		auj_p
organ_development	C=1538;O=167;E=112.45;R=1.49;rawP=1.54e-08	2.2
system_development	C=2158;O=220;E=157.79;R=1.39;rawP=8.08e-09	2.2
anatomical_structure_development	C=2485;O=244;E=181.70;R=1.34;rawP=2.67e-08	2.6
developmental_process	C=2782;O=266;E=203.41;R=1.31;rawP=5.38e-08	3.9
cell_migration	C=531;O=73;E=38.83;R=1.88;rawP=6.92e-08	4.0
localization_of_cell	C=562;O=75;E=41.09;R=1.83;rawP=1.56e-07	6.5
cell_motility	C=562;O=75;E=41.09;R=1.83;rawP=1.56e-07	6.5
anatomical_structure_morphogenesis	C=1331;O=143;E=97.32;R=1.47;rawP=4.63e-07	1.0
cellular_component_movement	C=754;O=92;E=55.13;R=1.67;rawP=3.84e-07	1.0
multicellular_organismal_development	C=2475;O=238;E=180.96;R=1.32;rawP=2.99e-07	1.0
locomotion	C=752;O=92;E=54.98;R=1.67;rawP=3.40e-07	1.0
tissue development	C=931;O=106;E=68.07;R=1.56;rawP=1.40e-06	3.0
regulation of developmental process	C=905:O=103:E=66.17:R=1.56:rawP=2.07e-06	5.0
multicellular organismal process	C=3152:O=286:F=230 47:R=1 24:rawP=2 17e-06	5.0
single-multicellular organism process	C=3142:O=284:E=229 73:B=1 24:rawP=3 51e-06	70
single-inditicential_organism_process	C=25142,O=204,L=225.75,R=1.24,Rawr=5.516-00	
coll_differentiation	C=1568:O=157:E=114 6E:D=1 27:rowD=8 4Eo 06	1 1 3
	C=1568;C=157;E=114.65;R=1.57;FawP=8.450-06	1.3
annune_system_process	C=1012;C=110;E=73.99;R=1.49;rawP=7.900-06	1.3
cellular_developmental_process	C=1686;U=167;E=123.28;R=1.35;FaWP=7.50e-06	1.3
regulation_of_blood_vessel_endothelial_cell_migration	C=31;O=10;E=2.27;R=4.41;rawP=4.41e-05	6.5
cell_fate_commitment	C=112;O=21;E=8.19;R=2.56;rawP=5.13e-05	7.2
enzyme_linked_receptor_protein_signaling_pathway	C=628;O=72;E=45.92;R=1.57;rawP=6.67e-05	8.6
single-organism_process	C=4462;O=374;E=326.25;R=1.15;rawP=6.95e-05	8.6
actin_cytoskeleton_organization	C=310;O=42;E=22.67;R=1.85;rawP=6.68e-05	8.6
biological_adhesion	C=533;O=62;E=38.97;R=1.59;rawP=0.0001	9.5
skeletal_system_morphogenesis	C=118;O=21;E=8.63;R=2.43;rawP=0.0001	9.5
blood_vessel_endothelial_cell_migration	C=48;O=12;E=3.51;R=3.42;rawP=0.0001	9.5
regulation_of_cellular_response_to_growth_factor_stimulus	C=34;O=10;E=2.49;R=4.02;rawP=0.0001	9.5
regionalization	C=174;O=27;E=12.72;R=2.12;rawP=0.0001	9.5
cell proliferation	C=990;O=103;E=72.39;R=1.42;rawP=9.87e-05	9.5
cell adhesion	C=532:O=62:E=38.90:R=1.59:rawP=0.0001	9.5
negative regulation of phosphorus metabolic process	C=188:O=28:E=13.75:R=2.04:rawP=0.0002	1.5
metanephros development	C=50:O=12:E=3.66:R=3.28:rawP=0.0002	1.5
cell surface receptor signaling pathway	C=1364:O=133:E=99.73:B=1.33:rawP=0.0002	1.9
organ morphogenesis	C=487:O=57:E=35 61:B=1 60:rawP=0 0002	1 1
connective tissue development	C-131:O-22:E-9 58:B-2 30:rawP-0.0002	1.0
enithelium_development	C=151,0=22,C=3:56,R=2:56,Raw1=0:0002	1.
pegative regulation of phosphate metabolic process	C-188:O-28:E-13 75:P-2 04:rowP-0.0002	1.
riegative_regulation_or_prospriate_metabolic_process	C=188,O=28,E=15.75,R=2.04,TawP=0.0002	1.5
vasculature_development	C=388;0=48;E=28.91;R=1.71;FawP=0.0002	1.:
peptide_cross-linking	C=14;O=6;E=1.02;R=5.86;FaWP=0.0003	2.0
growth_factor_binding	C=79;C=19;E=5.74;R=3.31;FawP=2.52e-06	1.4
actin_binding	C=269;0=40;E=19.55;R=2.05;rawP=9.92e-06	2.8
cytoskeletal_protein_binding	C=482;O=59;E=35.03;R=1.68;rawP=4.29e-05	8.1
cell_adhesion_molecule_binding	C=35;O=10;E=2.54;R=3.93;rawP=0.0001	1.:
calcium_ion_binding	C=350;O=45;E=25.44;R=1.77;rawP=0.0001	1.1
extracellular_matrix_binding	C=30;O=9;E=2.18;R=4.13;rawP=0.0002	1.8
glycosaminoglycan_binding	C=92;O=17;E=6.69;R=2.54;rawP=0.0003	2.4
extracellular_region_part	C=482;O=70;E=35.37;R=1.98;rawP=1.79e-08	6.7
cell_periphery	C=1999;O=205;E=146.68;R=1.40;rawP=4.59e-08	7.1
plasma_membrane	C=1946;O=200;E=142.79;R=1.40;rawP=6.14e-08	7.7
