



Supplementary Materials

G Protein-Coupled Receptor 15 Expression Is Associated with Myocardial Infarction

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Table S1. Study characteristics of the early-onset MI individuals and controls of the Gutenberg Young Infarction Study.

Characteristics	All	Controls	MI cases
n (%)	224	112 (50)	112 (50)
Females, n (%)	72 (32)	36 (32)	36 (32)
Age, years	45 (42-49)	45 (41-49)	45 (42-49)
Age at MI	43 (40-48)	NA	43 (40-48)
Cardiovascular risk factors			
Diabetes mellitus, n (%)	13 (6)	2 (2)	11 (10)
Body mass index, kg/m ²	27 (24-32)	26 (23-29)	29 (26-33)
Hypertension, n (%)	99 (44)	39 (35)	60 (54)
Current smoker, n (%)	61 (27)	30 (27)	31 (28)
Family history of MI, n (%)	47 (21)	10 (9)	37 (33)
Lipid status			
LDL-cholesterol, mg/dl	120 (93-146)	136 (119-162)	99 (79-121)
HDL-cholesterol, mg/dl	46 (39-55)	51 (42-58)	42 (37-52)
LDL/HDL-ratio	2.5 (1.8-3.3)	2.7 (2.2-3.4)	2.3 (1.7-3.1)

Continuous variables are described by median values (25th percentile to 75th percentile). Dichotomous variables are presented as total numbers (%). HDL = high-density lipoprotein, LDL=low-density lipoprotein, MI=myocardial infarction, NA=data not available

Table S2. Significantly differentially expressed genes between 112 early-onset MI patients and sex- and age-matched controls from the GIS discovery cohort.

Table S2 is provided at the end of this file

Table S3. Study characteristics of the early-onset MI patients and controls of the MIYoung study.

Characteristics	All	Controls	MI cases
n (%)	376	185 (49)	191 (51)
Females, n (%)	23 (6)	0 (0)	23 (12)
Age, years	65 (48-70)	70 (67-73)	48 (43-51)
Age at MI	44 (39-47)	NA	44 (39-47)
Cardiovascular risk factors			
Diabetes mellitus, n (%)	39 (10)	16 (9)	23 (12)
Body mass index, kg/m ²	27 (24-29)	26 (24-28)	28 (25-31)
Hypertension, n (%)	155 (41)	66 (36)	89 (47)
Current smoker, n (%)	55 (15)	8 (4)	47 (25)
Family history of MI, n (%)	106 (28)	29 (16)	77 (40)
Lipid status			
LDL-cholesterol, mg/dl	99 (74-125)	115 (93-137)	82 (55-103)
HDL-cholesterol, mg/dl	48 (41-60)	56 (46-66)	43 (36-51)
LDL/HDL-ratio	1.9 (1.4-2.5)	2.0 (1.5-2.5)	1.8 (1.2-2.3)

Continuous variables are described by median values (25th percentile to 75th percentile). Dichotomous variables are presented as total numbers (%). HDL = high-density lipoprotein, LDL=low-density lipoprotein, MI=myocardial infarction, NA=data not available

Table S4. Study characteristics of incident MI individuals and controls of the FIA2 cohort.

Characteristics	All	Controls	MI cases
n (%)	855	560 (65)	295 (35)
Females, n (%)	146 (17)	94 (17)	52 (18)
Age, years	53 (50-60)	53 (50-60)	54 (50-60)
Age at MI	57 (52-64)	NA	57 (52-64)
Current smoker, n (%)	252 (29)	118 (21)	134 (45)

Continuous variables are described by median values (25th percentile to 75th percentile). Dichotomous variables are presented as total numbers (%). MI=myocardial infarction, NA=data not available

Table S5. Incident myocardial infarction risk predicted by *GPR15* DNA methylation sites in the FIA2 study.

DNA methylation site	Hazard ratio	95% CI	p value
CpG1	0.988	0.973-1.004	0.3909
CpG2	0.993	0.982-1.004	0.6456
CpG3	0.993	0.987-0.999	0.0549
Adjusted for smoking			
CpG1	0.989	0.974-1.005	0.5685
CpG2	0.989	0.977-1.000	0.1812
CpG3	0.992	0.986-0.998	0.0177

Mixed effect Cox models, adjusted for age and sex, significance after Bonferroni correction, CI=confidence interval

Table S6. Study characteristics of the MIYoung study individuals selected for GPR15 sequencing.

Characteristics	Controls	MI cases
n (%)	32	32
Females, n (%)	32 (100)	32 (100)
Age, years	75 (71-76)	42 (38-49)
Age at MI	NA	38 (34-39)
<u>Cardiovascular risk factors</u>		
Diabetes mellitus, n (%)	3 (9)	5 (16)
Body mass index, kg/m ²	25 (24-28)	29 (27-33)
Hypertension, n (%)	15 (47)	18 (56)
Current smoker, n (%)	2 (6)	9 (28)
Family history of MI, n (%)	16 (50)	16 (50)
Lipid status		
LDL-cholesterol, mg/dl	113 (87-139)	87 (63-105)
HDL-cholesterol, mg/dl	55 (49-66)	45 (34-49)
LDL/HDL-ratio	1.8 (1.4-2.5)	2.0 (1.6-2.6)

Continuous variables are described by median values (25th percentile to 75th percentile). Dichotomous variables are presented as total numbers (%). HDL = high-density lipoprotein, LDL=low-density lipoprotein, MI=myocardial infarction, NA=data not available

Table S7. Associations between GPR15 DNA methylation and rs2230344 minor T allele in the MIYoung study.

DNA methylation site	Effect	Standard error	p value
CpG1	1.157	0.748	0.3690
CpG2	0.690	1.289	1.7790
CpG3	2.846	1.356	0.1083
Adjusted for smoking			
CpG1	0.972	0.749	0.5850
CpG2	1.292	1.293	0.9540
CpG3	2.821	1.367	0.1182

Linear mixed models with CpG methylation status as dependent and rs2230344 allele dosage as independent variable, adjusted for age and sex, effect=% difference per T allele, CI=confidence interval

Table S8. Risk prediction of incident myocardial infarction by rs2230344 minor T allele in the FIA2 cohort.

Subjects	Hazard ratio	95% CI	p value
All	0.947	0.757-1.185	0.6352
Never smokers only	0.837	0.544-1.288	0.4191
Ever smokers only	1.044	0.801-1.360	0.7522

Mixed effect Cox models, adjusted for age and sex, CI=confidence interval

Table S9. Significantly differentially expressed genes between *Gpr15^{gfp/gfp}* and wildtype mice in the myocardial infarction zone.

Table S9 is provided at the end of this file

Table S10. Cohort description.

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	GIS [32-36]	MIYoung	FIA2 [37, 38]
Rationale	Discovery	Replication	Risk prediction
Study design	Retrospective case-control study	Retrospective case-control study	Prospective cohort/retrospective case-control study
Country	Germany	Germany	Sweden
Ethnicity	Caucasian	Caucasian	Caucasian
Cases	<ul style="list-style-type: none"> ○ Prevalent early onset MI ○ n = 234 	<ul style="list-style-type: none"> ○ Prevalent early onset MI ○ n = 191 	<ul style="list-style-type: none"> ○ Incident MI ○ n = 295
Case definition	<ul style="list-style-type: none"> ○ Confirmed MI (positive ECG, increase in biomarkers of cardiac necrosis (troponin, creatine-kinase MB), or clinical symptoms) and proven by coronary angiography ○ Age at MI ≤ 50 years ○ Stable disease conditions for at least three months before study participation 	<ul style="list-style-type: none"> ○ Confirmed MI (positive ECG, increase in biomarkers of cardiac necrosis (troponin, creatine-kinase MB), or clinical symptoms) and proven by coronary angiography ○ Age at MI ≤ 50 years ○ Stable disease conditions for at least three months before study participation 	<ul style="list-style-type: none"> ○ MI events were identified through screening of hospital discharge records, general practitioners' reports, and death certificates. ○ Age at MI = 34-71 years ○ No prevalent MI cases
Eligible controls	<ul style="list-style-type: none"> ○ No manifest cardiovascular disease ○ n = 419 	<ul style="list-style-type: none"> ○ No manifest cardiovascular disease ○ Age > 65 years ○ n = 185 	<ul style="list-style-type: none"> ○ No prevalent and incident MI ○ n = 560
Matching criteria	<ul style="list-style-type: none"> ○ Sex ○ Age 	-	<ul style="list-style-type: none"> ○ Sex ○ Age ○ Date of health examination ○ Geographical region
Number of controls per case	1	-	2
Number of matched samples for analyses	112 (112 cases, 112 controls)	-	170 (170 cases, 340 controls)
Samples	<ul style="list-style-type: none"> ○ Smoking (self-reported) ○ RNA from PBMCs ○ DNA from leukocytes 	<ul style="list-style-type: none"> ○ Smoking (self-reported) ○ RNA from PBMCs ○ DNA from leukocytes 	<ul style="list-style-type: none"> ○ Smoking (self-reported) ○ DNA from whole blood

