



## Invasive Kühlung beim kardiogenen Schock



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# Indikationen der Hypothermie



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**Herz- und Neurochirurgie** ✓

**Koma nach Reanimation** ✓

**Kardiog. Schock nach Rean.: MH ist nicht schädlich** (✓)

**STEMI: protektiv in einer Pilotstudie** (✓)

**Apoplex: unklar** ?

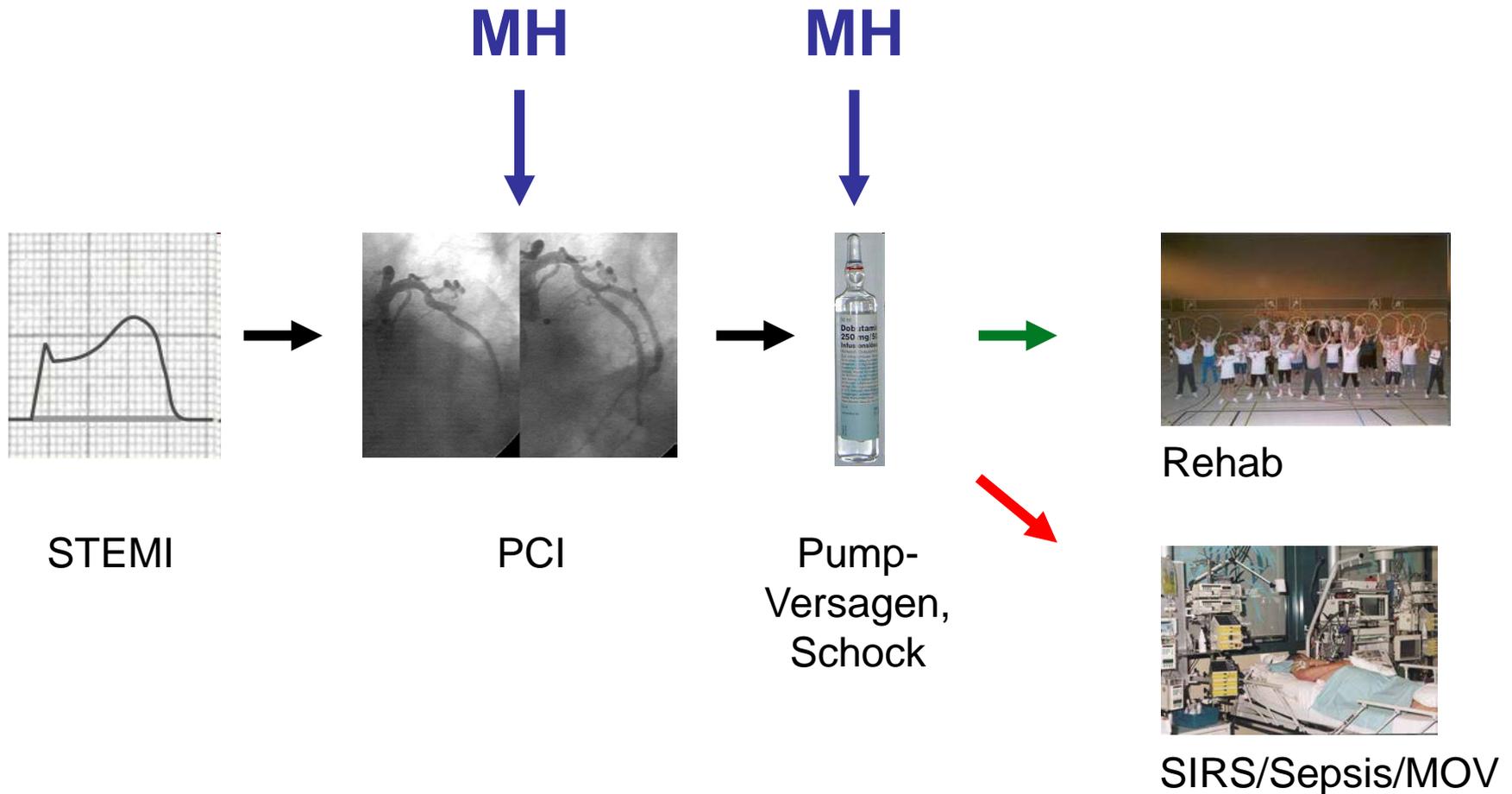
**SHT/ICB: unklar** ?

**Lungenversagen**

# Milde Hypothermie beim kardiogenen Schock: Wo kann sie indiziert sein?



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# Wie groß wird ein Herzinfarkt?



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nicht beeinflussbar:

- Größe des ischämischen Areals
- Restdurchblutung während der Ischämie

beeinflussbar:

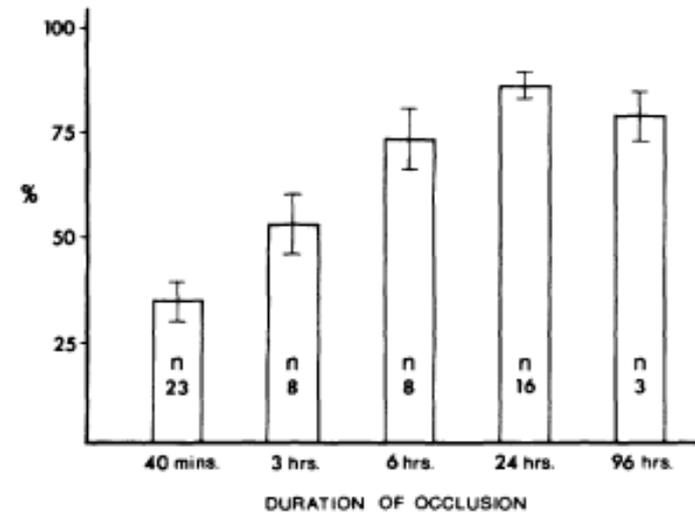
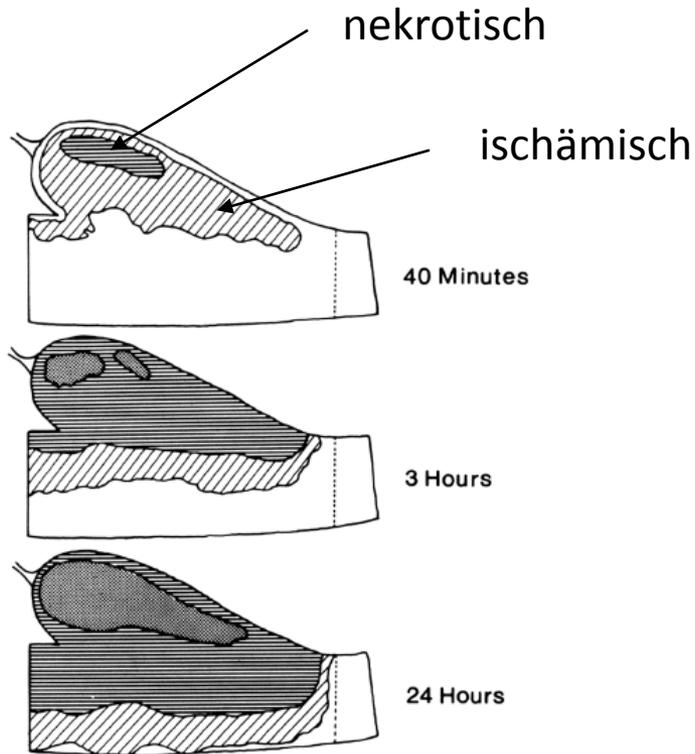
- Dauer der Ischämie
- Sympathotonus/metabolische Rate während Ischämie
- **Modus der Reperfusion**