extracellular_region	C=815;O=100;E=59.80;R=1.67;rawP=1.14e-07	1.0
plasma_membrane_part	C=889;O=102;E=65.23;R=1.56;rawP=2.15e-06	2.0
extracellular_space	C=357;O=51;E=26.20;R=1.95;rawP=2.77e-06	2.0
extracellular_matrix	C=226;O=37;E=16.58;R=2.23;rawP=2.85e-06	2.0
stress fiber	C=43;O=13;E=3.16;R=4.12;rawP=7.46e-06	4.0
proteinaceous extracellular matrix	C=187;O=31;E=13.72;R=2.26;rawP=1.40e-05	5.0
actin filament bundle	C=45:O=13:E=3.30:R=3.94:rawP=1.30e-05	5.0
actomyosin	C=52:O=14:E=3.82:R=3.67:rawP=1.48e-05	5.0
integral to plasma membrane	C=496:O=62:F=36.39:B=1.70:rawP=2.00e-05	6.0
intrinsic to plasma membrane	C=519:O=64:E=38.08:B=1.68:rawP=2.26e-05	7.0
cortical cytoskeleton	C=37:O=11:E=2.71:B=4.05:rawP=4.49e-05	1 1 2
adherens junction	C=161:O=26:F=11.81:B=2.20:raw/P=0.0001	21
	C-101,0-20,C-11.81,N-2.20,Raw1-0.0001	
	C=3;0=3;L=0.00;R=7:37;IawF=0.0002	4.
	C=30,0=10,E=2.04,K=3.79,TawP=0.0002	4.4
cell_cortex	C=134;O=22;E=9.83;R=2.24;rawP=0.0003	5.
anchoring_junction	C=1/0;0=26;E=12.47;R=2.08;rawP=0.0003	
cytoplasmic_vesicle_part	C=248;O=34;E=18.20;R=1.87;rawP=0.0003	
cell_cortex_part	C=61;O=13;E=4.48;R=2.90;rawP=0.0004	(
Golgi_lumen	C=33;O=9;E=2.42;R=3.72;rawP=0.0005	(
membrane	C=4169;O=348;E=305.90;R=1.14;rawP=0.0006	(
cytoplasmic_membrane-bounded_vesicle_lumen	C=34;O=9;E=2.49;R=3.61;rawP=0.0006	(
vesicle_lumen	C=35;O=9;E=2.57;R=3.50;rawP=0.0007	(
neuronal cell body	C=156;O=23;E=11.45;R=2.01;rawP=0.0010	(
lytic vacuole	C=253:O=32:E=18.56:R=1.72:rawP=0.0017	- (
lysosome	C=253;O=32;E=18.56;R=1.72:rawP=0.0017	+
cell body	C=172:O=24:E=12.62:B=1.90:rawP=0.0017	
secretory granule lumen	C=33:O=8:E=2.42:B=3.30:rawP=0.0022	+
coll_projection	C=33,0=8,E=2.42,N=3.30,Nawr=0.0022	+
cen_projection	C=/1/, U=/3;E=52.01;R=1.39;F3WP=U.UU23	+
nlatalat alpha granula	C=41,0=9;E=3.01;K=2.99;faWP=0.0024	+
platelet_alpha_granule	··· · · · · · · · · · · · · · · · · ·	1 (
platelet_alpha_granule platelet_alpha_granule_lumen	C=33;0=8;E=2.42;K=3.30;FaWP=0.0022	
platelet_alpha_granule platelet_alpha_granule_lumen cell_surface	C=348;O=31;E=18.20;R=1.70;rawP=0.0022 C=248;O=31;E=18.20;R=1.70;rawP=0.0024	(
platelet_alpha_granule platelet_alpha_granule_lumen cell_surface cell-substrate_junction	C=33;0=8;E=2.42;K=3.30;raWP=0.0022 C=248;0=31;E=18.20;R=1.70;rawP=0.0024 C=119;0=18;E=8.73;R=2.06;rawP=0.0025	
platelet_alpha_granule platelet_alpha_granule_lumen cell_surface cell-substrate_junction lamellipodium	C=35;0=8;E=2:42;K=3:30;rawP=0.0022 C=248;0=31;E=18.20;R=1.70;rawP=0.0024 C=119;0=18;E=8:73;R=2.06;rawP=0.0025 C=101;0=16;E=7.41;R=2.16;rawP=0.0027	
platelet_alpha_granule platelet_alpha_granule_lumen cell_surface cell-substrate_junction lamellipodium vesicle_membrane	C=33;0=8;E=2:42;K=3:30;rawP=0.0022 C=248;0=31;E=18.20;R=1.70;rawP=0.0024 C=119;0=18;E=8:73;R=2:06;rawP=0.0025 C=101;0=16;E=7:41;R=2:16;rawP=0.0027 C=220;0=28;E=16:14;R=1:73;rawP=0.0029	
platelet_alpha_granule platelet_alpha_granule_lumen cell_surface cell-substrate_junction lamellipodium vesicle_membrane apical_part_of_cell	C=33;0=8;E=2.42;K=3.30;rawP=0.0022 C=248;O=31;E=18.20;R=1.70;rawP=0.0024 C=119;O=18;E=8.73;R=2.06;rawP=0.0025 C=101;O=16;E=7.41;R=2.16;rawP=0.0027 C=220;O=28;E=16.14;R=1.73;rawP=0.0029 C=141;O=20;E=10.35;R=1.93;rawP=0.0033	
platelet_alpha_granule platelet_alpha_granule_lumen cell_surface cell-substrate_junction lamellipodium vesicle_membrane apical_part_of_cell cell-substrate_adherens_junction	C=33;0=8;E=2:42;R=3:30;rawP=0.0022 C=248;0=31;E=18:20;R=1.70;rawP=0.0024 C=119;0=18;E=8:73;R=2:06;rawP=0.0025 C=101;0=16;E=7:41;R=2:16;rawP=0.0027 C=220;0=28;E=16:14;R=1.73;rawP=0.0029 C=141;0=20;E=10.35;R=1.93;rawP=0.0033 C=114;0=17;E=8:36;R=2:03;rawP=0.0038	

