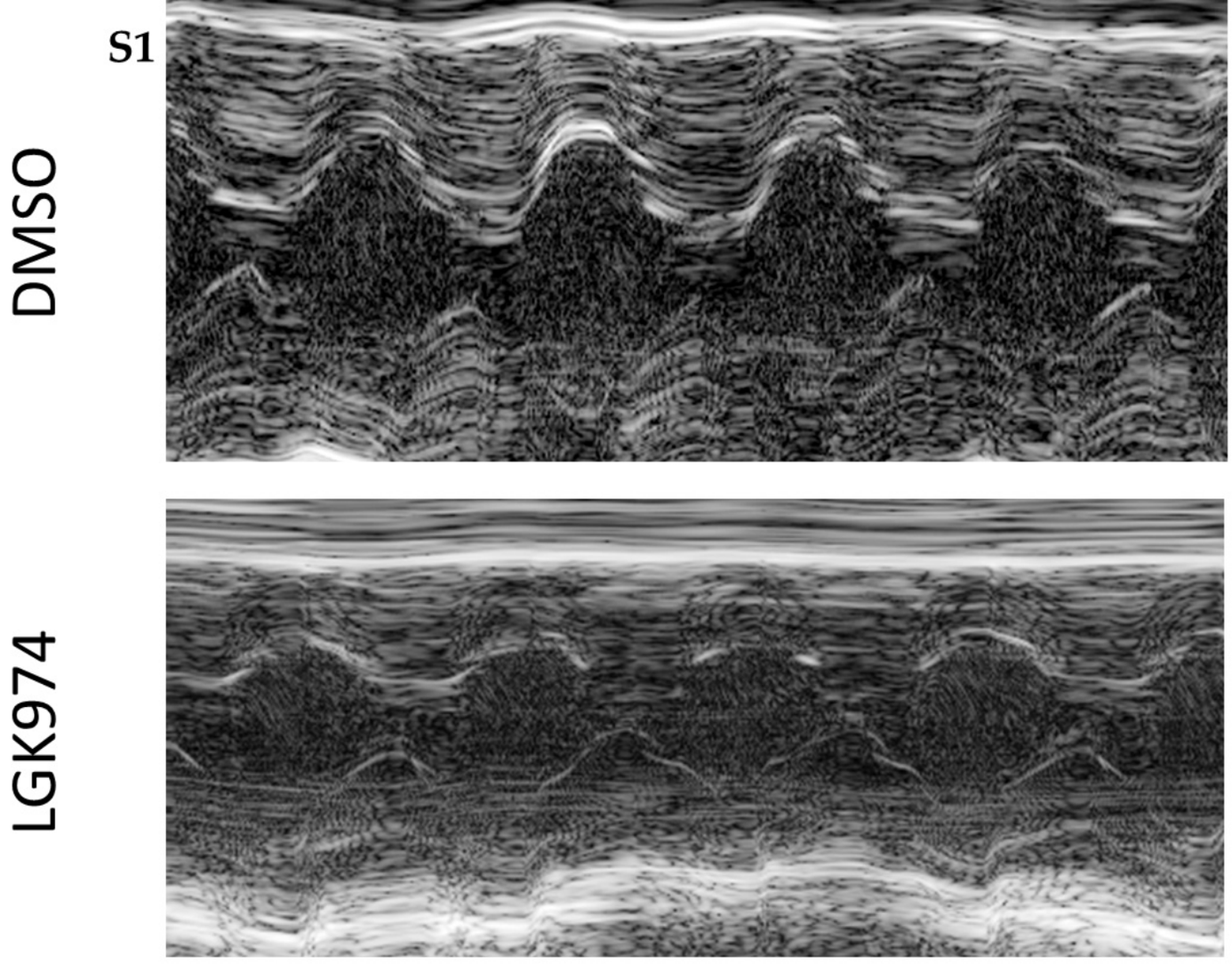
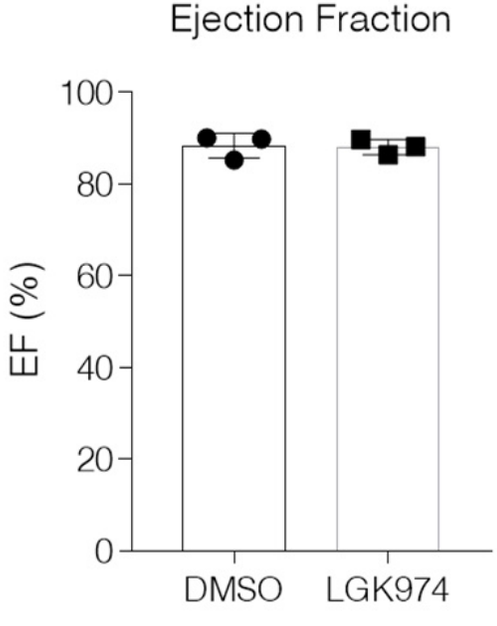
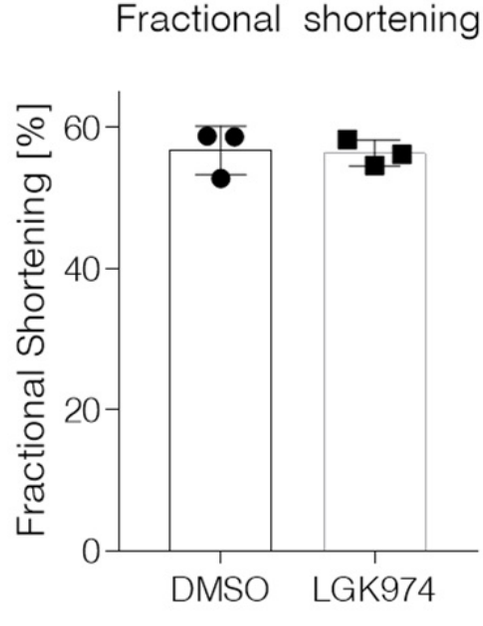
S1