# Infarktgröße: Dauer der Ischämie

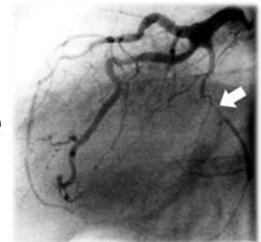


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Infarktgröße



Aspisol  
Efient/Brilique  
Heparine

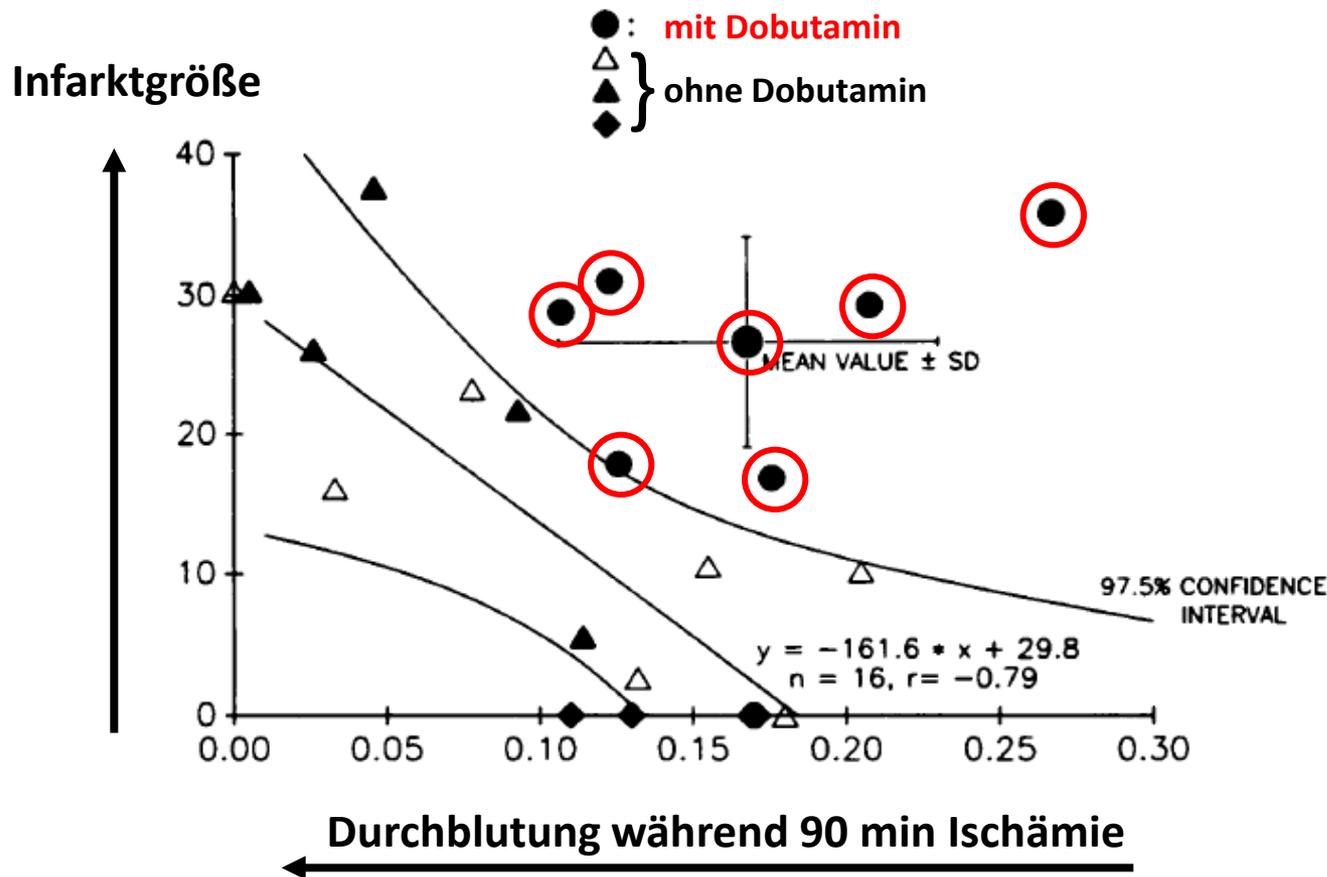


Reimer et al. , Circulation 1977

# Infarktgröße: Durchblutung und Katecholamine



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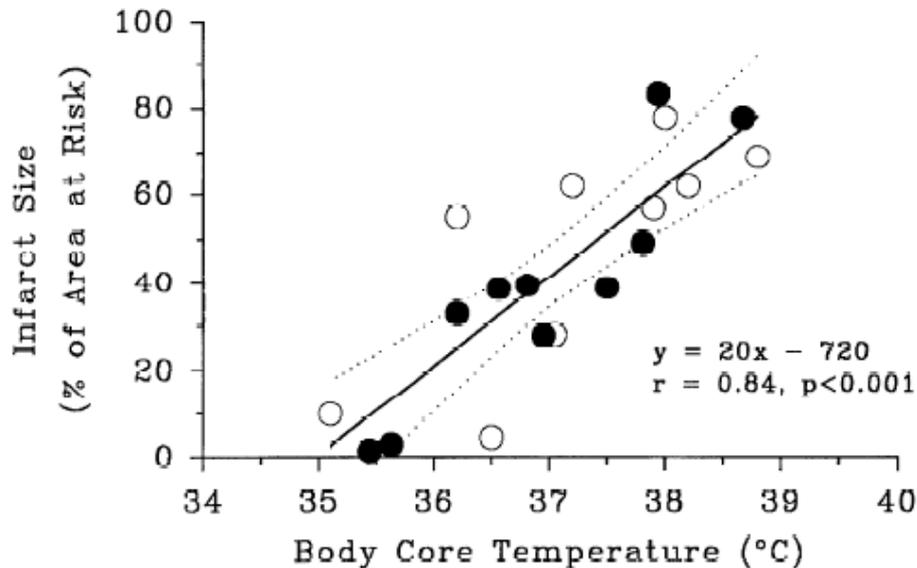
Schulz et al., Circulation 1993

# Hypothermie senkt die Infarktgröße: Es funktioniert im Experiment

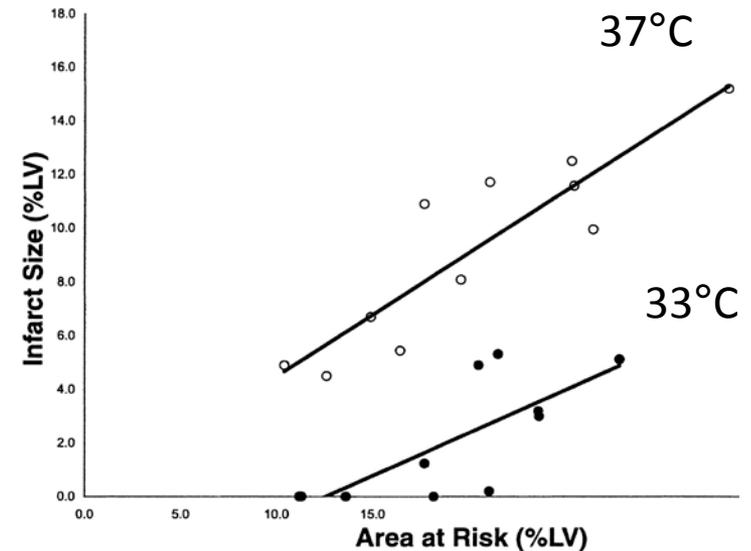


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Schwein in vivo, Kühlung während der Ischämie



Duncker et al., AJP 1996



Dae et al., AJP 2002

# Hypothermie bei Patienten: Die falsche Studie - Das Ende des Hype



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## Hypothermia in the prevention of MI – COOL-MI

### COOL-MI Trial (Radiant)

No difference in Day 30 SPECT infarct size (1<sup>o</sup> endpoint)

Control Arm (13.8%) vs. Hypothermia Arm (14.1%),  $p = 0.83$

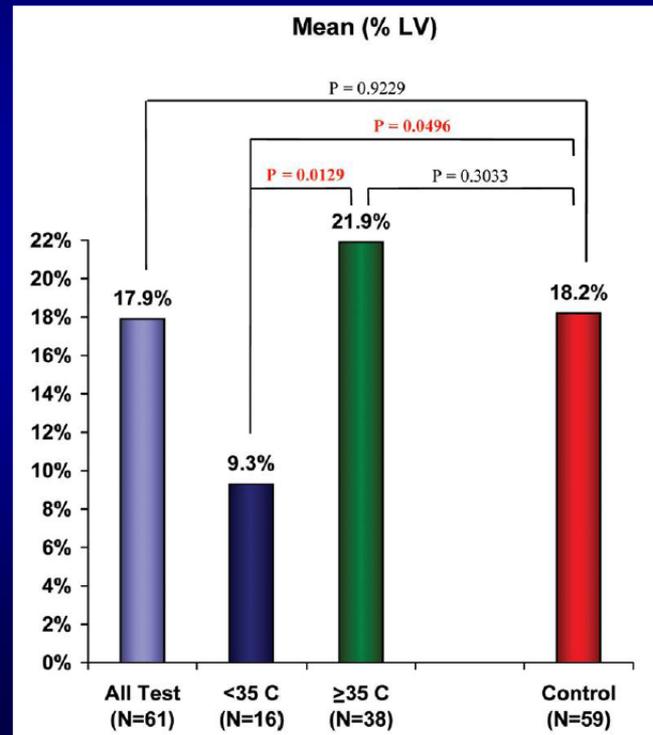
Cooling deemed safe and well-tolerated (94% tolerated hypothermia).