B Cells-S\_vs\_Cells-T, 2071 genes

CO torms	stats	adi mualua
doterms		auj_pvalue
multicellular_organismal_process	C=2814;O=616;E=485.36;R=1.27;rawP=2.19e-15	5.90E-12
single-multicellular_organism_process	C=2806;O=614;E=483.98;R=1.27;rawP=2.84e-15	5.90E-12
developmental_process	C=2516;O=557;E=433.96;R=1.28;rawP=1.67e-14	2.31E-11
multicellular_organismal_development	C=2232;O=500;E=384.97;R=1.30;rawP=1.18e-13	1.23E-10
anatomical_structure_morphogenesis	C=1189;O=297;E=205.08;R=1.45;rawP=1.78e-13	1.48E-10
single-organism_process	C=4040;O=820;E=696.82;R=1.18;rawP=7.17e-13	4.97E-10
system development	C=1935;O=439;E=333.75;R=1.32;rawP=1.22e-12	7.25E-10
anatomical structure development	C=2243:O=496:F=386.87:R=1.28:rawP=1.87e-12	9.72F-10
cellular developmental process	$C=1514\cdot O=352\cdot E=261 \ 13\cdot B=1 \ 35\cdot raw P=2 \ 27e-11$	1.05E-08
anotomical structure formation involved in morphogeneois	C=1314,0=332,E=201.13,N=1.35,Nawr=2.27e=11	1.05E-08
anatomical_structure_formation_involved_in_morphogenesis	C=928,O=253;E=100.06;R=1.47;TawP=2.08E-11	1.11E-08
	C=1400;O=326;E=241.47;R=1.35;raWP=1.39e-10	5.25E-08
organ_morphogenesis	C=436;O=126;E=75.20;R=1.68;rawP=4.16e-10	1.44E-07
nervous_system_development	C=954;O=235;E=164.55;R=1.43;rawP=4.88e-10	1.56E-07
cell_development	C=820;O=207;E=141.43;R=1.46;rawP=7.01e-10	2.08E-07
organ_development	C=1380;O=318;E=238.02;R=1.34;rawP=9.31e-10	2.58E-07
regulation of developmental process	C=810:O=203:E=139.71:B=1.45:rawP=2.04e-09	5.30E-07
response to wounding	C=570:0=152:E=98 31:B=1 55:rawP=3 48e-09	8 51F-07
cellular component morphogenesis	$C = 598 \cdot O = 157 \cdot E = 103 \cdot 14 \cdot B = 1 \cdot 52 \cdot r_{2} \cdot W = 6 \cdot 156 \cdot 09$	1.40E-06
	C=132;O=157;C=105:14;R=1:52;RawF=0:15e=05	1.400-00
extracellular_matrix_organization	C=123;O=48;E=21.21;R=2.26;FaWP=6.71e-09	1.40E-06
extracellular_structure_organization	C=123;O=48;E=21.21;R=2.26;rawP=6.71e-09	1.40E-06
cell_morphogenesis	C=567;O=150;E=97.80;R=1.53;rawP=8.01e-09	1.59E-06
cell_proliferation	C=913;O=221;E=157.47;R=1.40;rawP=9.25e-09	1.75E-06
tissue_development	C=833;O=204;E=143.68;R=1.42;rawP=1.43e-08	2.59E-06
regulation of cell differentiation	C=562;O=146;E=96.93;R=1.51;rawP=4.59e-08	7.95E-06
regulation of cell proliferation	C=688:O=172:E=118.67:R=1.45:rawP=5.05e-08	8,40E-06
nositive regulation of cell differentiation	$C=274 \cdot O=82 \cdot F=47 \cdot 26 \cdot R=1 \cdot 74 \cdot r_{2} \cdot WP=1 \cdot 04e_{-}07$	1.66F-05
positive regulation of developmental process	C-272:O-101/E-61 22:D-1 62:rourD-1 160 07	1 705 05
positive_regulation_of_developmental_process	C=373;O=104;E=64.33;R=1.62;FawP=1.16e-07	1.79E-05
locomotion	C=674;O=167;E=116.25;R=1.44;rawP=1.55e-07	2.30E-05
localization_of_cell	C=508;O=132;E=87.62;R=1.51;rawP=2.11e-07	2.74E-05
cell_motility	C=508;O=132;E=87.62;R=1.51;rawP=2.11e-07	2.74E-05
regulation of biological quality	C=1342;O=298;E=231.47;R=1.29;rawP=1.99e-07	2.74E-05
cellular component movement	C=676:O=167:E=116.60:B=1.43:rawP=1.91e-07	2.74E-05
regulation of multicellular organismal development	C=600:O=151:E=103:49:B=1:46:rawP=2:34e=07	2 95E-05
regulation_of_localization	C-722:O-179:E-126 42:B-1 41:rowB-2 560 07	2.552.05
	C=237,O=178,E=120.43,R=1.41,TawF=2.30e=07	2.97E-03
skeletal_system_development	C=227;O=70;E=39.15;R=1.79;rawP=2.57e-07	2.97E-05
positive_regulation_of_cell_proliferation	C=351;O=98;E=60.54;R=1.62;rawP=2.53e-07	2.97E-05
regulation_of_multicellular_organismal_process	C=914;O=214;E=157.65;R=1.36;rawP=2.81e-07	3.16E-05
cell_migration	C=478;O=125;E=82.45;R=1.52;rawP=3.12e-07	3.41E-05
cell_adhesion	C=484;O=126;E=83.48;R=1.51;rawP=3.64e-07	3.88E-05
biological adhesion	C=485:O=126:E=83.65:R=1.51:rawP=4.11e-07	4.27E-05
transmembrane signaling recentor activity	C=204·O=63·F=35 11·B=1 79·rawP=8 98e-07	3 00F-04
signal transducor activity	C=458:0=118:E=78 82:B=1 E0:row/D=1 200 06	3.00E 04
	C-458,0-118,E-78.83,R-1.30,TawF-1.39e-00	3.00E-04
molecular_transducer_activity	C=458;O=118;E=78.83;R=1.50;raWP=1.39e-06	3.00E-04
receptor_activity	C=343;O=95;E=59.03;R=1.61;rawP=5.26e-07	3.00E-04
signaling_receptor_activity	C=248;O=69;E=42.68;R=1.62;rawP=1.63e-05	2.60E-03
G-protein_coupled_receptor_activity	C=69;O=26;E=11.88;R=2.19;rawP=3.84e-05	5.20E-03
receptor binding	C=600;O=136;E=103.27;R=1.32;rawP=0.0002	2.31E-02
amino acid transmembrane transporter activity	C=35:O=15:E=6.02:R=2.49:rawP=0.0003	2.43E-02
protease binding	C=35:O=15:E=6 02:B=2 49:rawP=0 0003	2 43F-02
2' E' avalia nucleatida, phosphadiostorasa, activity	$C = 10 \cdot O = 7 \cdot E = 1 \cdot 72 \cdot P = 4 \cdot 07 \cdot r_2 \cdot v \cdot P = 0 \cdot 0003$	2.43E 02
linid hinding	C 10,0-7,L-1.72,R-4.07,RawF-0.0003	2.431-02
	C=408;O=95;E=70.22;R=1.35;rawP=0.0008	4.98E-02
neutral_amino_acid_transmembrane_transporter_activity	C=14;O=8;E=2.41;R=3.32;rawP=0.0008	4.98E-02
cyclic-nucleotide_phosphodiesterase_activity	C=11;O=7;E=1.89;R=3.70;rawP=0.0008	4.98E-02
extracellular_region	C=707;O=218;E=121.89;R=1.79;rawP=1.46e-20	6.88E-18
extracellular_region_part	C=416;O=128;E=71.72;R=1.78;rawP=2.93e-12	6.90E-10
proteinaceous_extracellular matrix	C=164;O=62;E=28.27;R=2.19;rawP=2.02e-10	3.17E-08
intrinsic to membrane	C=2210:O=477:E=381.01:R=1.25:rawP=7.28e-10	8.57E-08
cell periphery	C=1771·O=391·F=305 32·R=1 28·ram/P=2 9/a-09	2 77F_07
extracellular matrix	(-100.0-68.E-34.21.E-1.09.mu) = 4.120.00	3 245 07
	C 1722,O 200,E 200, D 1, 20 D 1, 20 D 2, 5 20 D 2	5.24E-07
piasma_membrane	C=1/22;O=380;E=296.88;R=1.28;rawP=5.80e-09	3.90E-07
integral_to_membrane	C=2176;O=464;E=375.15;R=1.24;rawP=8.90e-09	4.66E-07
extracellular_space	C=303;O=92;E=52.24;R=1.76;rawP=8.04e-09	4.66E-07
extracellular_matrix_part	C=92;O=38;E=15.86;R=2.40;rawP=4.25e-08	2.00E-06
plasma_membrane_part	C=759;O=185;E=130.85;R=1.41;rawP=1.19e-07	5.10E-06
membrane	C=3791;O=743;E=653.58:R=1.14:rawP=3.47e-07	1.36E-05
intrinsic to plasma membrane	C=434·O=115·F=74 82·R=1 54·rawP=4 81e-07	1 74F-05
membrane part	$C=2697 \cdot O=546 \cdot E=464.07 \cdot D=1.17 \cdot D=0.07$	2.7 46-05
internolate_part	C-415-C-108-E-71 EE-D-1 51	2.33E-05
Integral_to_plasma_memprane	C-413;U=108;E=/1.55;K=1.51;FaWP=2./2e-06	0.54E-05
collagen	C=37;O=16;E=6.38;R=2.51;rawP=0.0002	0.0059
membrane_raft	C=120;O=36;E=20.69;R=1.74;rawP=0.0004	0.0111
endoplasmic_reticulum_lumen	C=106;O=32;E=18.27;R=1.75;rawP=0.0007	0.0183
cell_projection	C=660;O=144;E=113.79;R=1.27;rawP=0.0010	0.0248
cell junction	C=379;O=88;E=65.34;R=1.35;rawP=0.0015	0.0336
basement membrane	C=51:O=18:E=8.79:R=2.05:rawP=0.0015	0.0336
axon	$C=137 \cdot O=37 \cdot F=23 \cdot 62 \cdot R=1 \cdot 57 \cdot r_{2100} - 0.0026$	0.0350
	C=45, 0=3, c=23, 02, n=1.3, iawr=0.0020	0.049
	C-43,0=10;C=7.70;K=2.00;FaWP=0.0024	0.049
GOIGI_STACK	C=87;O=26;E=15.00;R=1.73;rawP=0.0025	0.049
Imicrotibril	IC=7:0=5:F=1 21:R=4 14:rawP=0 0023	0.049