Ethical approval	local ethics committee of the Medical Chamber of Rhineland-Palatinate, Germany (ethical approval code 837.020.07 (5555))	local ethics committee of the University Medical Center Hamburg-Eppendorf, Germany (ethical approval code PV4137)	local ethics committee (Dnr 05-142M, with additional review in 2009-01-19)
Funding	Government of Rhineland-Palatinate ("Stiftung Rheinland-Pfalz für Innovation"), grant AZ 961-386261/733; "Wissen schafft Zukunft"; Johannes Gutenberg-University of Mainz "Center for Translational Vascular Biology"; Boehringer Ingelheimer, PHILIPS Medical Systems	-	Swedish Heart and Lung Foundation; Gustaf Sjölund Foundation; County councils of Northern Sweden (Visare Norr) and the Faculty of Medicine of Umeå University

MI = myocardial infarction, PBMCs = peripheral blood mononuclear cells

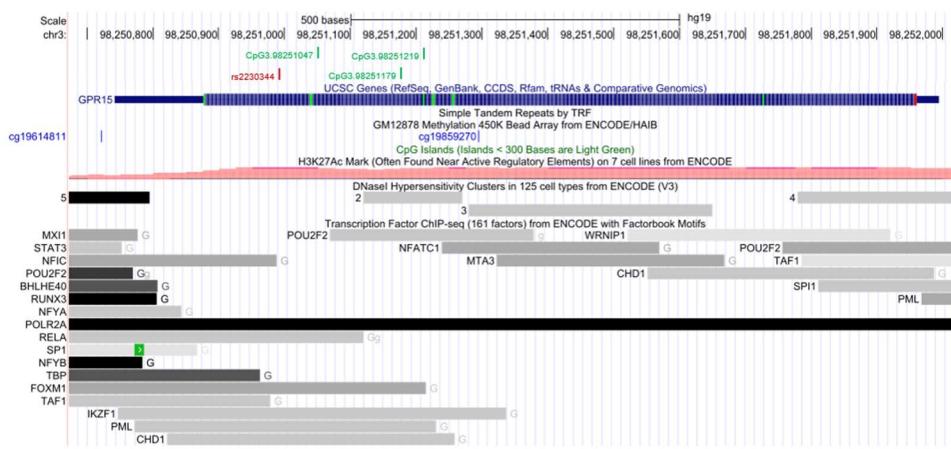


Figure S1. The *GPR15* locus in the UCSC genome browser. The *GPR15* locus depicted in UCSC Genome Browser on Human Feb. 2009 (GRCh37/hg19) Assembly, showing the *GPR15* gene, *GPR15* DNA methylation sites and rs2230344 SNP in close proximity to each other. <http://genome.ucsc.edu/>, 02/04/2020.

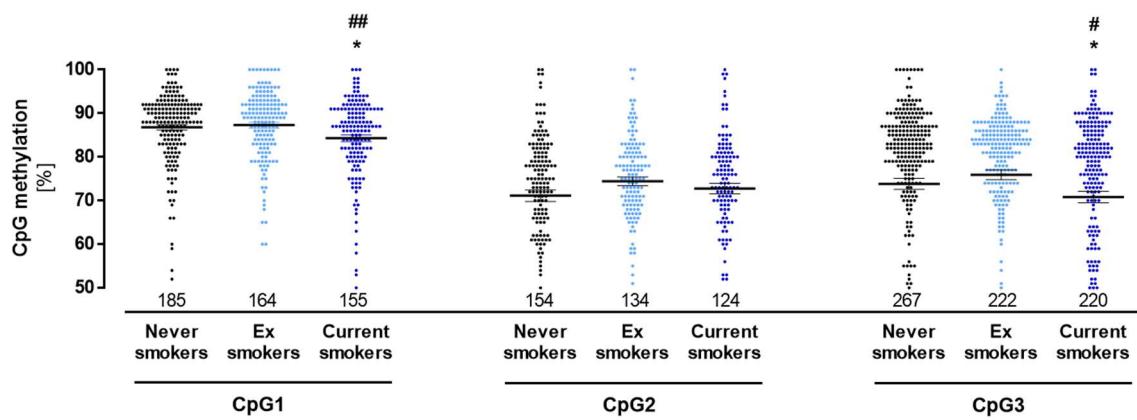


Figure S2. Increased *GPR15* expression and decreased *GPR15* DNA methylation in smokers. In the FIA2 study, *GPR15* DNA methylation was significantly lower in current smokers compared to never smokers and ex smokers for the CpG1 site ($p<0.05$ and $p<0.01$) and for the CpG3 site ($p<0.05$). Kruskal-Wallis test with Dunn's multiple comparison; significance threshold=0.0167 for Bonferroni correction, *#/#= $p<0.05$, ##= $p<0.01$ compared to the respective never smokers group/ex smokers group; Data are shown as mean \pm standard error of the mean.

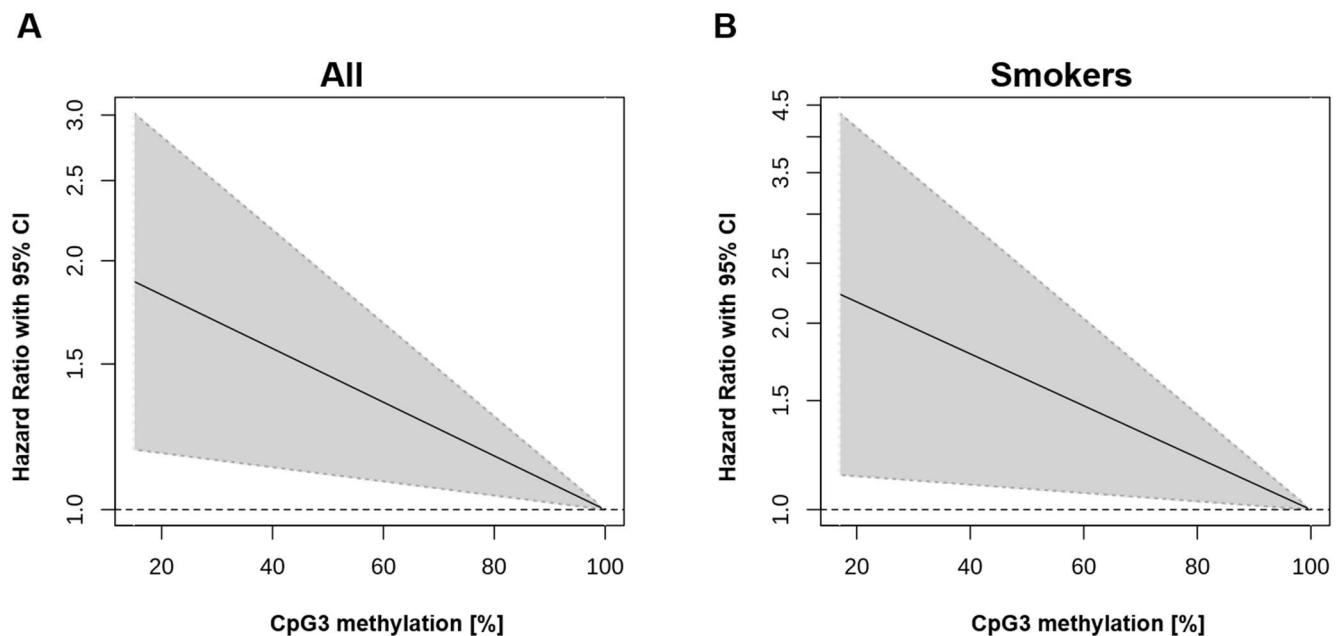


Figure S3. Prediction of risk of incident myocardial infarction by GPR15 DNA methylation in the FIA2 study. CpG3 methylation significantly predicted the risk for incident MI after adjustment for smoking ($HR=0.992$, $p=0.0177$). MI risk was lower in individuals with higher CpG3 DNA methylation levels compared to individuals with lower CpG3 DNA methylation levels. This was shown in the overall cohort (A) as well as in smokers only (B). *R* package *smoothHR*, Full CpG3 site methylation (100%) was used as reference. Grey areas represent 95% confidence intervals (CI).