Possible “dose-response” relationship in anterior MI subgroup

Anterior MIs –

Day 30 SPECT Infarct Size

49% reduction in infarction size in precooled patients

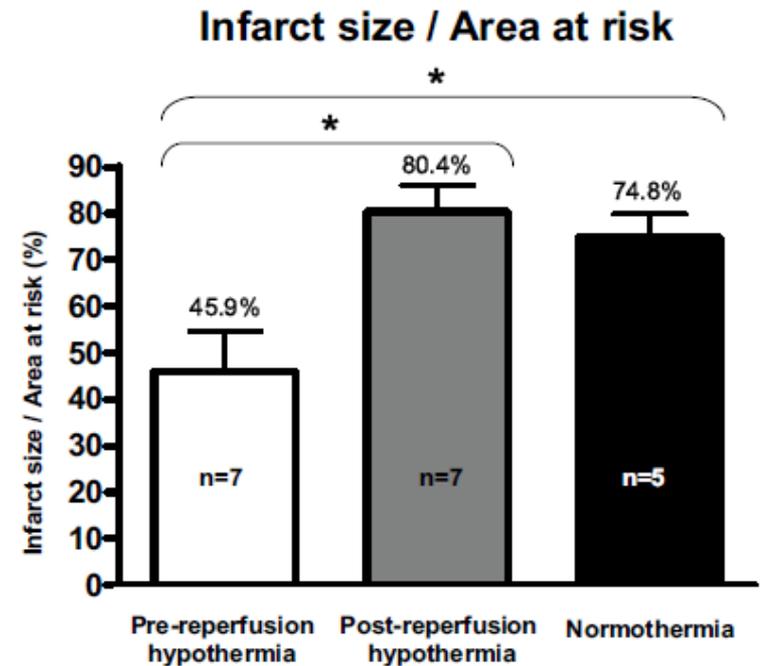
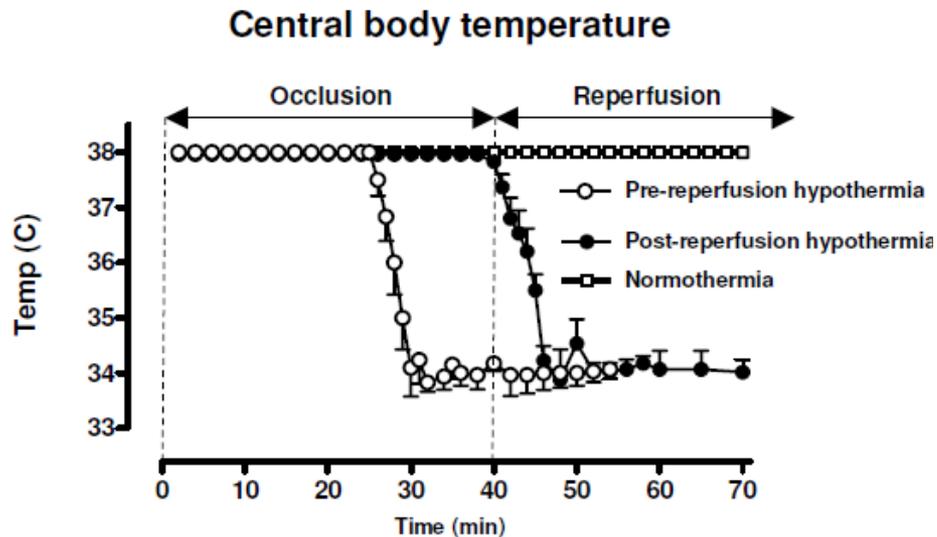


# MH vor vs nach der Reperfusion



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Schwein in vivo, Kühlung vor Reperfusion



Götberg, BMC 2008

# Schneller kühlen!

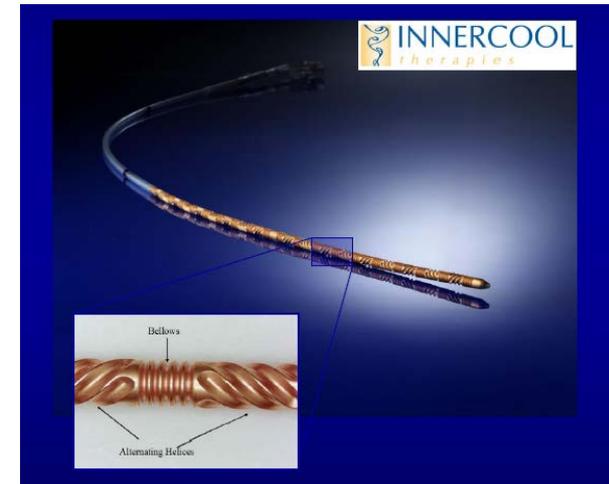


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**1000 ml cold saline (4°C)  
Quick initiation of hypothermia**

