Non-transgenic WT hCD59RBC

FigS1: The hCD59 protein was detected in the endothelium of middle-sized intramyocardial vessels in ThCD59RBC but not in the non-transgenic mouse hearts. Immunofluorescence was performed with anti-hCD59 antibody (Bric 229, BITS/IBGRL).









WT ThCD59^{END} ThCD59^{RBC}

FigS2: ILY injection did not result in direct endothelial cell

damage: **(A)** vWF levels after ILY injection in *ThCD59^{RBC}* mice. Data are represented by Mean <u>+</u> SEM **P* < 0.05 vs WT mice (n=4) 12 and 24 hrs after ILY injection. **(B)** Representative pictures from 4 independent mice in each group show the Evan's blue staining in the aorta of the mice. The Evan's blue staining of the aortas from WT and *ThCD59^{RBC}* mice treated with ILY₁₄₀SI or *ThCD59^{END}* mice treated with 4.5 ng/g ILY i.v. injection was performed at 2 hrs post-injection. Extensive Evan's blue staining was detected in the aorta of the *ThCD59^{END}* that express hCD59 in the endothelium, but not in the WT and *ThCD59^{RBC}* mice. Scale bar: 1 mm. **(C)** FACS analysis showed no hCD59 expression in the platelets of *ThCD59^{RBC}* or WT platelets but in human platelets. Red: WT platelets; Green: human platelets; Black: *ThCD59^{RBC}* platelets. The platelets were stained with anti-human CD59 Ab (Bric 229) and FITC-conjugated secondary Abs.



FigS3: Representative images of Pselectin staining in lung. There was more extensive P-selectin staining in the alveolar vessels in the $ThCD59^{RBC}$ with ILY₁₄₀FI treatment than in the $ThCD59^{RBC}$ with ILY₁₄₀SI treatment, or the WT with ILY₁₄₀FI treatment. The 10 X magnification images show the area of 4 X magnification images highlighted in the black squares.



FigS4. Thrombi were specifically formed in the lung, but not in the kidney, liver and heart of $ThCD59^{RBC}$ mice treated with ILY₁₄₀FI. Representative images (magnification: X 40) show Intergrin α IIb stainings in the blood vasculature of lung, kidney, liver and heart of $ThCD59^{RBC}$ mice treated with ILY₁₄₀FI. There were extensive Intergrin α IIb stainings in the thrombi of the blood vessels in the lungs but not the kidneyt, liver and heart of the mice. Arrows point to vasculatures in the tissues.



FigS5: Representative echocardiographic frames show the dynamic changes in the right ventricular volume in the *ThCD59*^{RBC} with platelet depletion (neuraminidase treatment) and inhibition (clopidogrel treatment) followed by ILY₁₄₀FI injection (the videos are shown in **Supplementary movies 4-6**).



FigS6: Increased percentage of maximal right ventricular area (at end-diastole) vs. baseline right ventricular area in the *ThCD59*^{*RBC*} mice treated with clopidogrel and neuraminidase followed by ILY₁₄₀FI. * *P*<0.01 vs. the mice without the pretreatment. The data are shown as mean <u>+</u> SEM

WT with ILY₁₄₀FI



60 X

20 X

FigS7. Representative images of vWF staining in lung. There were more extensive vWF stainings (arrows) in the thrombi of the blood vessels in the lungs of *ThCD59*^{RBC} mice but not WT treated with ÌLY₁₄₀FÍ.