C Biopsies-S\_vs\_Biopsies-T, 3725 genes

GO terms	stats	adj_pvalue
cellular_respiration	C=136;O=103;E=48.48;R=2.12;rawP=7.28e-22	3.65E-18
respiratory_electron_transport_chain	C=96;O=78;E=34.22;R=2.28;rawP=3.60e-20	9.02E-17
energy_derivation_by_oxidation_of_organic_compounds	C=233;O=147;E=83.06;R=1.77;rawP=4.65e-18	7.76E-15
generation_of_precursor_metabolites_and_energy	C=311;O=182;E=110.87;R=1.64;rawP=4.06e-17	5.08E-14
oxidation-reduction_process	C=370;O=208;E=131.90;R=1.58;rawP=1.02e-16	9.60E-14
electron_transport_chain	C=118;O=86;E=42.07;R=2.04;rawP=1.15e-16	9.60E-14
mitochondrial_ATP_synthesis_coupled_electron_transport	C=47;O=37;E=16.76;R=2.21;rawP=1.74e-09	1.09E-06
ATP_synthesis_coupled_electron_transport	C=47;O=37;E=16.76;R=2.21;rawP=1.74e-09	1.09E-06
acetyl-CoA_metabolic_process	C=41;O=33;E=14.62;R=2.26;rawP=4.81e-09	2.44E-06
oxidative_phosphorylation	C=57;O=42;E=20.32;R=2.07;rawP=4.88e-09	2.44E-06
homophilic_cell_adhesion	C=37;O=30;E=13.19;R=2.27;rawP=1.80e-08	8.20E-06
mitochondrial_electron_transport,_NADH_to_ubiquinone	C=36;O=29;E=12.83;R=2.26;rawP=4.13e-08	1.72E-05
aerobic_respiration	C=38;O=30;E=13.55;R=2.21;rawP=5.62e-08	2.17E-05
small_molecule_metabolic_process	C=1435;O=598;E=511.57;R=1.17;rawP=7.74e-08	2.77E-05
cofactor_catabolic_process	C=32;O=26;E=11.41;R=2.28;rawP=1.55e-07	5.18E-05
tricarboxylic_acid_cycle	C=24;O=21;E=8.56;R=2.45;rawP=2.18e-07	6.82E-05
muscle_system_process	C=182;O=98;E=64.88;R=1.51;rawP=2.93e-07	8.63E-05
catabolic_process	C=1301;O=542;E=463.80;R=1.17;rawP=4.22e-07	1.00E-04
striated_muscle_contraction	C=62;O=41;E=22.10;R=1.85;rawP=9.32e-07	2.00E-04
organic_substance_catabolic_process	C=1207;O=504;E=430.29;R=1.17;rawP=8.82e-07	2.00E-04
acetyl-CoA_catabolic_process	C=25;O=21;E=8.91;R=2.36;rawP=9.03e-07	2.00E-04
actin-myosin_filament_sliding	C=34;O=26;E=12.12;R=2.15;rawP=1.35e-06	2.00E-04
translational_termination	C=88;O=54;E=31.37;R=1.72;rawP=7.02e-07	2.00E-04
translational_elongation	C=104;O=61;E=37.08;R=1.65;rawP=1.25e-06	2.00E-04
organic_acid_catabolic_process	C=121;O=69;E=43.14;R=1.60;rawP=1.10e-06	2.00E-04
muscle_filament_sliding	C=34;O=26;E=12.12;R=2.15;rawP=1.35e-06	2.00E-04
carboxylic_acid_catabolic_process	C=121;O=69;E=43.14;R=1.60;rawP=1.10e-06	2.00E-04
muscle_contraction	C=156;O=85;E=55.61;R=1.53;rawP=9.29e-07	2.00E-04
coenzyme_catabolic_process	C=28;O=23;E=9.98;R=2.30;rawP=5.79e-07	2.00E-04
protein_complex_subunit_organization	C=567;O=255;E=202.13;R=1.26;rawP=1.19e-06	2.00E-04
skeletal_muscle_contraction	C=19;O=17;E=6.77;R=2.51;rawP=1.77e-06	3.00E-04
protein_complex_disassembly	C=122;O=69;E=43.49;R=1.59;rawP=1.65e-06	3.00E-04
cellular_protein_complex_disassembly	C=119;O=67;E=42.42;R=1.58;rawP=2.90e-06	4.00E-04
acyl-CoA_metabolic_process	C=64;O=41;E=22.82;R=1.80;rawP=3.21e-06	5.00E-04
thioester_metabolic_process	C=64;O=41;E=22.82;R=1.80;rawP=3.21e-06	5.00E-04
translational_initiation	C=140;O=76;E=49.91;R=1.52;rawP=4.19e-06	6.00E-04
cellular_catabolic_process	C=1145;O=473;E=408.18;R=1.16;rawP=8.92e-06	1.20E-03
single-organism_catabolic_process	C=149;O=79;E=53.12;R=1.49;rawP=8.92e-06	1.20E-03
small_molecule_catabolic_process	C=148;O=78;E=52.76;R=1.48;rawP=1.35e-05	1.70E-03
protein_targeting_to_membrane	C=133;O=71;E=47.41;R=1.50;rawP=1.85e-05	2.30E-03
oxidoreductase_activity	C=391;O=204;E=139.29;R=1.46;rawP=4.56e-12	5.34E-09
structural_constituent_of_ribosome	C=147;O=86;E=52.37;R=1.64;rawP=9.76e-09	5.71E-06
NADH_dehydrogenase_(quinone)_activity	C=37;O=29;E=13.18;R=2.20;rawP=1.24e-07	2.90E-05
NADH_dehydrogenase_activity	C=37;O=29;E=13.18;R=2.20;rawP=1.24e-07	2.90E-05
NADH_dehydrogenase_(ubiquinone)_activity	C=37;O=29;E=13.18;R=2.20;rawP=1.24e-07	2.90E-05
cofactor_binding	C=155;O=83;E=55.22;R=1.50;rawP=3.08e-06	5.00E-04
oxidoreductase activity, acting on NADH or NADPH, quinc	C=46;O=32;E=16.39;R=1.95;rawP=2.75e-06	5.00E-04
inorganic cation transmembrane transporter activity	C=171;O=90;E=60.92;R=1.48;rawP=3.13e-06	5.00E-04
