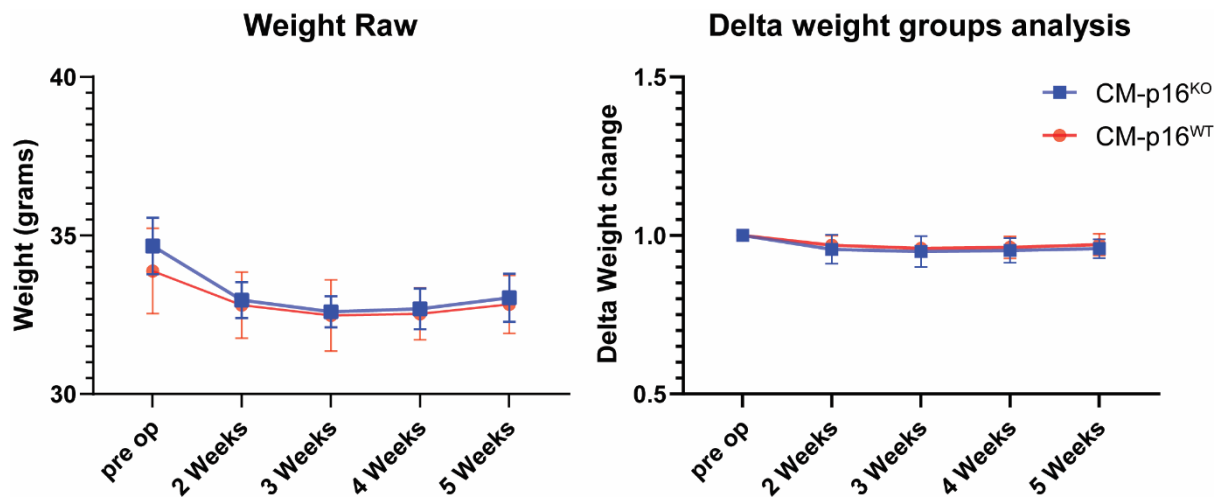
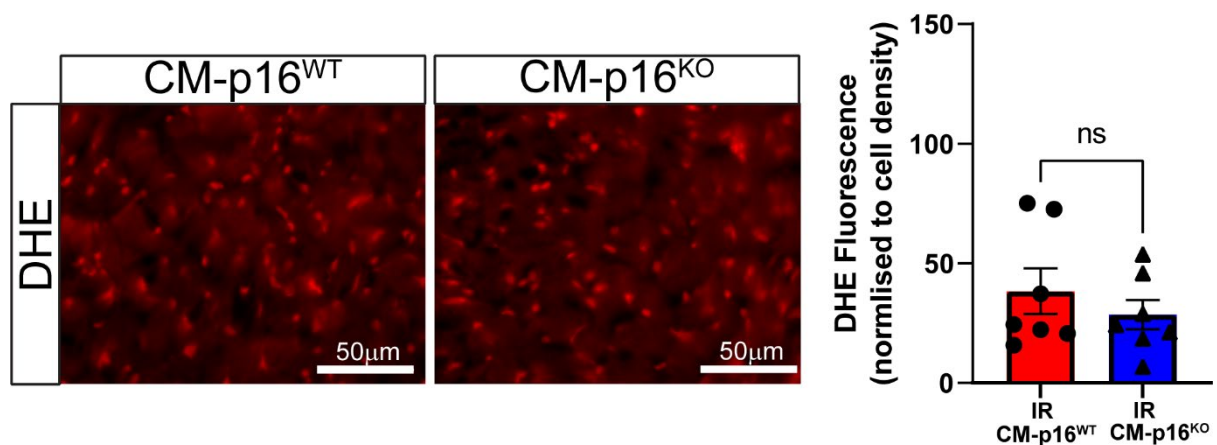
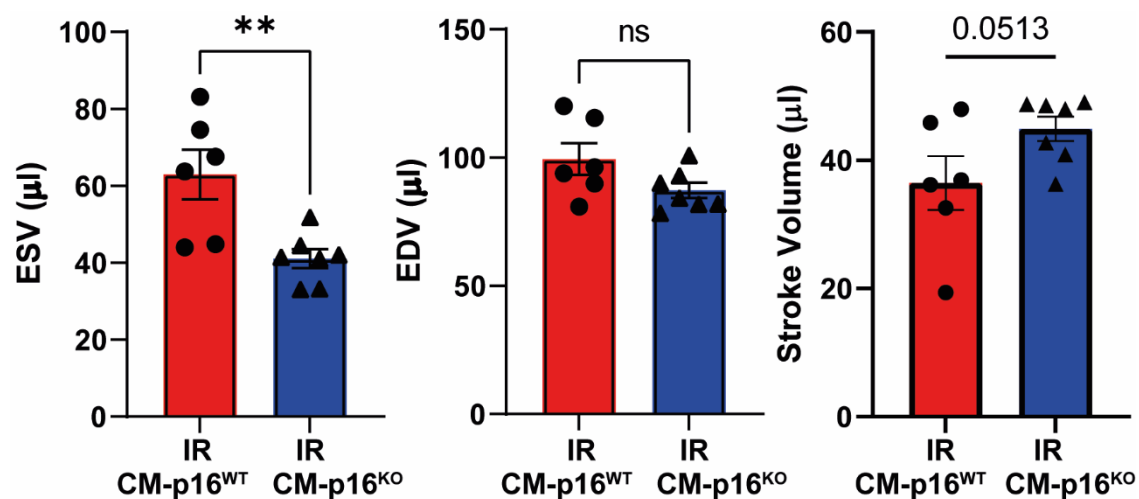




Supplementary Figure 1. **Body weight and delta change in body weight for all animals for the duration of the study.** No significant difference was observed in weights pre or post LAD-ligation between groups. No difference in weight change was observed between groups. Analysis performed by multiple student T-Test.