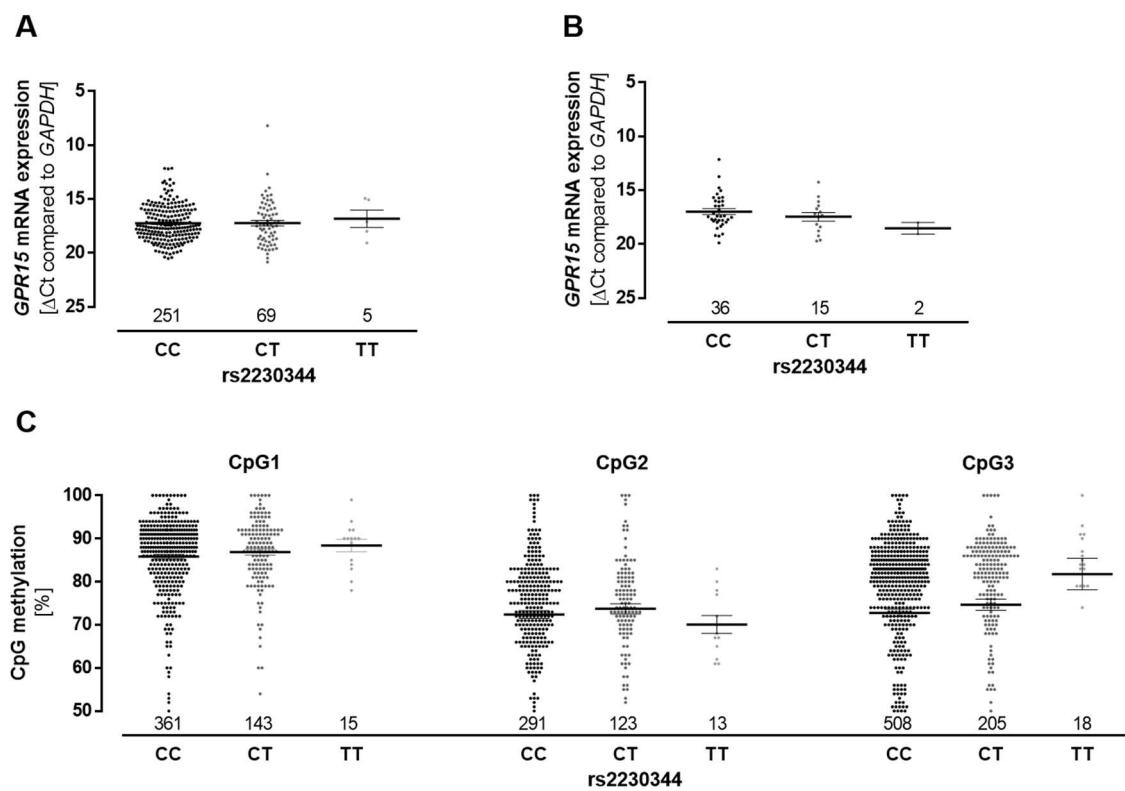


Figure S4. rs2230344 genotype is not associated with GPR15 DNA methylation or GPR15 mRNA expression. (A/B) In the MIYoung study, rs2230344 SNP genotype was not significantly associated with GPR15 mRNA expression in all individuals (A) or individuals with sequenced genotypes only (B). GPR15 mRNA expression is depicted as ΔCt values normalized to glyceraldehyde 3-phosphate dehydrogenase (GAPDH) mRNA expression. Lower ΔCt values indicate higher GPR15 mRNA expression. (C) In the FIA2 study, GPR15 DNA methylation of the CpG3 site was not significantly associated with rs2230344 genotype. n=13-508, linear mixed model, adjusted for age and sex; significance after Bonferroni correction Data are shown as mean \pm standard error of the mean.

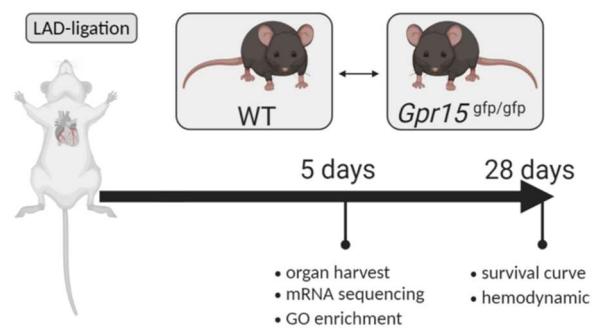


Figure S5. Overview *Gpr15^{gfp/gfp}* mice. *Gpr15^{gfp/gfp}* and wildtype (WT) mice were subjected to left anterior descending artery (LAD) ligation and either mRNA sequencing of the infarct zone (IZ) was performed 5 days after ligation or survival was determined 28 days after ligation.

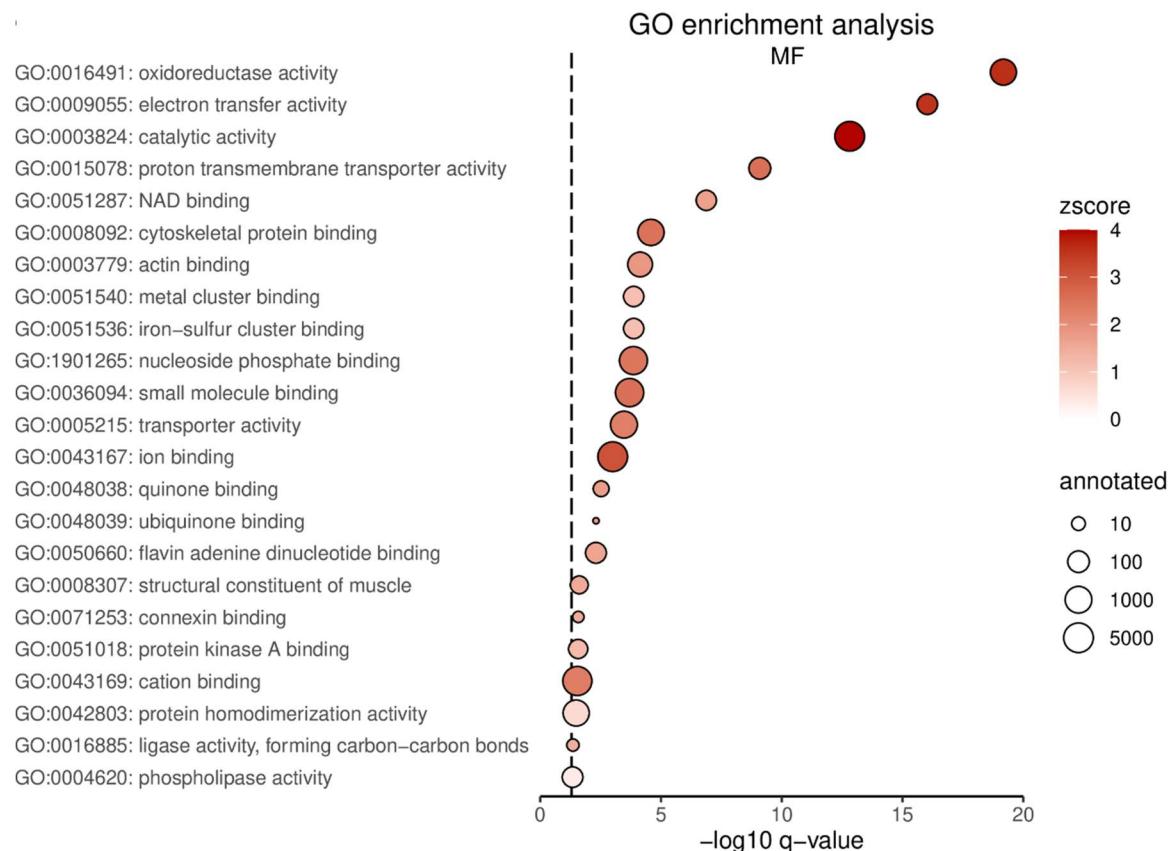


Figure S6. Altered GO terms for molecular functions in *Gpr15^{gfp/gfp}* mice. Gene expression measured by 3' mRNA sequencing of cardiac tissue from the IZ from 9 Bl6 wildtype (WT) and 8 *Gpr15^{gfp/gfp}* mice 5 days after MI surgery. Gene Ontology (GO) terms are displayed for molecular functions (MF).