ion_transmembrane_transporter_activity	C=284;O=137;E=101.17;R=1.35;rawP=5.78e-06	8.00E-04
hydrogen_ion_transmembrane_transporter_activity	C=70;O=43;E=24.94;R=1.72;rawP=8.97e-06	1.00E-03
structural_molecule_activity	C=357;O=165;E=127.18;R=1.30;rawP=1.56e-05	1.70E-03
transmembrane transporter activity	C=339;O=156;E=120.77;R=1.29;rawP=3.55e-05	2.80E-03
acyl-CoA_dehydrogenase_activity	C=13;O=12;E=4.63;R=2.59;rawP=3.59e-05	2.80E-03
coenzyme_binding	C=114;O=62;E=40.61;R=1.53;rawP=2.87e-05	0.0028
substrate-specific transmembrane transporter activity	C=319;O=148;E=113.64;R=1.30;rawP=3.38e-05	0.0028
monovalent inorganic cation transmembrane transporter a	C=115;O=62;E=40.97;R=1.51;rawP=4.09e-05	0.003
cation transmembrane transporter activity	C=211:O=103:E=75.17:R=1.37:rawP=4.53e-05	0.0031
oxidoreductase activity, acting on the CH-CH group of do	C=41:O=27:E=14.61:R=1.85:rawP=7.62e-05	0.005
oxidoreductase activity, acting on NADH or NADPH	C=65:O=38:E=23.16:R=1.64:rawP=0.0001	0.0062
NAD binding	C=37:O=24:E=13.18:R=1.82:rawP=0.0003	0.0175
flavin adenine dinucleotide binding	C=42;O=26;E=14.96;R=1.74;rawP=0.0005	0.0279
mitochondrial part	C=586:O=322:E=207.58:R=1.55:rawP=9.02e-24	5.88E-21
mitochondrial inner membrane	C=278:O=178:E=98.47:R=1.81:rawP=5.23e-23	1.70E-20
organelle inner membrane	C=300:O=182:E=106.27:R=1.71:rawP=1.06e-19	1.73E-17
mitochondrial membrane	C=388:O=223:E=137.44:R=1.62:rawP=9.37e-20	1.73E-17
mitochondrial envelope	C=406:O=230:E=143.82:R=1.60:rawP=2.85e-19	3.72E-17
mitochondrion	C=1156;O=543;E=409.48:R=1.33:rawP=1.44e-18	1.56E-16
mitochondrial membrane part	C=121;O=84;E=42.86:R=1.96:rawP=1.88e-14	1.75F-12
mitochondrial matrix	C=237;O=136;E=83.95:R=1.62:rawP=2.20e-12	1.79F-10
respiratory chain	C=65;O=50;E=23.02;R=2.17;rawP=8.33e-12	6.03E-10
organelle envelope	C=600;O=290;E=212.53:R=1.36:rawP=9.80e-12	6.39E-10
mitochondrial respiratory chain	C=60;O=47;E=21.25:R=2.21:rawP=1.12e-11	6.64E-10
envelope	C=601:O=290:E=212.89:R=1.36:rawP=1.25e-11	6.79F-10
mitochondrial respiratory chain complex I	C=39;O=32;E=13.81:R=2.32:rawP=2.81e-09	1.22F-07
respiratory chain complex I	C=39;O=32;E=13.81;R=2.32;rawP=2.81e-09	1.22E-07
NADH dehydrogenase complex	C=39:O=32:E=13.81:R=2.32:rawP=2.81e-09	1.22F-07
ribosomal subunit	C=128:O=76:E=45.34:R=1.68:rawP=2.27e-08	9.25F-07
ribosome	C=189:O=103:E=66.95:R=1.54:rawP=4.64e-08	1.78F-06
cvtosolic ribosome	C=86:O=53:E=30.46:R=1.74:rawP=5.90e-07	2.14F-05
cytosolic part	C=158:O=84:E=55.97:R=1.50:rawP=3.09e-06	0.0001
contractile fiber part	C=120:O=67:E=42.51:R=1 58:rawP=3 39e-06	0.0001
cvtoplasmic part	C=4185:O=1573;E=1482.43:R=1.06:rawP=1.12e=05	0.0003
contractile fiber	C=133:O=71:F=47.11·R=1 51·rawP=1 470-05	0.0003
mvofibril	C=124:O=67:E=43.92:R=1.53:rawP=1.51e-05	0.0004
striated muscle thin filament	C=13:O=12:E=4.60:R=2.61:rawP=3.37e-05	0 0009
sarcomere	C=105:O=57:E=37.19:R=1.53:rawP=5 37e-05	0.0013
small ribosomal subunit	C=59:O=36:E=20.90:R=1.72:rawP=5.15e-05	0.0013
cvtosolic large ribosomal subunit	C=49:O=31:E=17.36:R=1.79:rawP=6.36e-05	0.0015
cytoplasm	C=5398:O=1988:F=1912 10:R=1 04:rawP=6 94o 05	0.0015
mitochondrial proton-transporting ATP synthese complex	C=19:O=15:F=6 73:R=2 23:rawP=0.0001	0.0010
large_ribosomal_subupit	C=70:0=40:F=24 80:P=1 61:rowD=0.0002	0.0022
sarconlasm	C=/0,0-40,L=24.00,R=1.01,I dWf=0.0002	0.0042
proton-transporting ATP synthese complex	C=20:0=15:E=7.08:P=2.12:rou/P=0.0004	0.0042
mombrane_part	C-20,0-13,C-7.00,N-2.12;FdWY=0.0004	0.0082
nenorane_part	C=45:O=27:E=15.04:B=1.60:rowB=0.0000	0.0115
troponin complex	C=7:0=7:E=2 48:E=2 82:re:::D=0 0007	0.0115
a doponin_complex	C-7, U-7, E-2.40, R-2.62; IdWP=U.UUU/	0.013
mombrano	C-35,0-22,E-12.40;K=1.77;TaWP=0.0008	0.0141
ion channel complex	C-45-0-27-E-16 20-B-1 65-march 0.0010	0.0141
T tubulo	C=40,0=27;C=10.23;K=1.00;FaWP=0.0010	0.0172
mitochondrial nucleoid	C-10,0-10,E-0.30,R=2.04;IdWF=0.0016	0.0254
milliochondrial nucleold	1C-32,O-20;E=11.34;K=1.76;FaWP=0.0016	U.0254