**1-2 Liter eiskalte  
Lösung**

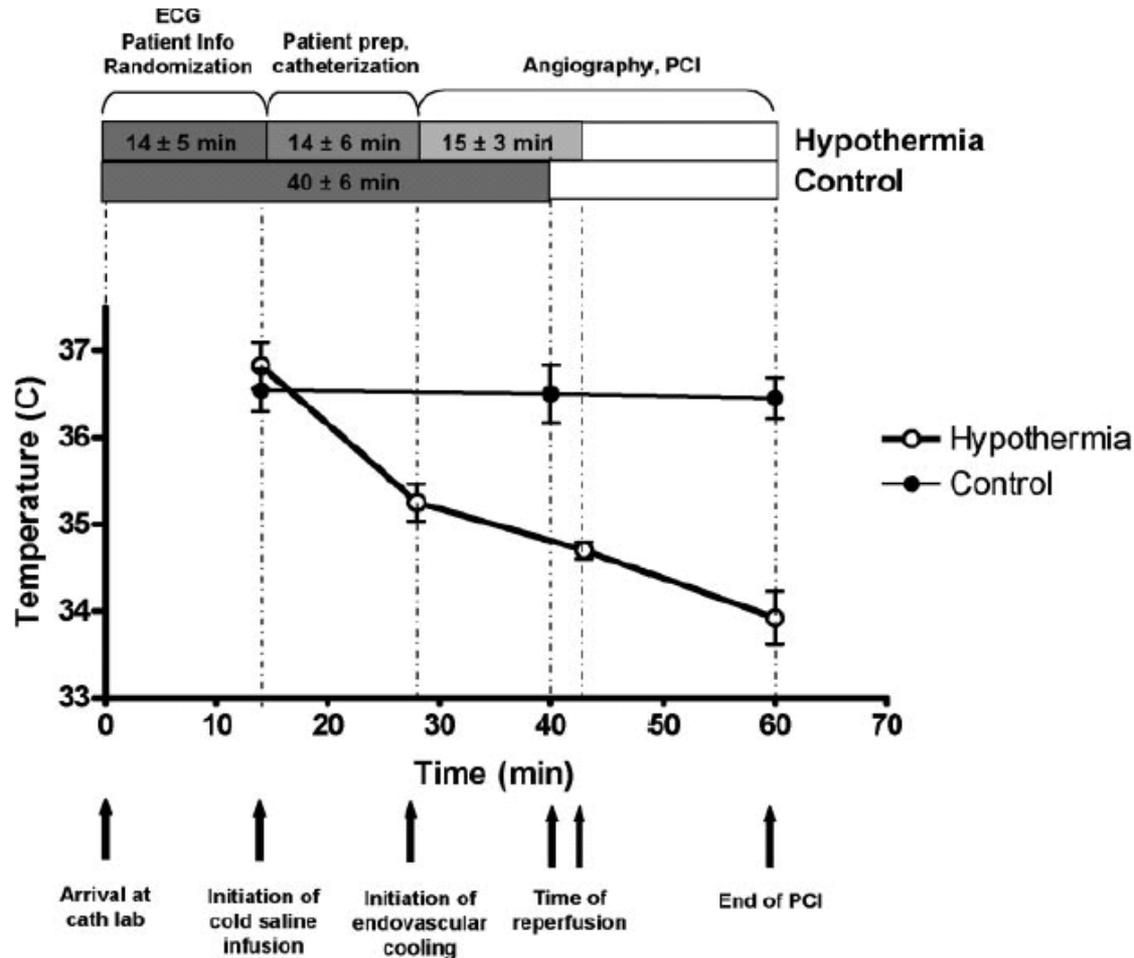


**intravaskuläres  
Kühlen**

# Induktion einer MH $< 35^{\circ}\text{C}$ bei Patienten mit STEMI



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Götberg, Circ CI 2010

# MH bei Patienten mit STEMI



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## time to reperfusion

## area-at-risk and infarct size (MRT)

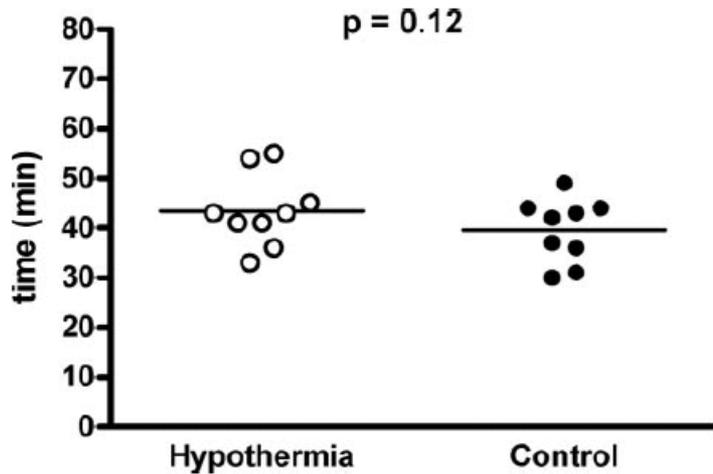
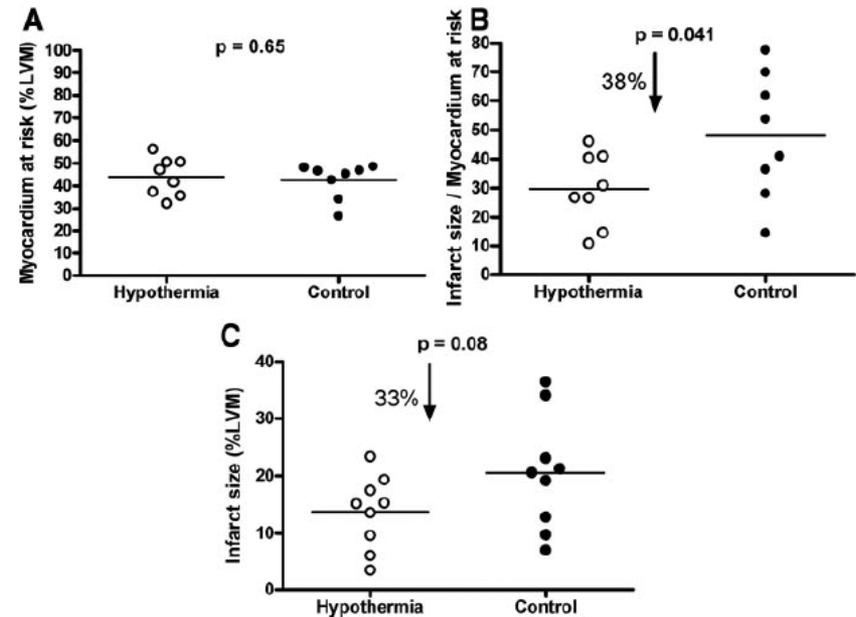


Figure 2. Time from arrival at the hospital until reperfusion (door-to-balloon time). There was no significant delay in reperfusion caused by hypothermia treatment.



Chill-MI, 60 vs 60 Pat. multizentrisch

Götberg, Circ CI 2010

# Oscar Langendorff (\*1853, †1908)



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(Aus dem physiologischen Institut in Rostock.)

## Untersuchungen am überlebenden Säugethierherzen.

Mitgetheilt von

**O. Langendorff.**

II. A b h a n d l u n g.

### Ueber den Einfluss von Wärme und Kälte auf das Herz der warmblütigen Thiere.

(Nach Versuchen von O. Langendorff und Dr. Czeslaw Nawrocki.)

Hierzu Tafel V, VI, VII u. VIII und 14 Textfiguren.

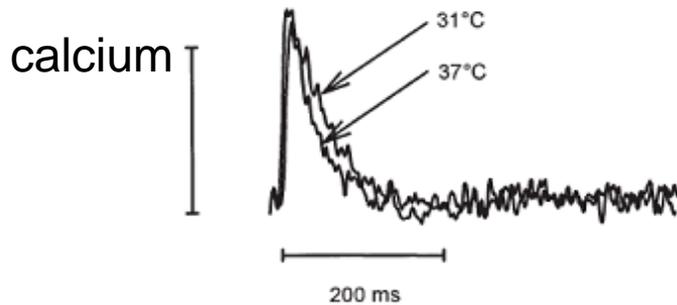
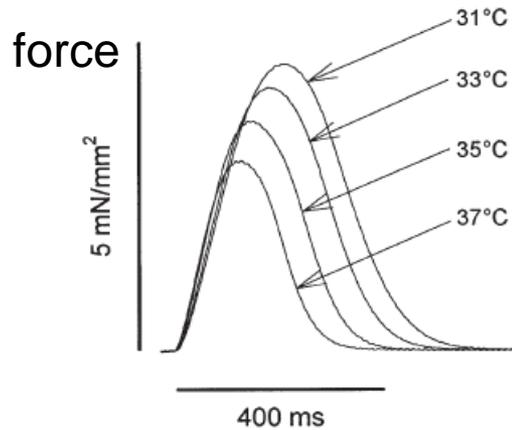
In meiner ersten Mittheilung über das isolirte Warmblüterherz (1) habe ich gezeigt, dass es gerade so wie das Froschherz von der Temperatur beeinflusst wird. Änderte man die Wärme des durch das Koronarsystem fließenden Blutes, so änderte sich entsprechend die Schlagzahl; dem rapiden Ablauf der Zusammenziehungen des warmen stand die träge Kontraktion des abgekühlten Herzens gegenüber. Ich zeigte ferner, dass das Herz noch bei Speisung mit stark abgekühltem Blut seine Schläge nicht einstellt.

# MH: normal heart

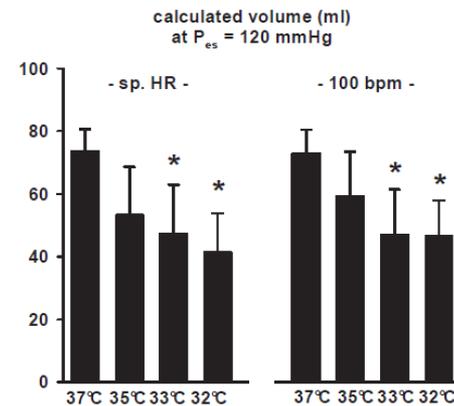
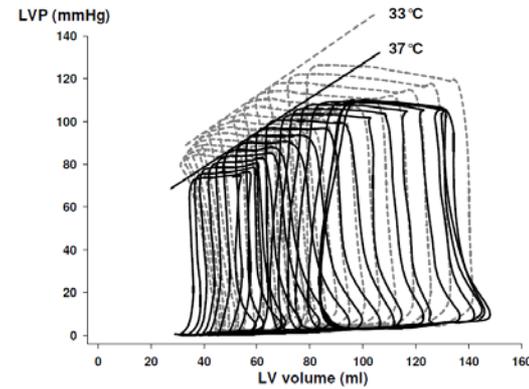


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isolated human cardiac muscle strips



pig in vivo



# Experimental setup



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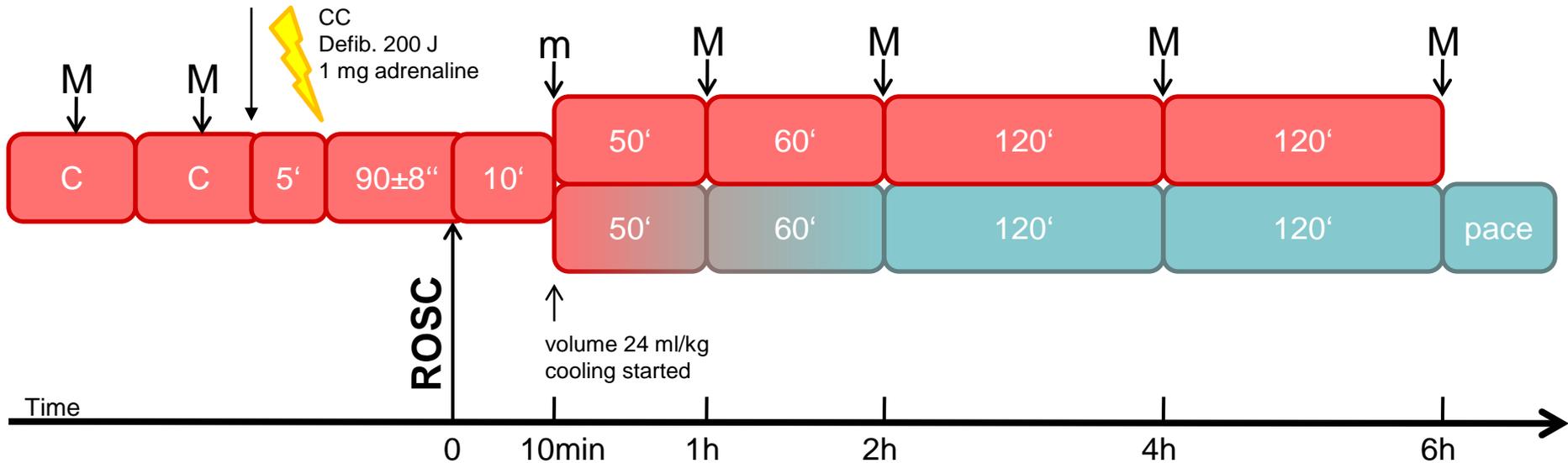
# Resuscitation



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- 38 °C, n=8
- 33 °C, n=8 (129±8 min)