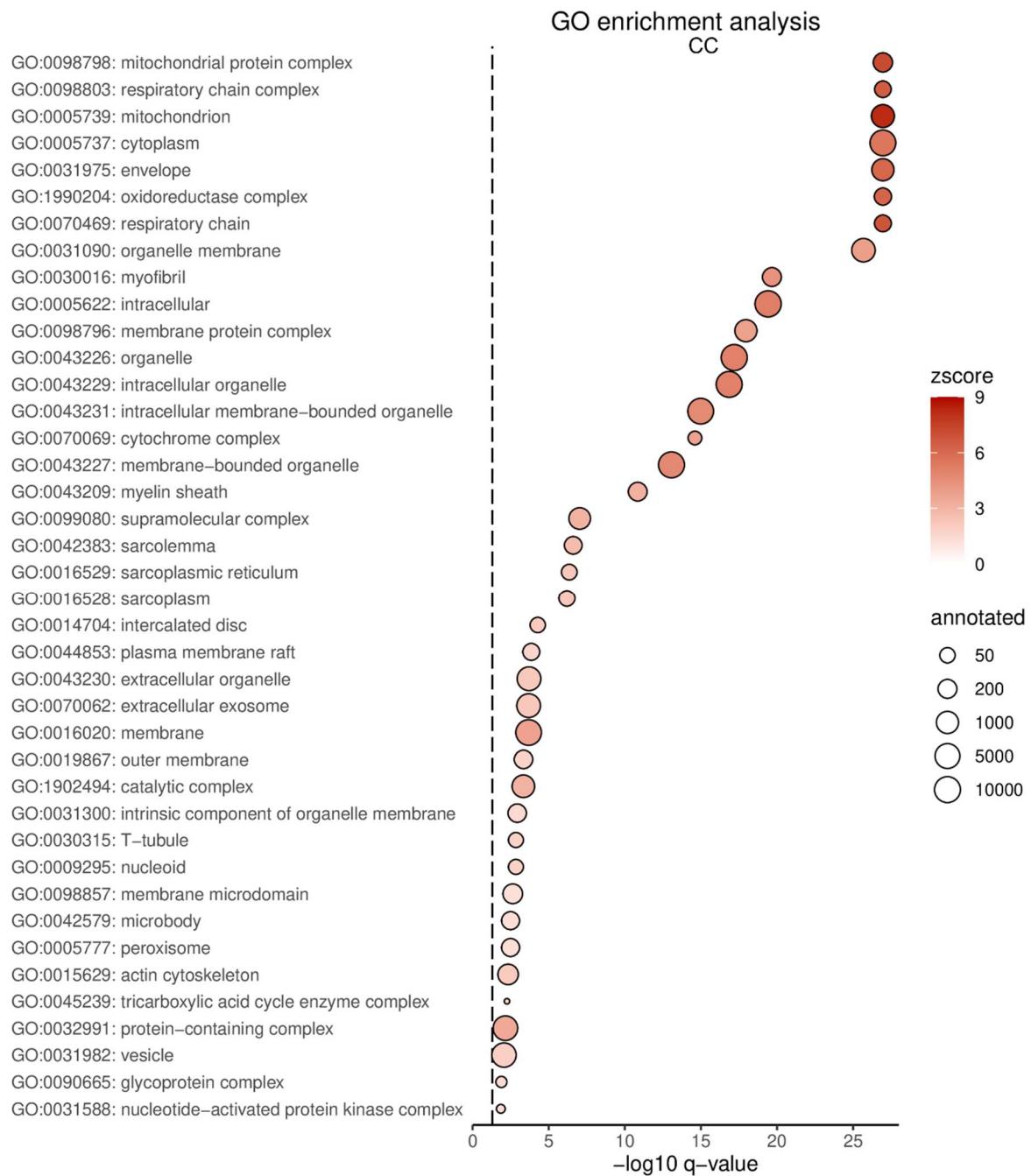


Figure S7. Altered GO terms for cellular components in *Gpr15^{gfp/gfp}* mice. Gene expression measured by 3' mRNA sequencing of cardiac tissue from the IZ from 9 Bl6 wildtype (WT) and 8 *Gpr15^{gfp/gfp}* mice 5 days after MI surgery. Gene Ontology (GO) terms are displayed for cellular components (CC).

Table S2: Significantly differentially expressed genes between 112 early-onset MI patients and sex- and age-matched controls from the GIS discovery cohort.