#### D Cells-S\_vs\_Cells-T/Biopsies-S\_vs\_Biopsies-T, 864 genes

KEGG pathway	stats	adj_pvalue
Regulation_of_actin_cytoskeleton	C=154;O=26;E=10.55;R=2.46;rawP=1.56e-05	2.20E-03
Focal_adhesion	C=158;O=24;E=10.82;R=2.22;rawP=0.0002	1.06E-02
Pathogenic_Escherichia_coli_infection	C=40;O=10;E=2.74;R=3.65;rawP=0.0003	1.06E-02
Leukocyte_transendothelial_migration	C=71;O=14;E=4.86;R=2.88;rawP=0.0003	1.06E-02

### E Cells-S\_vs\_Cells-T, 2071 genes

KEGG pathway	stats	adj_pvalue
Protein_digestion_and_absorption	C=32;O=17;E=5.37;R=3.17;rawP=2.71e-06	5.00E-04
Cytokine-cytokine_receptor_interaction	C=68;O=25;E=11.40;R=2.19;rawP=5.65e-05	5.50E-03
ECM-receptor_interaction	C=51;O=19;E=8.55;R=2.22;rawP=0.0004	0.0241
Focal_adhesion	C=143;O=40;E=23.98;R=1.67;rawP=0.0005	0.0241
Nitrogen_metabolism	C=9;O=6;E=1.51;R=3.98;rawP=0.0012	0.0386
Gap_junction	C=55;O=19;E=9.22;R=2.06;rawP=0.0010	0.0386
Amoebiasis	C=57;O=19;E=9.56;R=1.99;rawP=0.0017	0.041
Pathways_in_cancer	C=219;O=54;E=36.73;R=1.47;rawP=0.0017	0.041

## **F** Biopsies-S\_vs\_Biopsies-T, 3725 genes

KEGG pathway	stats	adj_pvalue
Oxidative_phosphorylation	C=104;O=76;E=36.63;R=2.08;rawP=2.89e-15	6.10E-13
Alzheimer's_disease	C=131;O=89;E=46.13;R=1.93;rawP=1.73e-14	1.83E-12
Parkinson's_disease	C=102;O=73;E=35.92;R=2.03;rawP=6.18e-14	4.35E-12
Huntington's_disease	C=139;O=84;E=48.95;R=1.72;rawP=9.38e-10	4.95E-08
Cardiac_muscle_contraction	C=48;O=36;E=16.90;R=2.13;rawP=2.04e-08	8.61E-07
Valine,_leucine_and_isoleucine_degradation	C=37;O=29;E=13.03;R=2.23;rawP=9.44e-08	2.85E-06
Ribosome	C=85;O=54;E=29.93;R=1.80;rawP=8.84e-08	2.85E-06
Metabolic_pathways	C=645;O=287;E=227.15;R=1.26;rawP=2.85e-07	7.52E-06
Citrate_cycle_(TCA_cycle)	C=26;O=21;E=9.16;R=2.29;rawP=2.50e-06	5.86E-05
Calcium_signaling_pathway	C=66;O=41;E=23.24;R=1.76;rawP=7.18e-06	0.0002
Propanoate_metabolism	C=25;O=19;E=8.80;R=2.16;rawP=3.71e-05	0.0007
Fatty_acid_metabolism	C=31;O=22;E=10.92;R=2.02;rawP=5.29e-05	0.0009
Glycolysis_/_Gluconeogenesis	C=37;O=25;E=13.03;R=1.92;rawP=6.03e-05	0.001
Pyruvate_metabolism	C=30;O=19;E=10.57;R=1.80;rawP=0.0015	0.0226
Tight_junction	C=80;O=41;E=28.17;R=1.46;rawP=0.0023	0.0324