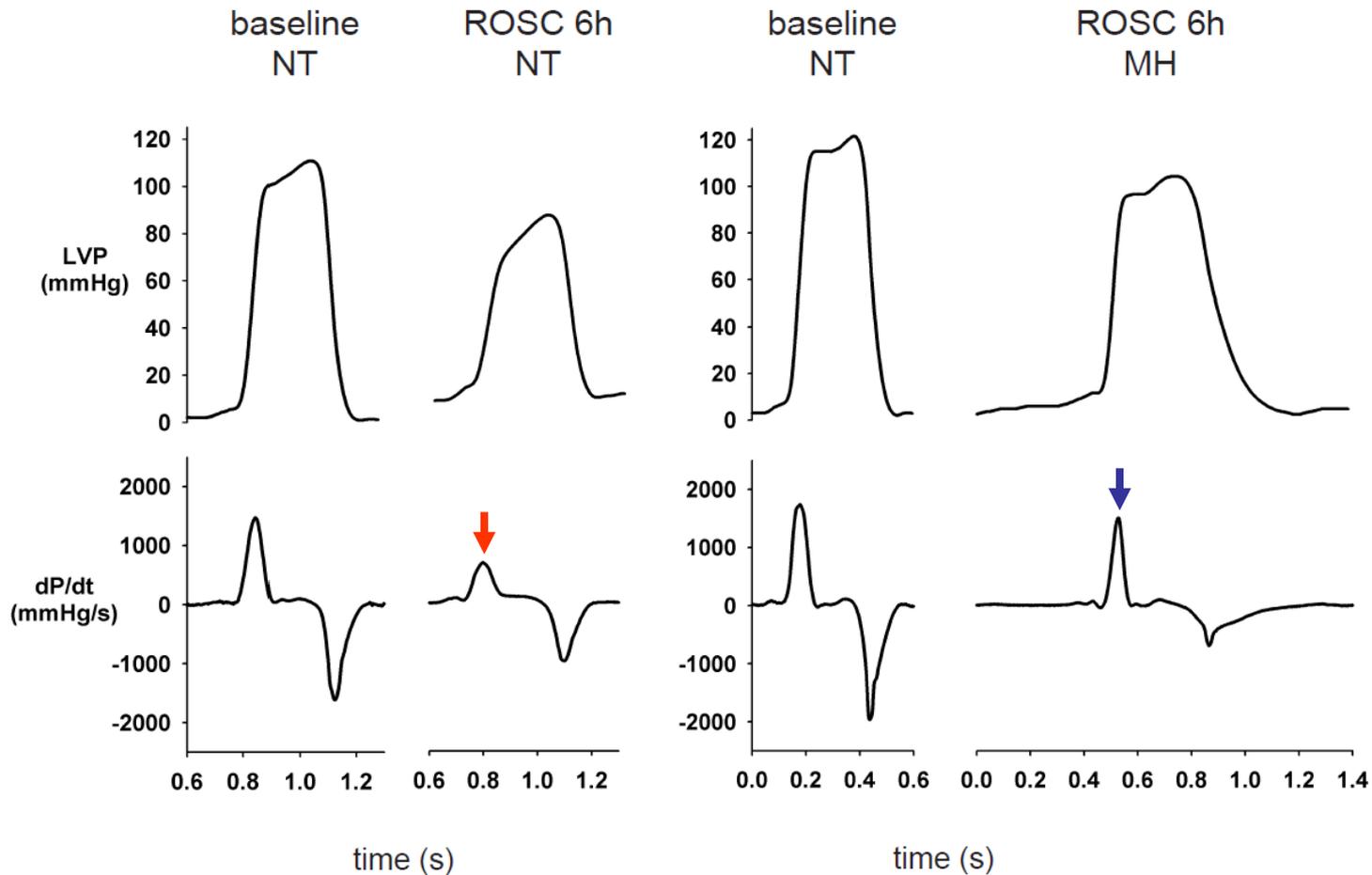


M: saline, CO, BGA art./ven., blood samples, aortic occlusion  
 m: saline, CO, BGA art./ven., blood samples

# Resuscitation: LVP



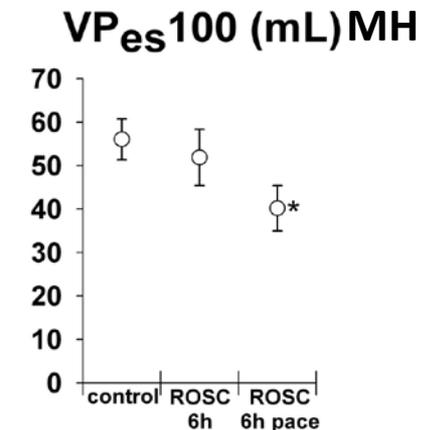
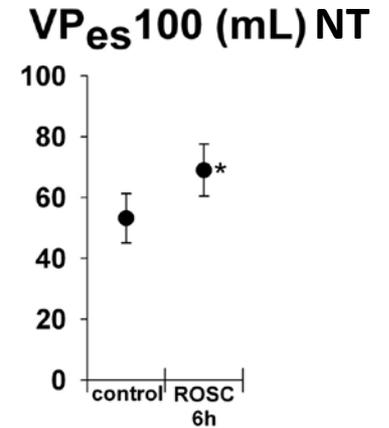
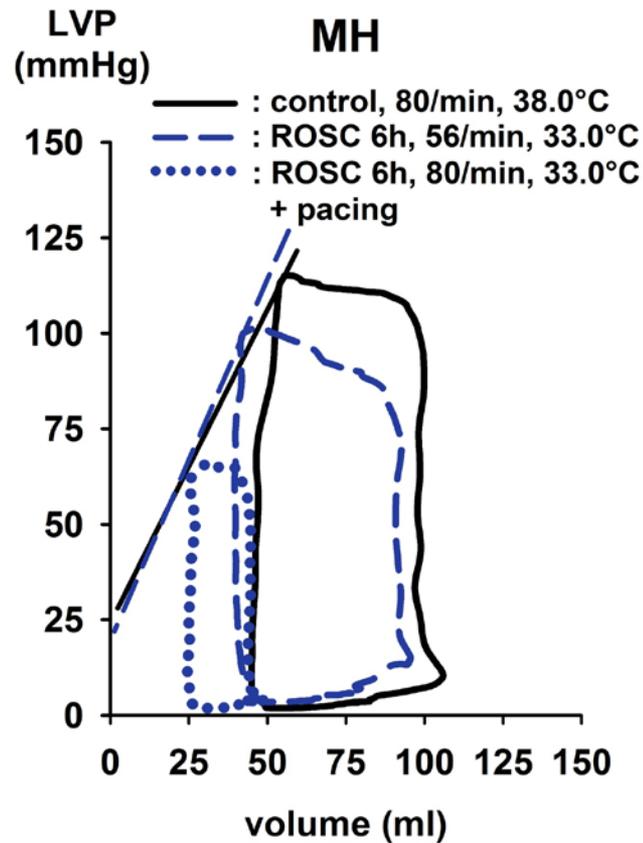
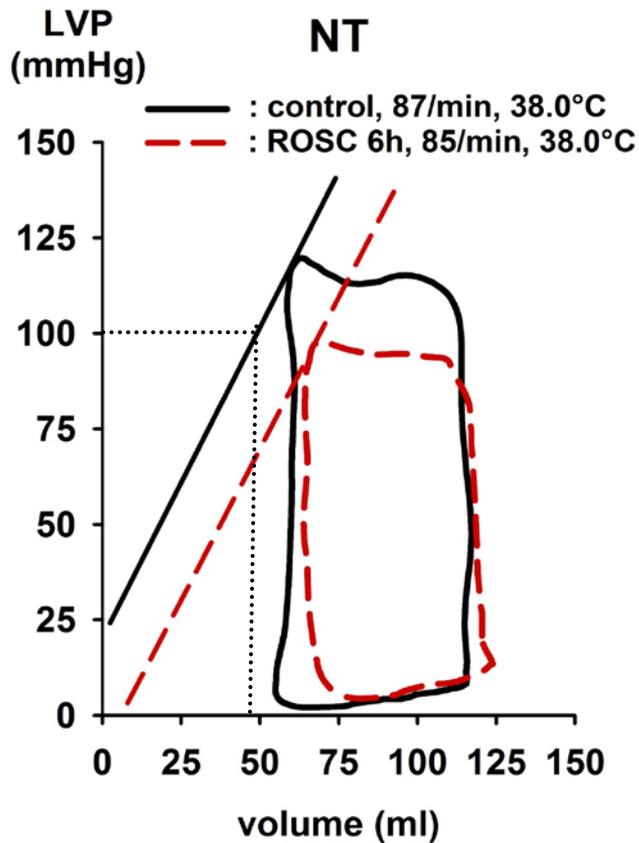
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# MH is a positive inotrope