gene	Ensembl_id	FC	p-value	q-value	gene description
GPR15	ENSG00000154165	1.43E+00	1.87E-07	2.88E-03	G protein-coupled receptor 15
ABCA1	ENSG00000165029	-1.14E+00	4.54E-07	3.51E-03	ATP-binding cassette, sub-family A (ABC1), member 1
SELO	ENSG00000073169	-1.18E+00	1.35E-06	6.05E-03	selenoprotein O
CTF1	ENSG00000150281	1.15E+00	1.61E-06	6.05E-03	cardiotrophin 1
DUSP1	ENSG00000120129	-1.34E+00	2.19E-06	6.05E-03	dual specificity phosphatase 12 (EC:3.1.3.16 3.1.3.48)
DNM2	ENSG00000079805	-1.08E+00	2.35E-06	6.05E-03	dynamin 2 (EC:3.6.5.5)
DYNLL2	ENSG00000121083	-1.14E+00	4.05E-06	7.37E-03	dynein, light chain, LC8-type 2
CMIP	ENSG00000153815	-1.11E+00	4.25E-06	7.37E-03	CMIP
TMEM127	ENSG00000135956	-1.14E+00	4.29E-06	7.37E-03	transmembrane protein 127
GALT	ENSG00000213930	1.09E+00	5.36E-06	8.28E-03	galactose-1-phosphate uridylyltransferase (EC:2.7.7.12)
CLK3	ENSG00000179335	1.09E+00	7.80E-06	9.42E-03	CDC-like kinase 3 (EC:2.7.12.1)
IGF2R	ENSG00000197081	-1.09E+00	7.96E-06	9.42E-03	insulin-like growth factor 2 receptor
ADCY7	ENSG00000121281	-1.08E+00	8.00E-06	9.42E-03	adenylate cyclase 7 (EC:4.6.1.1)
ST6GALNAC3	ENSG00000184005	-1.13E+00	9.63E-06	9.42E-03	ST6
PIK3CD	ENSG00000171608	-1.06E+00	9.75E-06	9.42E-03	phosphoinositide-3-kinase, catalytic, delta polypeptide
MERTK	ENSG00000153208	-1.18E+00	1.03E-05	9.42E-03	c-mer proto-oncogene tyrosine kinase (EC:2.7.10.1)
SIPA1L1	ENSG00000197555	-1.12E+00	1.09E-05	9.42E-03	signal-induced proliferation-associated 1 like 1
HIP1R	ENSG00000130787	1.09E+00	1.10E-05	9.42E-03	huntingtin interacting protein 1 related
HDAC9	ENSG00000048052	1.08E+00	1.22E-05	9.90E-03	histone deacetylase 9 (EC:3.5.1.98)
C1orf38	ENSG00000130775	-1.17E+00	1.30E-05	1.01E-02	chromosome 1 open reading frame 38
SPRN1	ENSG00000230786	-1.10E+00	1.46E-05	1.02E-02	NA
SPTAN1	ENSG00000197694	1.09E+00	1.51E-05	1.02E-02	spectrin, alpha, non-erythrocytic 1 (alpha-fodrin)
EPHA4	ENSG00000116106	1.13E+00	1.52E-05	1.02E-02	EPH receptor A4 (EC:2.7.10.1)
XXbac-B461K10.4	ENSG00000093100	1.09E+00	1.77E-05	1.07E-02	NA
OSGEP	ENSG00000092094	1.09E+00	1.80E-05	1.07E-02	O-sialoglycoprotein endopeptidase (EC:3.4.24.57)
BASP1	ENSG00000176788	-1.12E+00	1.80E-05	1.07E-02	brain abundant, membrane attached signal protein 1
SLC11A1	ENSG00000182820	-1.16E+00	2.57E-05	1.47E-02	solute carrier family 11 (proton-coupled divalent metal ion
PPP1R15B	ENSG00000158615	-1.09E+00	2.84E-05	1.57E-02	protein phosphatase 1, regulatory (inhibitor) subunit 15B
P2RY2	ENSG00000175591	-1.17E+00	2.94E-05	1.57E-02	purinergic receptor P2Y, G-protein coupled, 2
SNX12	ENSG00000147164	1.10E+00	3.08E-05	1.57E-02	sorting nexin 12
CD93	ENSG00000125810	-1.13E+00	3.16E-05	1.57E-02	CD93 molecule
MYO19	ENSG00000141140	1.07E+00	3.55E-05	1.71E-02	myosin XIX
VAMP7	ENSG00000124333	1.20E+00	4.03E-05	1.85E-02	vesicle-associated membrane protein 7
IL1R2	ENSG00000115590	-1.13E+00	4.19E-05	1.85E-02	interleukin 1 receptor, type II
IKZF3	ENSG00000161405	1.14E+00	4.20E-05	1.85E-02	IKAROS family zinc finger 3 (Aiolos)
CASK	ENSG00000147044	1.07E+00	4.50E-05	1.90E-02	calcium/calmodulin-dependent serine protein kinase (MAGUK family)
TNFAIP3	ENSG00000118503	-1.09E+00	4.54E-05	1.90E-02	tumor necrosis factor, alpha-induced protein 3
ADAM23	ENSG00000114948	1.13E+00	4.88E-05	1.98E-02	ADAM metallopeptidase domain 23
CYB5B	ENSG00000103018	1.10E+00	5.51E-05	2.13E-02	cytochrome b5 type B (outer mitochondrial membrane)
JARID2	ENSG00000008083	-1.07E+00	5.52E-05	2.13E-02	jumonji, AT rich interactive domain 2
SOAT1	ENSG00000057252	-1.11E+00	5.89E-05	2.22E-02	sterol O-acyltransferase 1 (EC:2.3.1.26)
AP000679.2	ENSG00000176984	1.25E+00	6.19E-05	2.26E-02	NA
STX6	ENSG00000135823	-1.07E+00	6.28E-05	2.26E-02	syntaxin 6
AC129778.1	ENSG00000230059	-1.10E+00	6.46E-05	2.27E-02	NA
U47924_25	ENSG00000237240	1.25E+00	6.84E-05	2.35E-02	NA
HIST1H2BK	ENSG00000197903	-1.15E+00	8.04E-05	2.70E-02	histone cluster 1, H2bk
PRKD3	ENSG00000115825	-1.08E+00	8.99E-05	2.96E-02	protein kinase D3 (EC:2.7.11.13)
TREM1	ENSG00000124731	-1.28E+00	9.31E-05	3.00E-02	triggering receptor expressed on myeloid cells 1
SPCS3	ENSG00000129128	1.16E+00	9.61E-05	3.03E-02	signal peptide complex subunit 3 homolog (S. cerevisiae)
ALDH2	ENSG00000111275	-1.15E+00	9.90E-05	3.03E-02	aldehyde dehydrogenase 2 family (mitochondrial) (EC:1.2.1.3)
TNFAIP2	ENSG00000185215	-1.13E+00	9.99E-05	3.03E-02	tumor necrosis factor, alpha-induced protein 2
ARAP1	ENSG00000186635	-1.08E+00	1.02E-04	3.04E-02	ArfGAP with RhoGAP domain, ankyrin repeat and PH domain 1
MAFK	ENSG00000198517	-1.10E+00	1.08E-04	3.14E-02	v-maf musculoaponeurotic fibrosarcoma oncogene homolog K (avian)
RPL23	ENSG00000125691	1.16E+00	1.10E-04	3.14E-02	mitochondrial ribosomal protein L23
EXOSC1	ENSG00000171311	1.09E+00	1.21E-04	3.36E-02	exosome component 1
NELL2	ENSG00000184613	1.15E+00	1.22E-04	3.36E-02	NEL-like 2 (chicken)
FAM122A	ENSG00000187866	1.11E+00	1.29E-04	3.51E-02	family with sequence similarity 122A
WDR49	ENSG00000174776	-1.09E+00	1.34E-04	3.57E-02	WD repeat domain 49
MYLIP	ENSG00000007944	-1.08E+00	1.42E-04	3.71E-02	myosin regulatory light chain interacting protein (EC:6.3.2.-)
BCKDHB	ENSG00000083123	-1.11E+00	1.49E-04	3.75E-02	branched chain keto acid dehydrogenase E1, beta polypeptide
NA	ENSG00000188666	-1.14E+00	1.56E-04	3.75E-02	NA
LRRK70	ENSG00000186105	-1.14E+00	1.56E-04	3.75E-02	leucine rich repeat containing 70
PGS1	ENSG00000087157	-1.07E+00	1.57E-04	3.75E-02	phosphatidylglycerophosphate synthase 1 (EC:2.7.8.5)
CXorf65	ENSG00000204165	1.13E+00	1.57E-04	3.75E-02	chromosome X open reading frame 65
C6orf52	ENSG00000137434	-1.11E+00	1.58E-04	3.75E-02	chromosome 6 open reading frame 52
DTW2D	ENSG00000169570	1.09E+00	1.68E-04	3.94E-02	DTW domain containing 2
CTSZ	ENSG00000101160	-1.09E+00	1.77E-04	4.04E-02	cathepsin Z (EC:3.4.18.1)
ZSWIM6	ENSG00000130449	-1.10E+00	1.78E-04	4.04E-02	zinc finger, SWIM-type containing 6
RNF145	ENSG00000145860	-1.08E+00	1.85E-04	4.15E-02	ring finger protein 145
MAK	ENSG00000111837	-1.10E+00	2.03E-04	4.42E-02	male germ cell-associated kinase (EC:2.7.11.22)
PIGM	ENSG00000143315	-1.09E+00	2.03E-04	4.42E-02	phosphatidylinositol glycan anchor biosynthesis, class M
AMFR	ENSG00000159461	-1.11E+00	2.07E-04	4.45E-02	autocrine motility factor receptor (EC:6.3.2.19)
MIOS	ENSG00000164654	-1.08E+00	2.26E-04	4.67E-02	missing oocyte, meiosis regulator, homolog (Drosophila)
CRISPLD2	ENSG00000103196	-1.18E+00	2.26E-04	4.67E-02	cysteine-rich secretory protein LCLL domain containing 2
SERPINE2	ENSG00000135919	-1.09E+00	2.27E-04	4.67E-02	serpin peptidase inhibitor, clade E (nexin, plasminogen activator
BMP6	ENSG00000153162	-1.15E+00	2.36E-04	4.80E-02	bone morphogenetic protein 6
KIAA0930	ENSG00000100364	-1.09E+00	2.45E-04	4.88E-02	NA
HINT1	ENSG00000169567	1.13E+00	2.47E-04	4.88E-02	histidine triad nucleotide binding protein 1 (EC:3.-.-.-)
C7orf66	ENSG00000205174	1.12E+00	2.50E-04	4.88E-02	chromosome 7 open reading frame 66
DUS4L	ENSG00000105865	-1.09E+00	2.64E-04	5.07E-02	dihydrodouridine synthase 4-like (S. cerevisiae)
KLHL24	ENSG00000114796	-1.08E+00	2.66E-04	5.07E-02	kelch-like 24 (Drosophila)
GALNT2	ENSG00000143641	-1.06E+00	2.70E-04	5.09E-02	UDP-N-acetyl-alpha-D-galactosamine:polypeptide
RNF125	ENSG00000101695	1.10E+00	2.79E-04	5.20E-02	ring finger protein 125
STX11	ENSG00000135604	-1.14E+00	2.87E-04	5.25E-02	syntaxin 11
AGER	ENSG00000204305	-1.08E+00	2.89E-04	5.25E-02	advanced glycosylation end product-specific receptor
HSPB1	ENSG00000106211	1.10E+00	2.92E-04	5.25E-02	heat shock 27kDa protein 1
ARF4	ENSG00000168374	1.15E+00	3.03E-04	5.34E-02	ADP-ribosylation factor 4
TMEM116	ENSG00000198270	1.11E+00	3.04E-04	5.34E-02	transmembrane protein 116
TNFRSF21	ENSG00000146072	-1.09E+00	3.18E-04	5.52E-02	tumor necrosis factor receptor superfamily, member 21
MYH13	ENSG00000006788	1.07E+00	3.23E-04	5.54E-02	myosin, heavy chain 13, skeletal muscle
TSC22D3	ENSG00000157514	-1.06E+00	3.28E-04	5.57E-02	TSC22 domain family, member 3
WDFY1	ENSG00000085449	-1.08E+00	3.40E-04	5.66E-02	WD repeat and FYVE domain containing 1
RALGPS1	ENSG00000136828	1.06E+00	3.40E-04	5.66E-02	Ral GEF with PH domain and SH3 binding motif 1
ITGB7	ENSG00000139626	1.09E+00	3.49E-04	5.71E-02	integrin, beta 7
NDUFV2	ENSG00000178127	1.16E+00	3.51E-04	5.71E-02	NADH dehydrogenase (ubiquinone) flavoprotein 2, 24kDa (EC:1.6.5.3)