**Table S2:** Cytokine assay of the CM of SKMC-S and SKMC-T cells in controlconditions of after 24h TNF-alpha treatment.

	C-SKMC-S-C	м	C-SKMC-T-CM		TNF-SKMC-S-CM		TNF-SKMC-T-CM	
Cytokine	Protein level (fold C-SKMC-S-CM)	ESM (±)						
Angiogenin	1.00	0.22	4.92	1.54	1.30	0.28	3.17	1.05
Acrp30	1.00	0.31	1.06	0.16	1.02	0.12	1.20	0.13
AgRP	1.00	0.25	1.04	0.21	1.01	0.21	1.15	0.13
ANGPT2	1.00	0.23	1.00	0.18	1.06	0.17	1.21	0.07
AREG	1.00	0.22	0.97	0.22	0.98	0.22	0.99	0.18
Axl	1.00	0.23	1.03	0.17	0.98	0.19	1.10	0.11
b-NGF	1.00	0.17	1.03	0.17	1.05	0.35	1.13	0.12
BDNF	1.00	0.18	0.84	0.07	0.96	0.07	0.99	0.06
bFGF	1.00	0.26	1.00	0.15	1.02	0.20	1.15	0.09
BLC	1.00	0.15	0.90	0.07	1.04	0.07	1.08	0.12
BMP-4	1.00	0.12	0.97	0.05	1.09	0.09	1.19	0.11
BMP-6	1.00	0.16	0.89	0.14	1.13	0.08	1.16	0.06
BTC	1.00	0.27	1.03	0.21	0.98	0.14	1.21	0.12
CCL11	1.00	0.17	1.04	0.27	1.19	0.29	2.11	0.70
CCL24	1.00	0.12	0.89	0.09	1.05	0.06	1.17	0.10
CCL26	1.00	0.13	0.98	0.10	1.14	0.08	1.21	0.10
CCL28	1.00	0.21	1.09	0.23	1.05	0.25	1.12	0.10
CKB 8-1	1.00	0.15	0.94	0.06	1.06	0.07	1.23	0.11
CNTF	1.00	0.14	0.92	0.08	1.11	0.08	1.17	0.10
CSF2	1.00	0.19	1.09	0.27	1.55	0.27	1.54	0.41
CSF3	1.00	0.28	1.00	0.22	1.06	0.30	1.18	0.23
СТАСК	1.00	0.22	0.95	0.22	0.92	0.21	1.21	0.13
CX3CL1	1.00	0.17	0.93	0.08	1.05	0.06	1.18	0.10
Dtk	1.00	0.21	0.95	0.21	0.93	0.14	1.16	0.06
EGF	1.00	0.16	0.92	0.05	1.06	0.06	1.18	0.09
EGFR	1.00	0.22	1.05	0.15	0.95	0.13	1.17	0.11
ENA-78	1.00	0.23	1.13	0.32	1.17	0.41	2.63	1.01
Fas	1.00	0.38	1.09	0.35	1.23	0.48	1.12	0.31
FGF4	1.00	0.25	0.97	0.19	0.95	0.25	1.12	0.12
FGF6	1.00	0.14	0.98	0.10	1.10	0.07	1.13	0.13
FGF7	1.00	0.15	0.89	0.08	1.00	0.08	1.13	0.09
FGF9	1.00	0.19	0.93	0.22	1.01	0.20	1.04	0.10
Flt-3L	1.00	0.15	0.96	0.10	1.06	0.06	1.20	0.12
GCP-2	1.00	0.16	0.99	0.05	1.10	0.08	1.12	0.11
GDNF	1.00	0.17	0.95	0.07	1.16	0.10	1.24	0.10
GITR	1.00	0.18	0.87	0.14	0.87	0.21	1.04	0.11
GITRL	1.00	0.21	0.99	0.18	0.93	0.08	1.06	0.09
GRO	1.00	0.20	1.71	0.79	9.12	5.16	19.09	10.13
GRO-a	1.00	0.24	1.06	0.25	1.10	0.35	1.22	0.23
HCC-4	1.00	0.16	0.96	0.17	0.97	0.20	1.12	0.12
HGF	1.00	0.24	1.26	0.28	0.97	0.13	1.11	0.16
1-309	1.00	0.15	0.94	0.07	1.04	0.06	1.13	0.05
I-TAC	1.00	0.24	0.94	0.23	1.35	0.46	1.13	0.17
ICAM1	1.00	0.26	1.04	0.18	0.98	0.19	1.17	0.09
ICAM3	1.00	0.20	0.94	0.13	0.86	0.13	1.01	0.08
IFN-g	1.00	0.20	0.94	0.31	1.14	0.27	1.21	0.33
IGF-1	1.00	0.15	0.91	0.09	1.04	0.10	1.21	0.08
IGFBP1	1.00	0.16	0.92	0.04	1.01	0.04	1.16	0.12
IGFBP2	1.00	0.19	0.91	0.07	1.12	0.16	1.13	0.08
IGFBP4	1.00	0.16	0.99	0.08	1.12	0.09	1.22	0.09
IGFBP6	1.00	0.22	2.00	0.60	1.23	0.40	1.77	0.56
IGFPB3	1.00	0.21	1.02	0.13	0.90	0.16	1.11	0.12
IL-1 R1	1.00	0.25	0.93	0.22	1.05	0.12	1.11	0.11
IL-10	1.00	0.14	0.93	0.11	1.11	0.08	1.21	0.10
IL-11	1.00	0.27	0.83	0.20	0.90	0.13	0.92	0.13
	1		1		1			