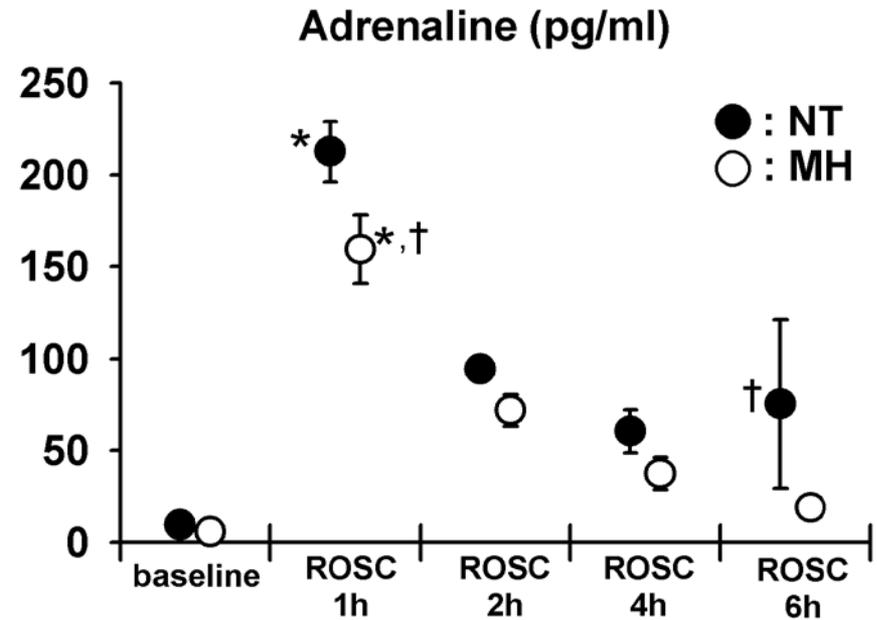
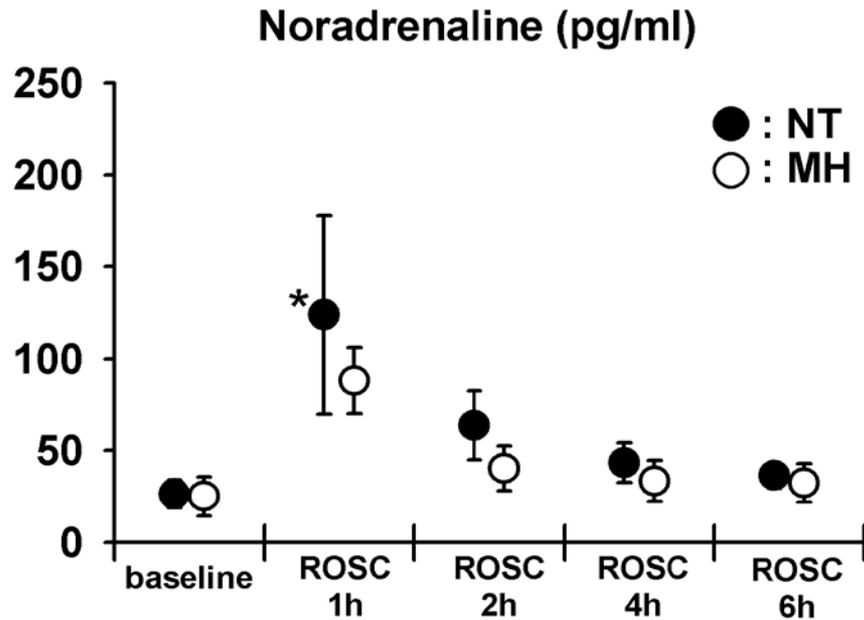
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# Resuscitated porcine heart: No further sympathetic activation during MH



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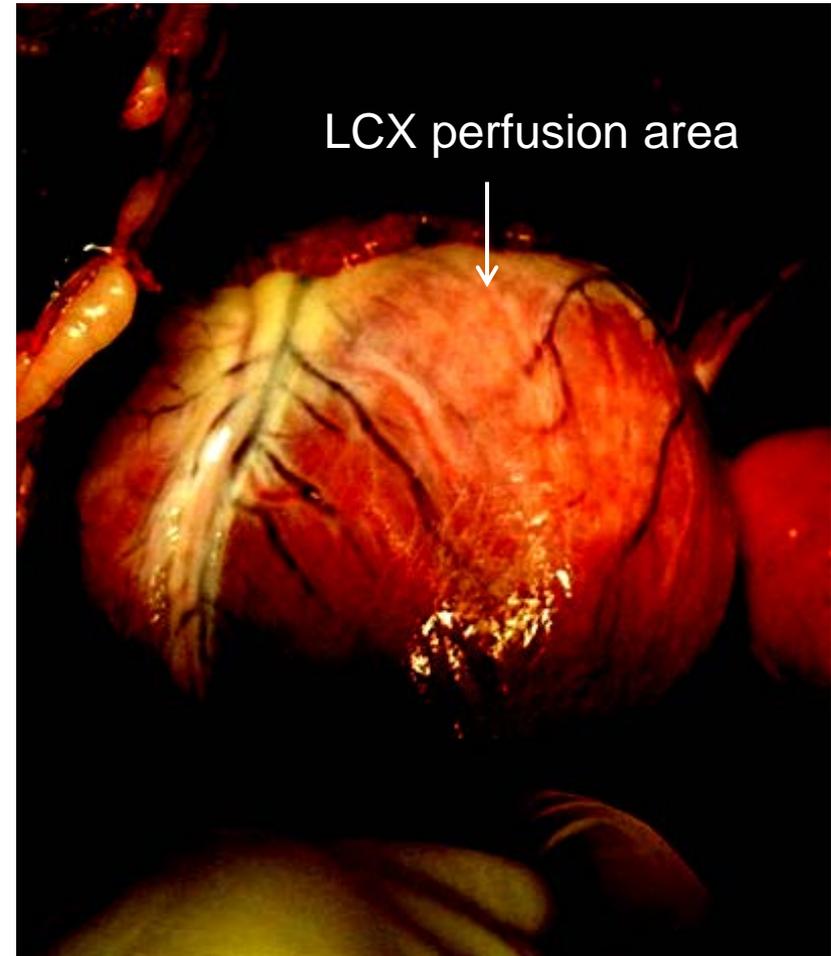
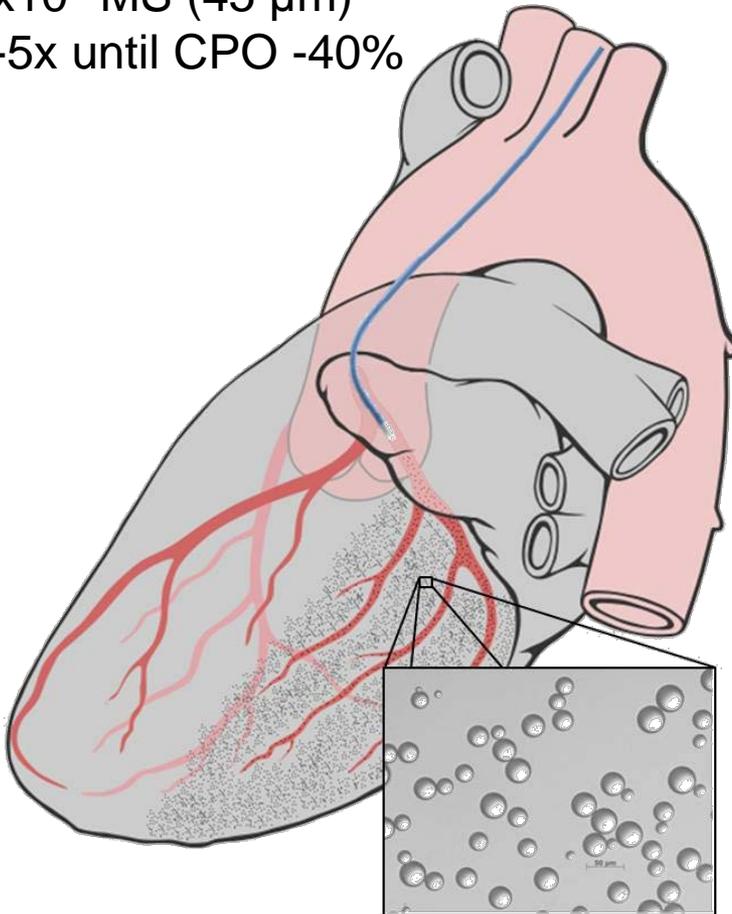