<i>OPRD1</i>	ENSG00000116329	-1,11E+00	3,58E-04	5,77E-02	opioid receptor, delta 1
<i>SPINK8</i>	ENSG00000229453	-1,13E+00	3,63E-04	5,79E-02	serine peptidase inhibitor, Kazal type 8 (putative)
<i>MYCL1</i>	ENSG00000116990	-1,10E+00	3,70E-04	5,84E-02	v-myc myelocytomatosis viral oncogene homolog 1, lung carcinoma
<i>ABCD2</i>	ENSG00000173208	1,10E+00	4,16E-04	6,32E-02	ATP-binding cassette, sub-family D (ALD), member 2
<i>STX2</i>	ENSG00000111450	1,13E+00	4,18E-04	6,32E-02	syntaxin 2
<i>MAP3K6</i>	ENSG00000142733	1,06E+00	4,21E-04	6,32E-02	mitogen-activated protein kinase kinase kinase 6 (EC:2.7.11.25)
<i>PLCB4</i>	ENSG00000101333	-1,07E+00	4,21E-04	6,32E-02	phospholipase C, beta 4 (EC:3.1.4.11)
<i>DOK3</i>	ENSG00000146094	-1,13E+00	4,28E-04	6,32E-02	docking protein 3
<i>MGEA5</i>	ENSG00000198408	1,11E+00	4,33E-04	6,32E-02	meningioma expressed antigen 5 (hyaluronidase) (EC:3.2.1.52)
<i>MTO1</i>	ENSG00000135297	1,08E+00	4,34E-04	6,32E-02	mitochondrial translation optimization 1 homolog (S. cerevisiae)
<i>SH2B3</i>	ENSG00000111252	-1,06E+00	4,37E-04	6,32E-02	SH2B adaptor protein 3
<i>UNC93B1</i>	ENSG00000110057	-1,11E+00	4,47E-04	6,32E-02	unc-93 homolog B1 (C. elegans)
<i>C19orf28</i>	ENSG00000161091	-1,07E+00	4,53E-04	6,32E-02	chromosome 19 open reading frame 28
<i>VSTM1</i>	ENSG00000189068	-1,18E+00	4,57E-04	6,32E-02	V-set and transmembrane domain containing 1
<i>SLC6A6</i>	ENSG00000131389	-1,12E+00	4,59E-04	6,32E-02	solute carrier family 6 (neurotransmitter transporter, taurine),
<i>ESRP1</i>	ENSG00000104413	-1,08E+00	4,62E-04	6,32E-02	epithelial splicing regulatory protein 1
<i>PTRF</i>	ENSG00000177469	1,10E+00	4,67E-04	6,32E-02	polymerase I and transcript release factor
<i>IL23A</i>	ENSG00000110944	1,07E+00	4,68E-04	6,32E-02	interleukin 23, alpha subunit p19
<i>STS</i>	ENSG00000101846	-1,10E+00	4,72E-04	6,32E-02	steroid sulfatase (microsomal), isozyme S (EC:3.1.6.2)
<i>PCYT1B</i>	ENSG00000102230	-1,11E+00	4,74E-04	6,32E-02	phosphate cytidyltransferase 1, choline, beta (EC:2.7.7.15)
<i>MCL1</i>	ENSG00000143384	-1,12E+00	4,74E-04	6,32E-02	myeloid cell leukemia sequence 1 (BCL2-related)
<i>CDK5RAP1</i>	ENSG00000101391	1,07E+00	4,92E-04	6,48E-02	CDK5 regulatory subunit associated protein 1
<i>TRPS1</i>	ENSG00000104447	-1,10E+00	4,98E-04	6,48E-02	trichorhinophalangeal syndrome I
<i>ZNF812</i>	ENSG00000224689	1,15E+00	4,99E-04	6,48E-02	zinc finger protein 812
<i>DAPK1</i>	ENSG00000196730	-1,11E+00	5,03E-04	6,48E-02	death-associated protein kinase 1 (EC:2.7.11.1)
<i>TMEM110</i>	ENSG00000213533	-1,10E+00	5,14E-04	6,51E-02	transmembrane protein 110
<i>TMEM110-MUSTN1</i>	ENSG00000248592	-1,10E+00	5,14E-04	6,51E-02	NA
<i>JAZF1</i>	ENSG00000153814	1,09E+00	5,24E-04	6,58E-02	JAZF zinc finger 1
<i>HSPA2</i>	ENSG00000126803	1,10E+00	5,43E-04	6,77E-02	heat shock 70kDa protein 2
<i>POGK</i>	ENSG00000143157	-1,07E+00	5,62E-04	6,95E-02	pogo transposable element with KRAB domain
<i>HIST2H2BE</i>	ENSG00000184678	-1,26E+00	5,71E-04	7,00E-02	histone cluster 2, H2be
<i>NA</i>	ENSG00000141325	1,07E+00	5,83E-04	7,05E-02	NA
<i>SLC43A2</i>	ENSG00000167703	-1,11E+00	5,84E-04	7,05E-02	solute carrier family 43, member 2
<i>MCF2L</i>	ENSG00000126217	1,05E+00	5,98E-04	7,15E-02	MCF-2 cell line derived transforming sequence-like
<i>ATF7IP2</i>	ENSG00000166669	1,13E+00	6,02E-04	7,15E-02	activating transcription factor 7 interacting protein 2
<i>ABCG1</i>	ENSG00000160179	-1,08E+00	6,06E-04	7,15E-02	ATP-binding cassette, sub-family G (WHITE), member 1
<i>OR52J3</i>	ENSG00000205495	1,16E+00	6,22E-04	7,28E-02	olfactory receptor, family 52, subfamily J, member 3
<i>ASB13</i>	ENSG00000196372	1,13E+00	6,34E-04	7,37E-02	ankyrin repeat and SOCS box-containing 13
<i>AGTRAP</i>	ENSG00000177674	-1,10E+00	6,42E-04	7,41E-02	angiotensin II receptor-associated protein
<i>ETAA1</i>	ENSG00000143971	1,11E+00	6,58E-04	7,54E-02	Ewing tumor-associated antigen 1
<i>CXCL16</i>	ENSG00000161921	-1,18E+00	6,67E-04	7,58E-02	chemokine (C-X-C motif) ligand 16
<i>SLC40A1</i>	ENSG00000138449	-1,14E+00	6,80E-04	7,63E-02	solute carrier family 40 (iron-regulated transporter), member 1
<i>PPM1G</i>	ENSG00000115241	1,07E+00	6,81E-04	7,63E-02	protein phosphatase, Mg ²⁺ /Mn ²⁺ dependent, 1G (EC:3.1.3.16)