	C-SKMC-S-C	м	C-SKMC-T-CM		TNF-SKMC-S-CM		TNF-SKMC-T-CM	
Cytokine	Protein level (fold C-SKMC-S-CM)	ESM (±)	Protein level (fold C-SKMC-S-CM)	ESM (±)	Protein level (fold C-SKMC-S-CM)	ESM (±)	Protein level (fold C-SKMC-S-CM)	ESM (±)
IL-12 p40	1.00	0.17	1.05	0.16	0.96	0.18	1.01	0.13
IL-12 p70	1.00	0.21	1.00	0.14	0.91	0.18	1.04	0.15
IL-13	1.00	0.06	0.98	0.17	1.12	0.09	1.25	0.17
IL-15	1.00	0.14	0.86	0.10	1.05	0.08	1.18	0.11
IL-16	1.00	0.17	0.89	0.09	1.02	0.06	1.14	0.07
IL-17	1.00	0.22	1.04	0.13	1.04	0.20	1.11	0.12
IL-1a	1.00	0.12	0.97	0.16	1.22	0.08	1.21	0.11
IL-1b	1.00	0.12	0.84	0.09	1.12	0.07	1.09	0.12
IL-1ra	1.00	0.16	0.88	0.10	1.10	0.09	1.17	0.10
IL-2	1.00	0.13	0.84	0.09	1.12	0.07	1.14	0.14
IL-2 Ra	1.00	0.22	0.99	0.19	0.99	0.21	1.20	0.10
IL-3	1.00	0.16	0.99	0.11	1.23	0.04	1.30	0.11
IL-4	1.00	0.15	0.90	0.10	1.07	0.06	1.16	0.07
1L-5	1.00	0.18	0.83	0.11	1.00	0.09	1.10	0.07
IL-6	1.00	0.08	4.92	3.49	14.56	7.47	20.38	9.55
IL-7	1.00	0.16	1.03	0.13	1.03	0.08	1.26	0.10
IL-8	1.00	0.21	1.18	0.35	2.13	1.24	3.29	1.51
IL1R4	1.00	0.24	1.08	0.25	1.17	0.33	1.27	0.39
Leptin	1.00	0.15	0.85	0.05	1.00	0.18	1.25	0.19
LIGHT	1.00	0.15	1.02	0.20	1.08	0.08	1.16	0.09
M-CSF	1.00	0.13	0.93	0.06	1.06	0.07	1.10	0.07
MCP1	1.00	0.34	1.19	0.54	3.12	0.63	3.00	0.48
MCP2	1.00	0.15	1.03	0.24	1.66	0.39	2.35	0.80
МСР3	1.00	0.18	1.05	0.29	7.96	3.85	9.81	4.32
MCP4	1.00	0.16	0.96	0.08	1.05	0.06	1.12	0.09
MDC	1.00	0.19	0.88	0.07	1.00	0.06	1.12	0.09
MIF	1.00	0.30	1.76	0.61	0.96	0.33	1.23	0.38
MIG	1.00	0.15	0.85	0.12	1.31	0.03	1.17	0.10
MIP-1a	1.00	0.21	1.07	0.14	0.90	0.15	1.21	0.07
MIP-1b	1.00	0.23	1.02	0.14	0.82	0.15	1.09	0.11
MIP-1d	1.00	0.14	0.98	0.07	1.10	0.08	1.14	0.08
MIP-3a	1.00	0.17	1.11	0.27	1.31	0.26	2.49	0.81
MIP-3b	1.00	0.18	0.97	0.16	0.89	0.11	1.00	0.17
MSP-a	1.00	0.21	0.97	0.18	0.92	0.21	1.02	0.14
NAP-2	1.00	0.12	1.00	0.04	1.07	0.07	1.42	0.21
N1-3	1.00	0.17	0.93	0.06	1.03	0.07	1.12	0.08
NI-4	1.00	0.31	0.92	0.16	0.87	0.15	0.90	0.10
OPG	1.00	0.39	2.31	0.54	1.45	0.22	5.44	1.38
USM	1.00	0.16	0.81	0.08	0.81	0.15	0.78	0.10
PARC	1.00	0.15	0.95	0.06	1.06	0.08	1.16	0.07
PDGF-BB	1.00	0.14	0.86	0.09	1.01	0.06	1.12	0.09
PLGF	1.00	0.20	1.01	0.14	0.93	0.18	1.06	0.18
RANTES	1.00	0.18	1.03	0.28	24.27	3.43	27.75	2.13
	1.00	0.20	0.91	0.08	1.02	0.06	1.10	0.08
5DF-1	1.00	0.15	0.85	0.10	1.05	0.06	1.14	0.08
sgp130	1.00	0.22	0.99	0.10	0.94	0.20	1.08	0.11
siGF-1R	1.00	0.17	0.98	0.15	1.00	0.11	1.01	0.15
SIL-OR	1.00	0.22	0.94	0.20	1.00	0.10	1.12	0.09
STNERS	1.00	0.25	1.06	0.13	0.90	0.19	1.08	0.05
57/NFR2	1.00	0.24	1.00	0.12	1.02	0.19	1.07	0.15
TARC	1.00	0.15	0.96	0.06	1.02	0.08	1.17	0.12
TCE b1	1.00	0.21	1.01	0.24	1 10	0.10	1.15	0.10
	1.00	0.12	1.00	0.12	1.10	0.05	1.19	0.12
	1.00	0.15	1.00	0.04	1.07	0.08	1.20	0.08
	1.00	0.23	1 26	0.14	1 20	0.22	1.07	0.12
TIMDO	1.00	0.40	1 21	0.30	1 17	0.05	1 19	0.70
	1.00	0.30	1 17	0.30	10.25	0.41 1 01	10 10	4.01
TRAIL 02	1.00	0.12	1.00	0.25	0.35	+.21 Ο 12	1 09	4.01 0.11
TRAIL RA	1.00	0.24	1.00	0.20	0.07	0.12	1.03	0.11
INAIL N4	1.00	0.31	1.05	0.20	1.07	0.22	1 14	0.19
VECE A	1.00	0.25	0.82	0.22	1.07	0.10	0.05	0.17
VEGE D	1.00	0.24	0.02	0.17	0.00	0.15	0.95	0.07
VCGF-D	1.00	0.18	0.90	0.17	0.95	0.10	1 11	0.10
ACLI	1.00	0.10	0.95	0.17	0.00	0.13	1 1.11	0.11

 Table S3: Muscle biopsies donors informations.

	MEAN	SEM
Age	22.79	0.38
Height	182.79	1.71
Weight	75.96	1.89
BMI	22.71	0.39
Р	51.36	1.84
S-BP (mmHg)	135.71	1.87
D-BP (mmHg)	78.43	1.62
Hb	9.10	0.16
Hct	0.43	0.01
Leuco	5.67	0.36
Trombo	214.21	10.42
Crp	1.43	0.43
Na	142.50	0.57
к	3.91	0.05
TG	0.89	0.11
Total chol	4.15	0.21
HDL	1.44	0.06
LDL	2.34	0.18
Creatinin	82.64	2.68
Carbamid	5.24	0.43
ALAT	18.18	1.51
Bilirubin	13.36	1.91
Alcalic phos	61.43	4.52
F II+VII+X	0.66	0.02
HbA1c	5.11	0.05
TSH	2.13	0.22