DMSO
LGK974

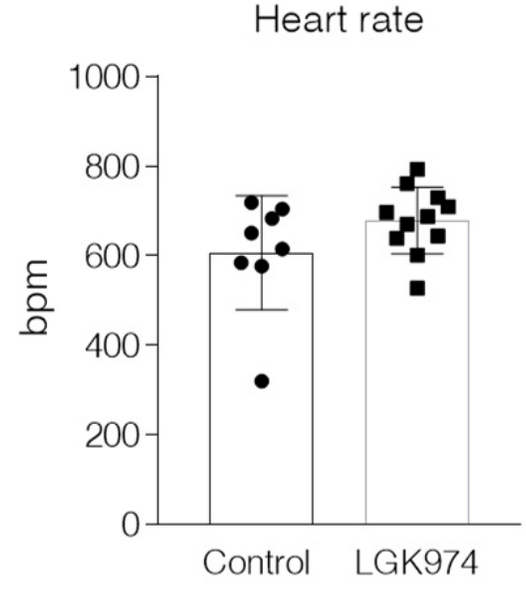
S2


Ejection Fraction
EF (%)
100
80
60
40
20
0
DMSO
LGK974

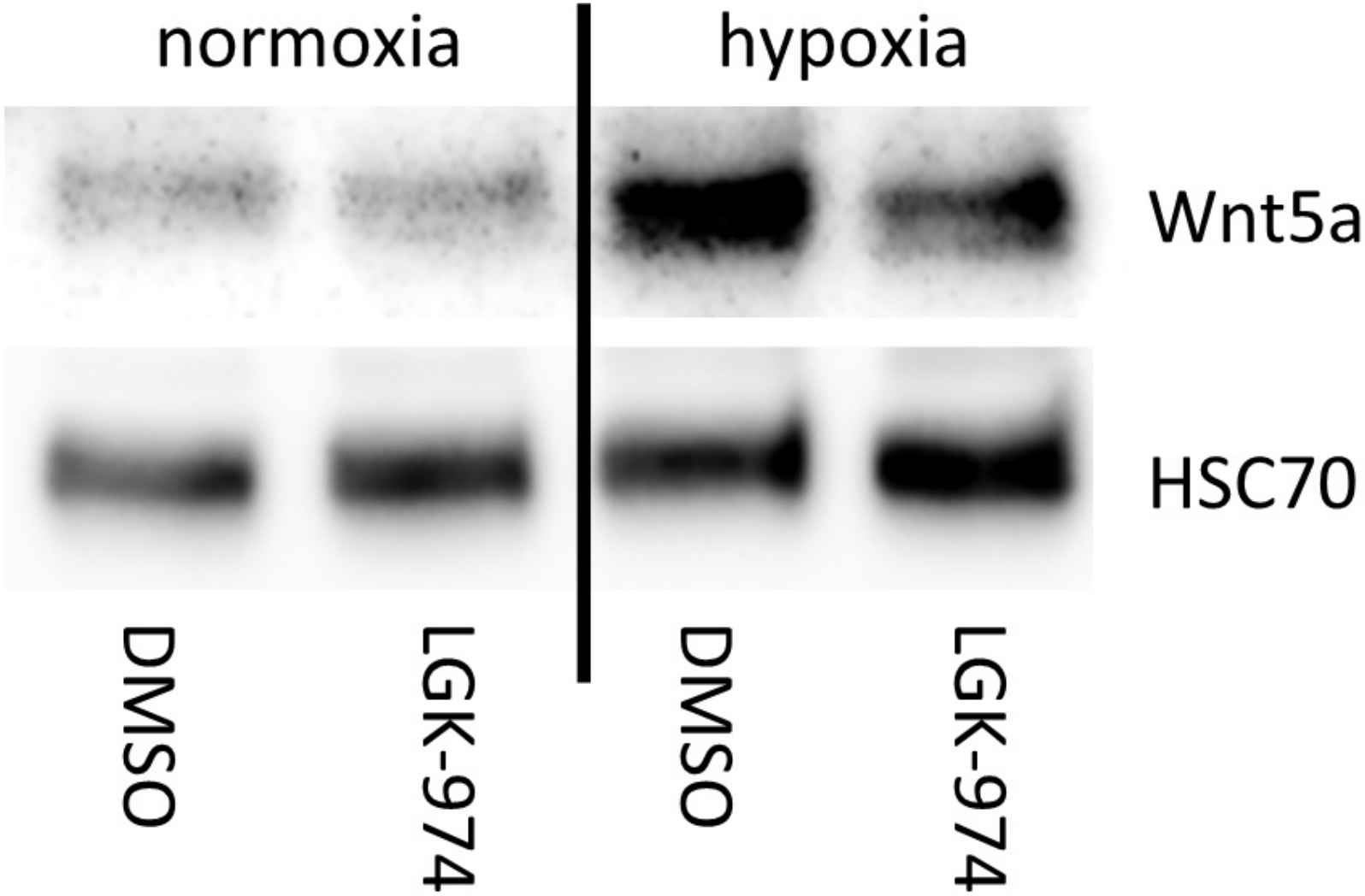
S3


Fractional shortening
Fractional Shortening [%]
60
40
20
0
DMSO
LGK974

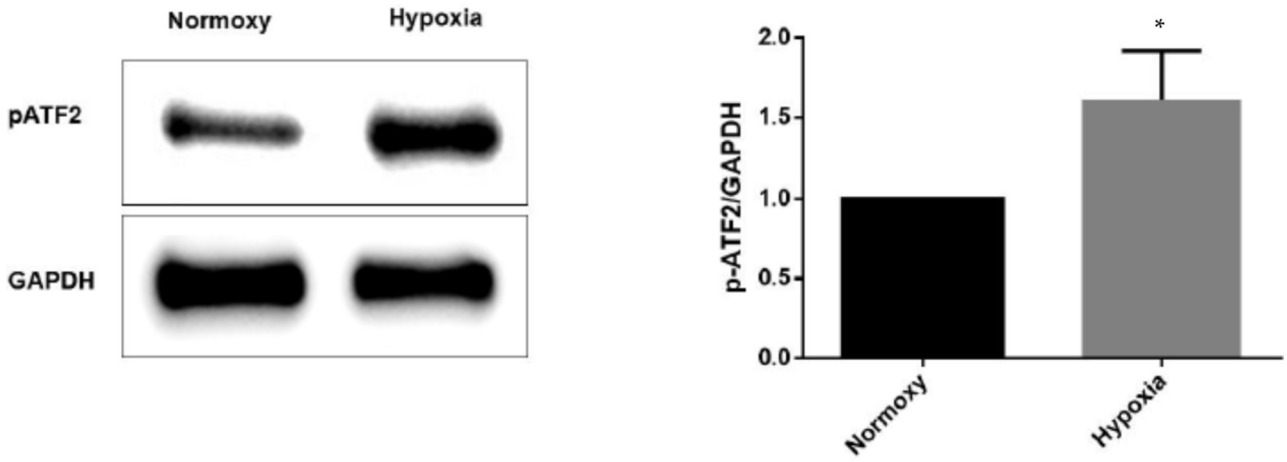
S4


Heart rate
bpm
1000
800
600
400
200
0
Control
LGK974

S5



S6


Normoxy
Hypoxia
pATF2
GAPDH
p-ATF2/GAPDH
2.0
1.5
1.0
0.5
0.0
*
Normoxy
Hypoxia

Supplemental figure S 1-4: S1: Representative M-Mode echocardiographic images of DMSO (top) and LGK-974 treated (bottom) animals under basal conditions. S2 and S3: Echocardiographic results from DMSO and LGK-974 treated animals under basal conditions. S4: Heart rate of LGK-974 and DMSO treated animals 28 days after I/R injury.

Supplemental figure S5: Western blot of Wnt5A from supernatant of cardiomyocytes cultured under hypoxic or normoxic condition in the presence or absence of LGK-974. HSC70 were used as a secreted control protein. LGK-974 effectively inhibited Wnt5A secretion.

Supplemental figure S 6: Stimulation of RAW264.7 cell with supernatant of hypoxic cardiomyocytes leads to activation of non-canonical Wnt signaling as seen by increased pATF2 expression levels.