<i>C19orf18</i>	ENSG00000177025	1,14E+00	6,97E-04	7,74E-02	chromosome 19 open reading frame 18
<i>PTGER2</i>	ENSG00000125384	1,12E+00	7,05E-04	7,78E-02	prostaglandin E receptor 2 (subtype EP2), 53kDa
<i>CCDC77</i>	ENSG00000120647	-1,11E+00	7,18E-04	7,87E-02	coiled-coil domain containing 77
<i>PRKAG2</i>	ENSG00000106617	-1,06E+00	7,27E-04	7,91E-02	protein kinase, AMP-activated, gamma 2 non-catalytic subunit
<i>RELA</i>	ENSG00000173039	1,09E+00	7,64E-04	8,26E-02	v-rel reticuloendotheliosis viral oncogene homolog A (avian)
<i>SGMS2</i>	ENSG00000164023	-1,10E+00	7,92E-04	8,50E-02	spingomyelin synthase 2 (EC:2.7.8.27)
<i>ZDHHC18</i>	ENSG00000204160	1,06E+00	8,10E-04	8,63E-02	zinc finger, DHHC-type containing 18
<i>AC008021.1</i>	ENSG00000206043	-1,15E+00	8,21E-04	8,69E-02	NA
<i>C2CD3</i>	ENSG00000168014	1,05E+00	8,51E-04	8,95E-02	C2 calcium-dependent domain containing 3
<i>EDEM3</i>	ENSG00000116406	-1,08E+00	8,62E-04	8,97E-02	ER degradation enhancer,mannosidase alpha-like 3
<i>RIPK3</i>	ENSG00000129465	1,08E+00	8,80E-04	8,97E-02	receptor-interacting serine-threonine kinase 3 (EC:2.7.11.1)
<i>ST14</i>	ENSG00000149418	-1,08E+00	8,85E-04	8,97E-02	suppression of tumorigenicity 14 (colon carcinoma) (EC:3.4.21.109)
<i>IRF2BP2</i>	ENSG00000168264	-1,07E+00	8,92E-04	8,97E-02	interferon regulatory factor 2 binding protein 2
<i>CD300A</i>	ENSG00000167851	-1,13E+00	8,94E-04	8,97E-02	CD300a molecule
<i>USP14</i>	ENSG00000101557	1,07E+00	9,10E-04	8,97E-02	ubiquitin specific peptidase 14 (tRNA-guanine transglycosylase)
<i>CCDC56</i>	ENSG00000183978	1,23E+00	9,10E-04	8,97E-02	coiled-coil domain containing 56
<i>MARK1</i>	ENSG00000116141	1,08E+00	9,12E-04	8,97E-02	MAP/microtubule affinity-regulating kinase 1 (EC:2.7.11.1)
<i>C5orf17</i>	ENSG00000248874	1,11E+00	9,19E-04	8,97E-02	NA
<i>MICAL3</i>	ENSG00000243156	1,08E+00	9,20E-04	8,97E-02	microtubule associated monooxygenase, calponin and LIM domain
<i>EIF4ENIF1</i>	ENSG00000184708	1,05E+00	9,25E-04	8,97E-02	eukaryotic translation initiation factor 4E nuclear import factor 1
<i>ECE1</i>	ENSG00000117298	-1,08E+00	9,27E-04	8,97E-02	endothelin converting enzyme 1 (EC:3.4.24.71)
<i>INPP5D</i>	ENSG00000168918	-1,06E+00	9,29E-04	8,97E-02	inositol polyphosphate-5-phosphatase, 145kDa
<i>POU3F4</i>	ENSG00000196767	1,10E+00	9,37E-04	8,99E-02	POU class 3 homeobox 4
<i>MRO</i>	ENSG00000134042	-1,09E+00	9,46E-04	8,99E-02	maestro
<i>RP11-698N11.3</i>	ENSG00000186082	1,11E+00	9,48E-04	8,99E-02	NA
<i>NKX3-1</i>	ENSG00000167034	1,07E+00	9,58E-04	9,03E-02	NK3 homeobox 1
<i>DDX19A</i>	ENSG00000168872	1,08E+00	9,78E-04	9,11E-02	DEAD (Asp-Glu-Ala-As) box polypeptide 19A
<i>LDLR</i>	ENSG00000130164	1,09E+00	9,78E-04	9,11E-02	low density lipoprotein receptor
<i>CPA4</i>	ENSG00000128510	-1,08E+00	9,85E-04	9,12E-02	carboxypeptidase A4
<i>DEPD5</i>	ENSG00000100150	1,05E+00	1,00E-03	9,21E-02	DEP domain containing 5
<i>SLC26A8</i>	ENSG00000112053	-1,08E+00	1,02E-03	9,26E-02	solute carrier family 26, member 8
<i>ROMO1</i>	ENSG00000125995	1,15E+00	1,02E-03	9,26E-02	reactive oxygen species modulator 1
<i>PARP6</i>	ENSG00000137817	1,05E+00	1,04E-03	9,40E-02	poly (ADP-ribose) polymerase family, member 6 (EC:2.4.2.30)
<i>ARHGAP17</i>	ENSG00000140750	1,07E+00	1,06E-03	9,42E-02	Rho GTPase activating protein 17
<i>MARVELD1</i>	ENSG00000155254	-1,06E+00	1,03E-03	9,42E-02	MARVEL domain containing 1
<i>RAP2B</i>	ENSG00000181467	-1,09E+00	1,06E-03	9,42E-02	RAP2B, member of RAS oncogene family
<i>AC027763.2</i>	ENSG00000215067	-1,08E+00	1,07E-03	9,42E-02	NA
<i>HBM</i>	ENSG00000206177	1,15E+00	1,07E-03	9,42E-02	hemoglobin, mu
<i>SERHL2</i>	ENSG00000183569	1,11E+00	1,08E-03	9,42E-02	serine hydrolase-like 2
<i>ALG12</i>	ENSG00000182858	-1,06E+00	1,11E-03	9,66E-02	asparagine-linked glycosylation 12, alpha-1,6-mannosyltransferase
<i>PLCG1</i>	ENSG00000124181	1,08E+00	1,12E-03	9,66E-02	phospholipase C, gamma 1 (EC:3.1.4.11)
<i>MAP3K9</i>	ENSG0000006432	1,06E+00	1,13E-03	9,66E-02	mitogen-activated protein kinase kinase kinase 9 (EC:2.7.11.25)
<i>CLIC5</i>	ENSG0000012782	1,09E+00	1,14E-03	9,72E-02	chloride intracellular channel 5
<i>RASGEF1A</i>	ENSG00000198915	1,06E+00	1,15E-03	9,75E-02	RasGEF domain family, member 1A
<i>CDK20</i>	ENSG00000156345	1,08E+00	1,18E-03	9,95E-02	cyclin-dependent kinase 20 (EC:2.7.11.22)