Supplementary Figure 2. **Superoxide levels post ischemia reperfusion.** Representative image of Dihydroethidium (DHE) staining at 5-weeks post LAD-Ligation. Quantification of DHE fluorescence intensity in the peri-infarct myocardium of CM-p16KO and CM-p16WT mice. n=7/group. All scale bars as indicated. Data are mean±SEM, analysis by Student T-Test.

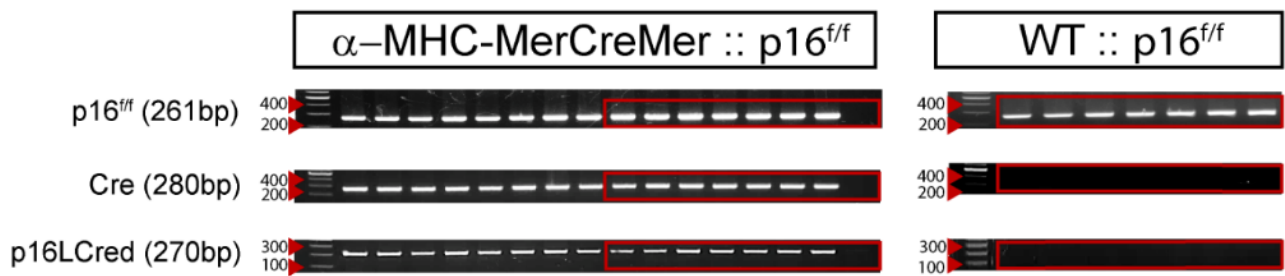


Supplementary Figure 3. **Magnetic resoance image analysis.** End diastolic volume (ESV), End systolic volumes (EDV) and stroke volume 5 weeks post-MI, n>6/group. Data are mean±SEM, **P<0.01. Using Mann-Whitney or student T-Test as appropriate.

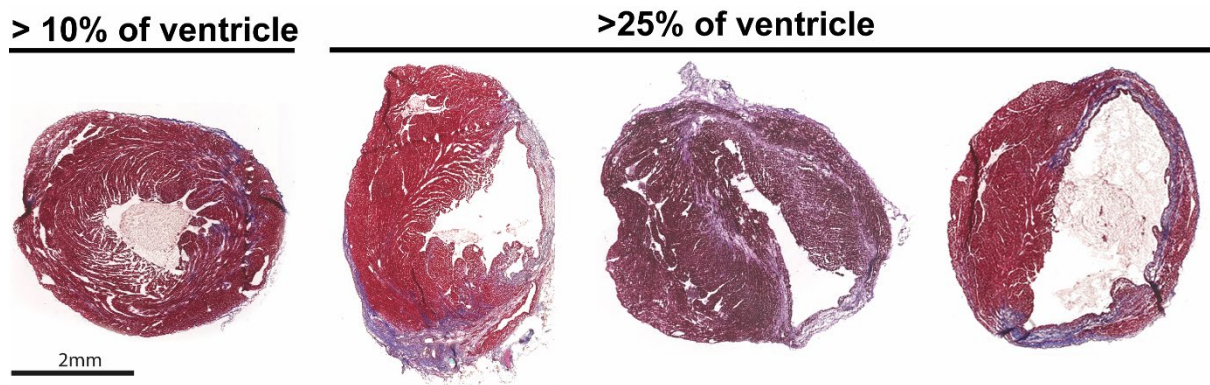
Sequences for primers used for genotyping polymerase chain reactions.

Name	Primer Sequence
p16 ^{Ink4a} f/f F	GTATGCTATACGAAGTTATTAGGTACTGC
p16 ^{Ink4a} wild-type F	GTTTTGGAGCAGCAGGGATT
p16 ^{Ink4a} common R	CTATGTCAGATTTGGCTAGGGAGT
S1X-A Cre F	TAACCAGTGAAACAGCATTGCTG
S1X-B Cre R	GGACATGTTTCAGGGATCGCCAGGCG
p16 ^{Ink4a} LCred F	TACCACAGTTTGAACAGCGTGA
p16 ^{Ink4a} LCred R	AACCAACTTCCTCCTTCCCC

Supplementary Figure 4. **Primer sequences used in the study.**



Supplementary Figure 5. **Confirmation of the excision of p16 exon 1α.** Mouse genotypes were established using primers specific to the floxed p16^{inka4} allele (p16^{f/f}) and αMHC-MerCreMer (cre). Bitransgenic MerCreMer-p16^{f/f} mice and WT-p16^{f/f} control mice were given 4-OH-tamoxifen for 14 days. Cre-mediated excision of exon 1α from the MerCreMer :: p16^{f/f} mice was confirmed by PCR using p16LCre primers which span the excised region. Red outline identifies animals used in this study.