# No-reflow myocardial infarction



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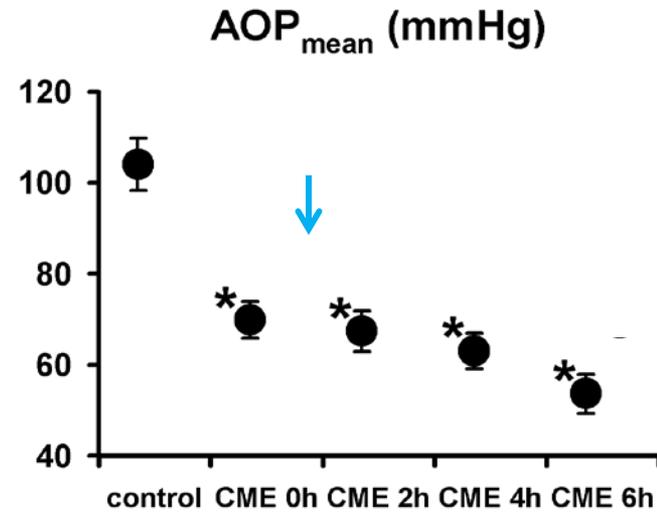
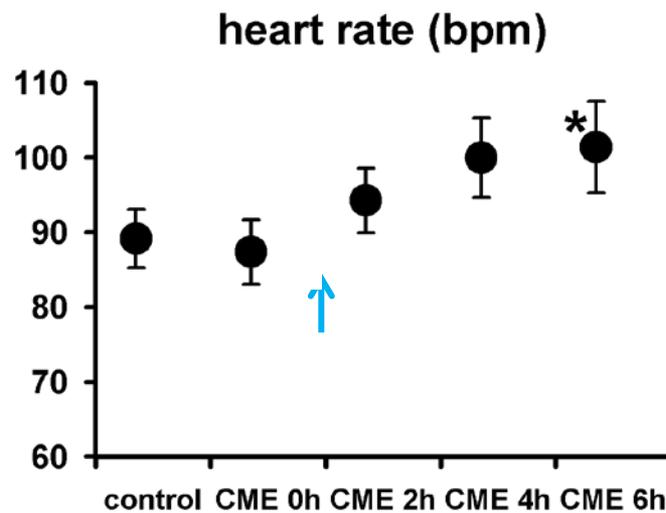
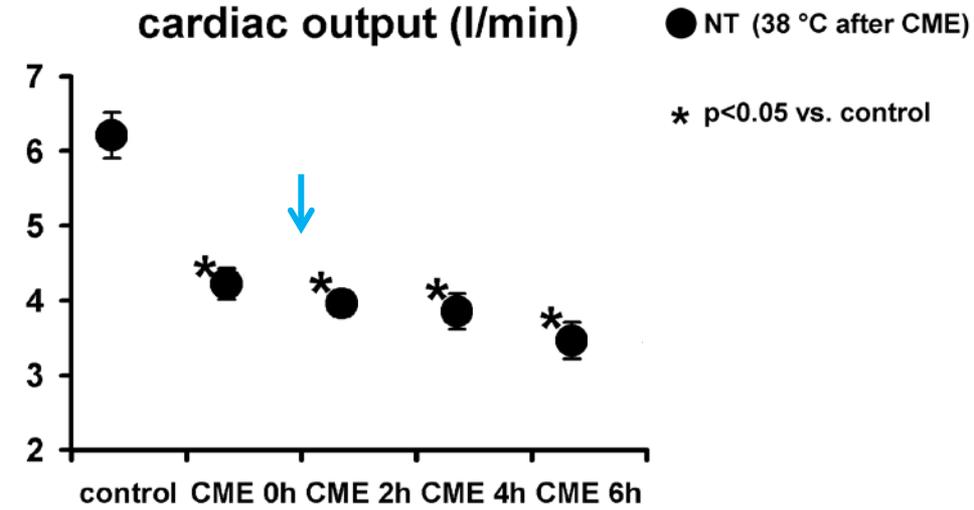
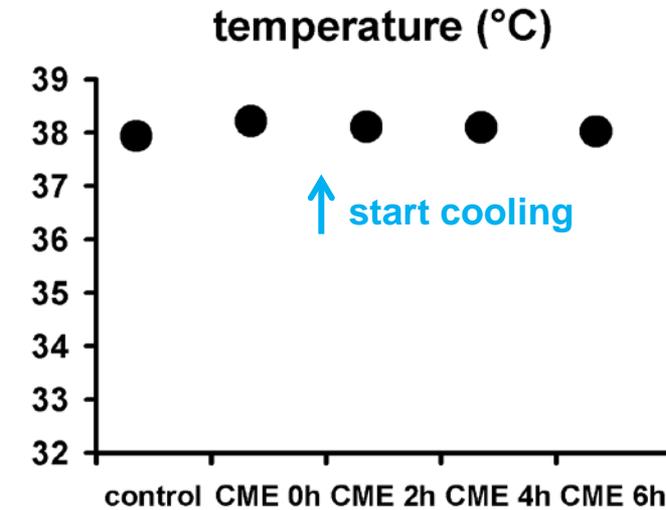
$5 \times 10^5$  MS (45  $\mu\text{m}$ )  
3-5x until CPO -40%



# MH in acute MI



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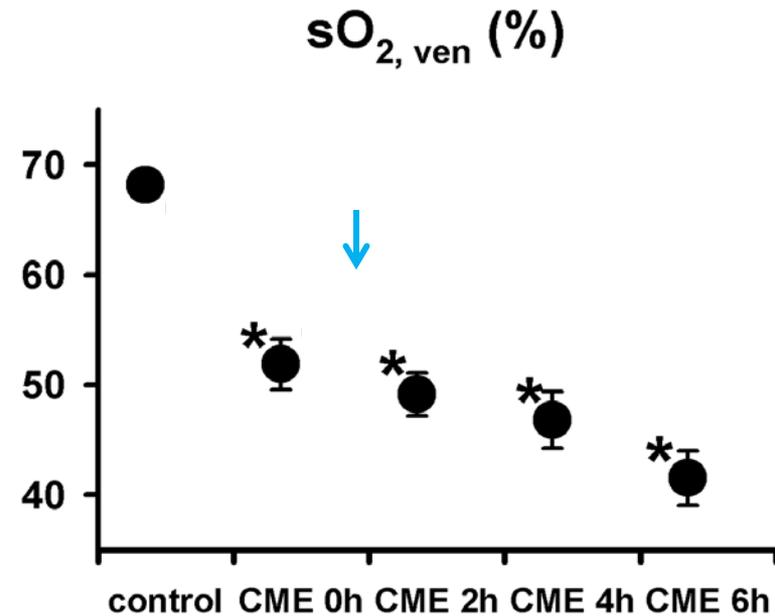
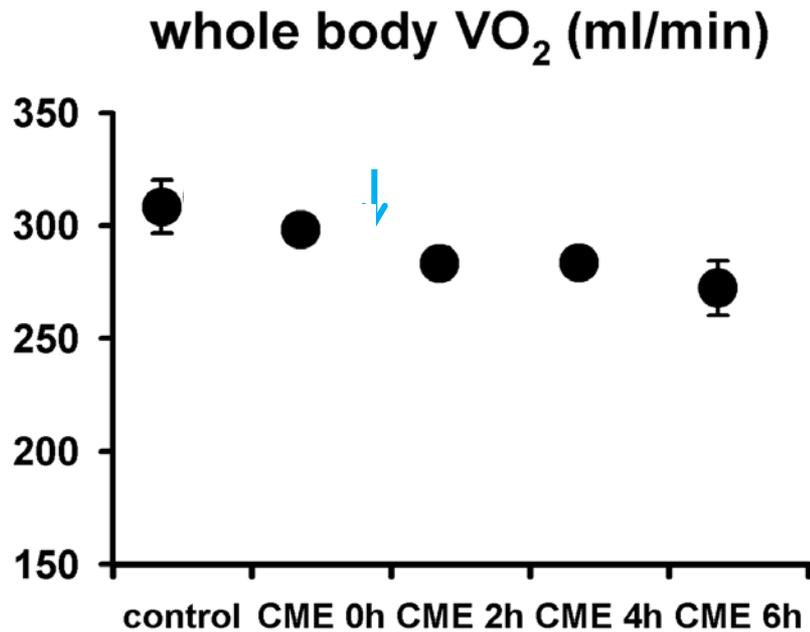


# MH in acute MI



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- NT (38 °C after CME)
- MH (33 °C after CME)
- \* p<0.05 vs. control
- † p<0.05 vs. NT