Table S2: Differential gene expression in PBMCs between the sex and age- matched groups was identified by applying a linear mixed model. The model was adjusted for the classical cardiovascular risk factors hypertension, smoking status, body mass index, diabetes and low-density lipoprotein/high-density lipoprotein ratio n=112, FC=fold change, MI=myocardial infarction

Table S9: Significantly differentially expressed genes between *Gpr15*^{gfp/gfp} and wildtype mice in the myocardial infarction zone.

gene	Ensembl_id	log2FC	p-value	gene description
<i>Muc16</i>	ENSMUSG00000109564	3.47E+00	1.65E-04	mucin 16
<i>Cxcl14</i>	ENSMUSG0000021508	1.03E+00	6.49E-04	chemokine (C-X-C motif) ligand 14
<i>Mertk</i>	ENSMUSG0000014361	6.51E-01	8.18E-04	c-mer proto-oncogene tyrosine kinase
<i>Abcb4</i>	ENSMUSG0000042476	-1.48E+00	5.27E-06	ATP-binding cassette, sub-family B (MDR/TAP), member 4
<i>Fam162a</i>	ENSMUSG0000003955	-6.43E-01	5.43E-06	family with sequence similarity 162, member A
<i>E230013L22Rik</i>	ENSMUSG00000096957	-1.59E+00	4.31E-05	RIKEN cDNA E230013L22 gene
<i>Slc16a1</i>	ENSMUSG00000032902	-1.09E+00	4.43E-05	solute carrier family 16 (monocarboxylic acid transporters), member 1
<i>4631405J19Rik</i>	ENSMUSG0000075027	-1.74E+00	4.91E-05	RIKEN cDNA 4631405J19 gene
<i>Kcnj12</i>	ENSMUSG0000042529	-1.55E+00	7.08E-05	potassium inwardly-rectifying channel, subfamily J, member 12
<i>Cap2</i>	ENSMUSG0000021373	-8.27E-01	8.05E-05	CAP, adenylate cyclase-associated protein, 2 (yeast)
<i>Ndufa6</i>	ENSMUSG0000050323	-8.96E-01	9.27E-05	NADH dehydrogenase (ubiquinone) complex I, assembly factor 6
<i>Fgf13</i>	ENSMUSG0000031137	-1.20E+00	9.58E-05	fibroblast growth factor 13
<i>Krt22</i>	ENSMUSG0000035849	-1.71E+00	9.72E-05	keratin 22
<i>Ldhd</i>	ENSMUSG0000031958	-1.11E+00	1.16E-04	lactate dehydrogenase D
<i>Cacnb2</i>	ENSMUSG0000057914	-8.32E-01	1.17E-04	calcium channel, voltage-dependent, beta 2 subunit
<i>Slc25a33</i>	ENSMUSG0000028982	-9.66E-01	1.26E-04	solute carrier family 25, member 33
<i>Tmem74b</i>	ENSMUSG0000044364	-2.62E+00	1.31E-04	transmembrane protein 74B
<i>Gm6123</i>	ENSMUSG00000103922	-1.92E+00	1.33E-04	predicted gene 6123
<i>Pip5k1b</i>	ENSMUSG0000024867	-1.07E+00	1.37E-04	phosphatidylinositol-4-phosphate 5-kinase, type 1 beta
<i>Akap1</i>	ENSMUSG0000018428	-8.07E-01	1.43E-04	A kinase (PRKA) anchor protein 1
<i>Gm17546</i>	ENSMUSG0000078648	-3.21E+00	1.55E-04	predicted gene_17546
<i>Lgr6</i>	ENSMUSG0000042793	-1.31E+00	1.61E-04	leucine-rich repeat-containing G protein-coupled receptor 6
<i>Gm11716</i>	ENSMUSG0000086298	-1.12E+00	1.64E-04	predicted gene_11716
<i>Acss2</i>	ENSMUSG0000027605	-8.16E-01	1.66E-04	acyl-CoA synthetase short-chain family member 2
<i>Cdk2</i>	ENSMUSG0000029403	-9.42E-01	1.88E-04	cyclin-dependent kinase-like 2 (CDC2-related kinase)
<i>0610040B10Rik</i>	ENSMUSG0000089889	-1.05E+00	1.98E-04	RIKEN cDNA 0610040B10 gene
<i>Ube2ql1</i>	ENSMUSG0000052981	-1.21E+00	2.04E-04	ubiquitin-conjugating enzyme E2Q family-like 1
<i>Grb14</i>	ENSMUSG0000026888	-9.15E-01	2.04E-04	growth factor receptor bound protein 14
<i>Kbtbd12</i>	ENSMUSG000003182	-1.23E+00	2.14E-04	kelch repeat and BTB (POZ) domain containing 12
<i>Fhl2</i>	ENSMUSG000008136	-1.16E+00	2.25E-04	four and a half LIM domains 2
<i>Pygo1</i>	ENSMUSG0000034910	-8.24E-01	2.39E-04	pygopus 1
<i>Atp5s</i>	ENSMUSG0000054894	-7.28E-01	2.69E-04	ATP synthase, H+ transporting, mitochondrial F0 complex, subunit S
<i>Bves</i>	ENSMUSG0000071317	-1.07E+00	2.84E-04	blood vessel epicardial substance
<i>Cox7c</i>	ENSMUSG0000017778	-6.95E-01	3.05E-04	cytochrome c oxidase subunit VIIc
<i>Smpx</i>	ENSMUSG0000041476	-8.88E-01	3.08E-04	small muscle protein, X-linked
<i>Tfrc</i>	ENSMUSG0000022797	-1.28E+00	3.17E-04	transferrin receptor
<i>Ppp1r3d</i>	ENSMUSG0000049999	-8.52E-01	3.19E-04	protein phosphatase 1, regulatory subunit 3D
<i>Rab3ip</i>	ENSMUSG0000064181	-6.68E-01	3.19E-04	RAB3A interacting protein
<i>Klhdc1</i>	ENSMUSG0000051890	-8.62E-01	3.22E-04	kelch domain containing 1
<i>Gm16399</i>	ENSMUSG0000097790	-7.88E-01	3.37E-04	predicted pseudogene_16399
<i>Tmem246</i>	ENSMUSG0000039611	-9.89E-01	3.67E-04	transmembrane protein 246
<i>Esrrib</i>	ENSMUSG0000021255	-1.25E+00	4.13E-04	estrogen related receptor, beta
<i>Mfn1</i>	ENSMUSG0000027668	-8.03E-01	4.13E-04	mitofusin 1
<i>Cox6c</i>	ENSMUSG0000014313	-6.16E-01	4.18E-04	cytochrome c oxidase subunit VIc
<i>Halr1</i>	ENSMUSG0000085412	-3.17E+00	4.61E-04	Hoxa adjacent long noncoding RNA 1
<i>Timp4</i>	ENSMUSG0000030317	-2.10E+00	5.04E-04	tissue inhibitor of metalloproteinase 4
<i>Armc2</i>	ENSMUSG0000071324	-1.18E+00	5.10E-04	armadillo repeat containing 2
<i>Grip2</i>	ENSMUSG0000030098	-1.62E+00	5.23E-04	glutamate receptor interacting protein 2
<i>Acat1</i>	ENSMUSG0000032047	-7.39E-01	5.31E-04	acetyl-Coenzyme A acetyltransferase 1
<i>Frmd5</i>	ENSMUSG0000027238	-1.17E+00	5.56E-04	FERM domain containing 5
<i>Gm43032</i>	ENSMUSG00000106498	-3.01E+00	5.72E-04	predicted gene_43032
<i>D3Ertd751e</i>	ENSMUSG0000025766	-9.20E-01	5.79E-04	DNA segment, Chr 3, ERATO Doi 751, expressed
<i>Egln1</i>	ENSMUSG0000031987	-7.02E-01	5.79E-04	egl-9 family hypoxia-inducible factor 1
<i>Usp28</i>	ENSMUSG0000032267	-1.05E+00	6.16E-04	ubiquitin specific peptidase 28
<i>Mipep-ps</i>	ENSMUSG0000093624	-2.75E+00	6.21E-04	mitochondrial intermediate peptidase, pseudogene
<i>D83039M14Rik</i>	ENSMUSG0000043126	-1.79E+00	6.26E-04	RIKEN cDNA D83039M14 gene
<i>Arhgef9</i>	ENSMUSG0000025656	-9.91E-01	6.37E-04	CDC42 guanine nucleotide exchange factor (GEF) 9
<i>Asb11</i>	ENSMUSG0000031382	-1.48E+00	6.56E-04	ankyrin repeat and SOCS box-containing 11
<i>CU074400.1</i>	ENSMUSG00000114254	-9.36E-01	6.60E-04	NA
<i>Gm16793</i>	ENSMUSG0000097357	-2.37E+00	6.63E-04	predicted gene_16793
<i>Gm6238</i>	ENSMUSG0000082996	-1.58E+00	6.71E-04	predicted pseudogene_6238
<i>Cycs</i>	ENSMUSG0000063694	-7.64E-01	6.76E-04	cytochrome c, somatic
<i>Pdk1</i>	ENSMUSG0000006494	-5.92E-01	6.94E-04	pyruvate dehydrogenase kinase, isoenzyme 1
<i>2210408F21Rik</i>	ENSMUSG0000087380	-7.06E-01	7.03E-04	RIKEN cDNA 2210408F21 gene
<i>Ndufb2</i>	ENSMUSG0000024216	-6.10E-01	7.26E-04	NADH dehydrogenase (ubiquinone) 1 beta subcomplex, 2
<i>Gm10053</i>	ENSMUSG0000058927	-7.19E-01	7.43E-04	predicted gene_10053
<i>Eml6</i>	ENSMUSG0000044072	-1.38E+00	7.44E-04	echinoderm microtubule associated protein like 6
<i>Nlrp10</i>	ENSMUSG0000049709	-1.08E+00	7.46E-04	NLR family, pyrin domain containing 10
<i>Slc25a4</i>	ENSMUSG0000031633	-9.71E-01	7.51E-04	solute carrier family 25 (mitochondrial carrier, adenine nucleotide translocator), member 4
<i>Ppif</i>	ENSMUSG0000021868	-1.06E+00	7.94E-04	peptidylprolyl isomerase F (cyclophilin F)
<i>Gm26672</i>	ENSMUSG0000097330	-2.20E+00	8.05E-04	predicted gene_26672
<i>Kcnd2</i>	ENSMUSG0000060882	-1.79E+00	8.05E-04	potassium voltage-gated channel, Shal-related family, member 2
<i>Gm17021</i>	ENSMUSG0000091661	-1.40E+00	8.21E-04	predicted gene_17021
<i>Pacsin3</i>	ENSMUSG0000027257	-7.54E-01	8.45E-04	protein kinase C and casein kinase substrate in neurons 3
<i>6430571L13Rik</i>	ENSMUSG0000037977	-1.89E+00	8.60E-04	RIKEN cDNA 6430571L13 gene
<i>Scn5a</i>	ENSMUSG0000032511	-7.42E-01	8.71E-04	sodium channel, voltage-gated, type V, alpha
<i>Dld</i>	ENSMUSG0000020664	-6.52E-01	8.81E-04	dihydrolipoamide dehydrogenase
<i>Lppos</i>	ENSMUSG00000097867	-6.12E-01	8.86E-04	LIM domain containing preferred translocation partner in lipoma, opposite strand
<i>Mterf2</i>	ENSMUSG0000049038	-9.85E-01	9.28E-04	mitochondrial transcription termination factor 2
<i>Cavin4</i>	ENSMUSG0000028348	-1.32E+00	9.44E-04	caveolae associated 4
<i>Mdh1</i>	ENSMUSG0000020321	-8.58E-01	9.52E-04	malate dehydrogenase 1, NAD (soluble)
<i>Trp53inp2</i>	ENSMUSG0000038375	-6.80E-01	9.65E-04	transformation related protein 53 inducible nuclear protein 2
<i>Acyp2</i>	ENSMUSG0000060923	-7.28E-01	9.78E-04	acylphosphatase 2, muscle type
<i>Neb1</i>	ENSMUSG0000053702	-8.90E-01	9.82E-04	nebulette

Table S9: Differential cardiac gene expression between *Gpr15*^{gfp/gfp} and wildtype mice was calculated using DESeq2. n=8/9, FC=fold change