Supplementary Figure 6. **Hearts excluded from study due to incomplete ligation or reperfusion.** Scar size under 10% area of left ventricle indicative of unsuccessful Ligation. Transmural scar which comprises >25% of the left ventricular myocardium indicative of unsuccessful reperfusion.

Cytokine	Concentration (pg/ml $\pm$ SD)		p Value
	IR CM-p16 ^{WT}	IR CM-p16 ^{KO}	
Eotaxin	0.06 $\pm$ 0.10	4.2 $\pm$ 7.00	0.18
G-CSF	0.59 $\pm$ 0.54	0.07 $\pm$ 0.98	0.07
GM-CSF	OOOR	OOOR	-
IFN γ	14.20 $\pm$ 4.2	6.90 $\pm$ 6.023	0.08
IL-1 α	161.2 $\pm$ 54.55	176.9 $\pm$ 99.87	0.41
IL-1 β	2.78 $\pm$ 2.44	2.713 $\pm$ 3.39	0.49
IL-2	22.30 $\pm$ 13.57	20.12 $\pm$ 12.08	0.42
IL-3	0.413 $\pm$ 0.7	0.05 $\pm$ 0.08	0.22
IL-4	0.05 $\pm$ 0.03	0.07 $\pm$ 0.12	0.41
IL-5	0.026 $\pm$ 0.05	OOOR	-
IL-6	2.93 $\pm$ 0.92	1.477 $\pm$ 0.49	0.04 (*)
IL-7	1.75 $\pm$ 0.91	1.047 $\pm$ 0.90	0.20
IL-9	72.10 $\pm$ 19.30	79.36 $\pm$ 11.88	0.30
IL-10	31.56 $\pm$ 26.88	33.35 $\pm$ 20.28	0.47
IL-11	2.21 $\pm$ 0.84	1.74 $\pm$ 0.46	0.22
IL-12/p40	39.94 $\pm$ 27.27	30.91 $\pm$ 11.00	0.31
IL-12/p70	1.66 $\pm$ 0.675	0.85 $\pm$ 1.472	0.22
IL-13	6.287 $\pm$ 2.01	5.09 $\pm$ 0.94	0.20
IL-15	12.86 $\pm$ 17.49	24.23 $\pm$ 12.18	0.36
IL-16	151.9 $\pm$ 123.2	138.7 $\pm$ 158.9	0.46
IL-17	28.84 $\pm$ 17.49	24.23 $\pm$ 13.18	0.37
IL-20	OOOR	OOOR	-
IP-10	3.53 $\pm$ 2.36	3.54 $\pm$ 1.69	0.50
KC	25.16 $\pm$ 15.95	21.32 $\pm$ 6.58	0.34
LIF	0.12 $\pm$ 0.19	0.15 $\pm$ 0.25	0.45
LIX	2753 $\pm$ 478.5	2822 $\pm$ 1056	0.46
MCP-1	4.107 $\pm$ 4.63	2.93 $\pm$ 2.80	0.36
MCP-5	5.53 $\pm$ 4.252	12.36 $\pm$ 8.59	0.12
M-CSF	5.063 $\pm$ 2.55	2.55 $\pm$ 2.011	0.12
MIG	1.783 $\pm$ 0.95	2.50 $\pm$ 1.274	0.24
MIP-1a	3.34 $\pm$ 3.47	3.1 $\pm$ 5.38	0.48
MIP-1 β	OOOR	OOOR	-
MIP-2	29.76 $\pm$ 14.09	25.62 $\pm$ 19.79	0.39
MIP-3 α	3.82 $\pm$ 1.34	4.45 $\pm$ 0.86	0.26
MIP-3 β	0.74 $\pm$ 1.29	OOOR	-
RANTES	2.183 $\pm$ 0.41	1.84 $\pm$ 0.86	0.28
TARC	0.46 $\pm$ 1.89	1.89 $\pm$ 2.26	0.17
TIMP-1	OOOR	OOOR	-
TNF α	1.35 $\pm$ 0.82	0.89 $\pm$ 0.96	0.29
VEGF	3.9 $\pm$ 3.58	4.9 $\pm$ 6.06	0.41
6CKine/Exodus2	306.20 $\pm$ 194.3	510.86 $\pm$ 199.2	0.13
EPO	OOOR	OOOR	-
Fractalkine	12.82 $\pm$ 3.81	12.93 $\pm$ 3.09	0.49
MCD	0.77 $\pm$ 0.24	0.75 $\pm$ 0.074	0.45
IFNB-1	62.12 $\pm$ 5.62	60.45 $\pm$ 3.81	0.35

Supplementary Table 1. Values from MD-44 Cytokine array. Below detectable range = OOR (Out of Range). Data from n=3 per experimental group. Following tests of normality, data analysed by Student's t-test.