# MH in cardiogenic shock



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systemic oxygen demand

systemic oxygen supply

hypothermia

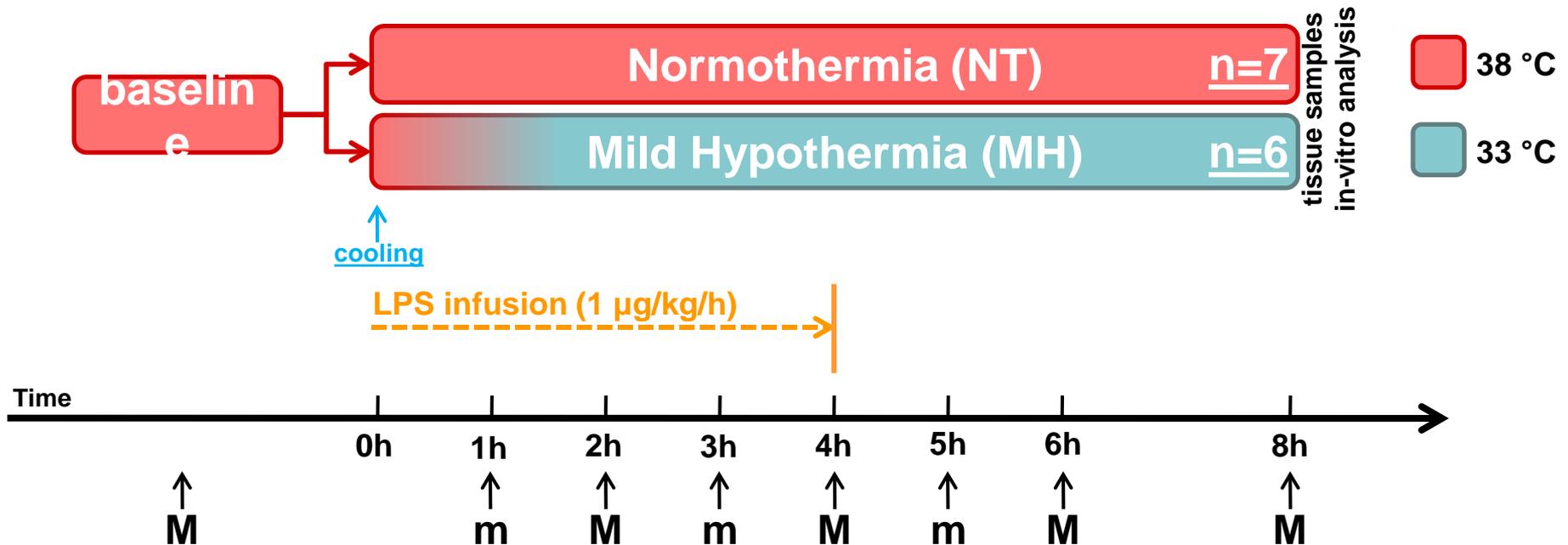


preload  
inotropes  
pacing  
IABP  
Impella  
ECMO

# MH in endotoxinemia (LPS)



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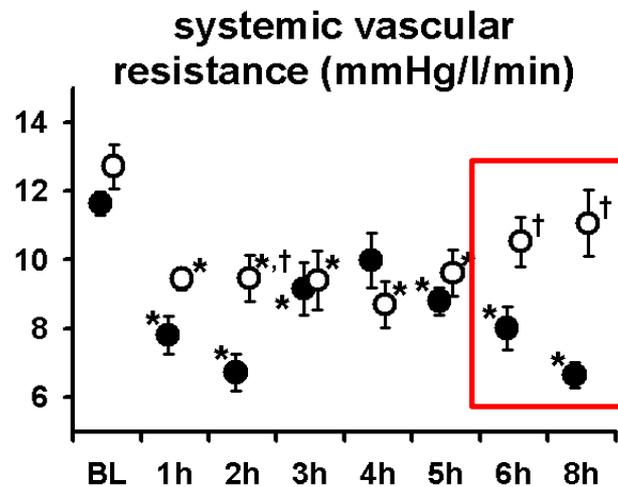
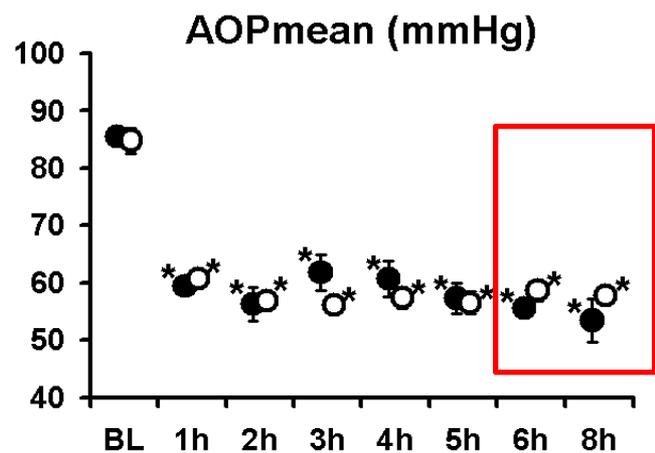
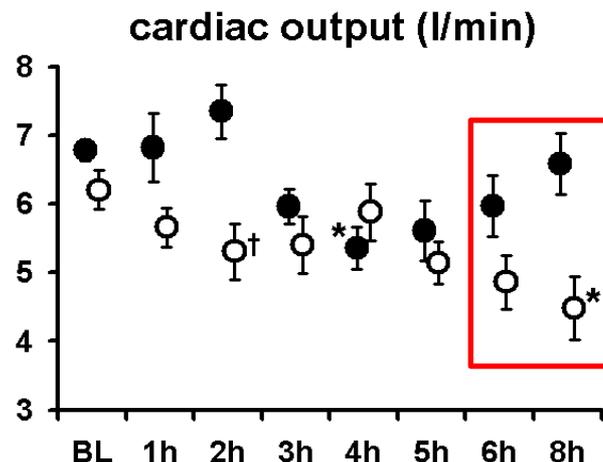
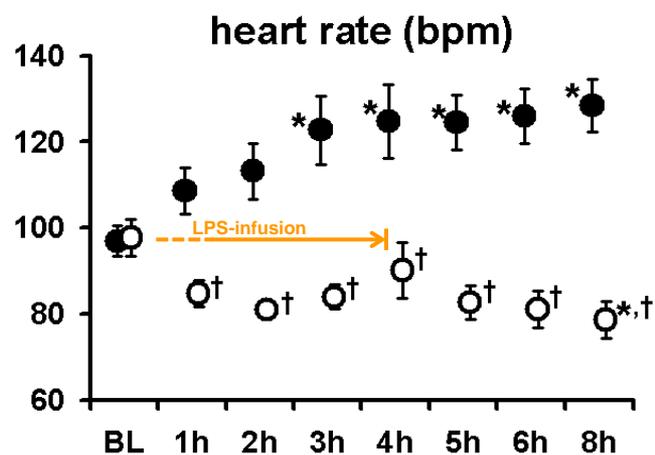


M: haemodynamics, aortic occlusion, pacing, cytokines, blood samples, catecholamines, heart rate variability  
m: haemodynamics, cytokines

# LPS + MH/Systemic haemodynamics: MH preserves systemic vascular resistance



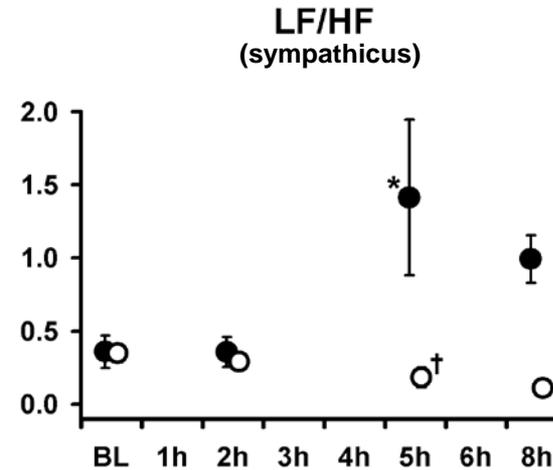
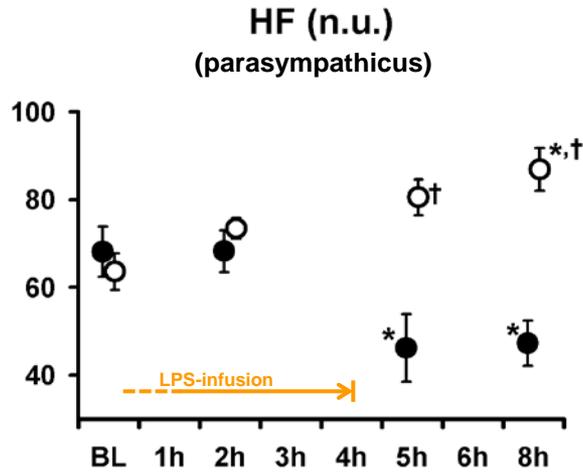
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# LPS + MH: autonomic tone, $\beta$ -adrenergic myocardial responsiveness



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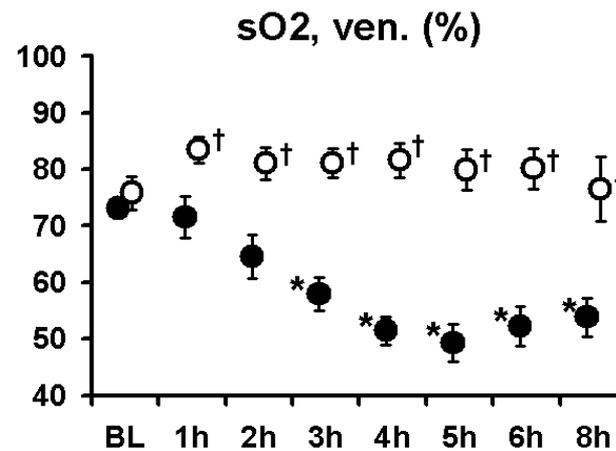
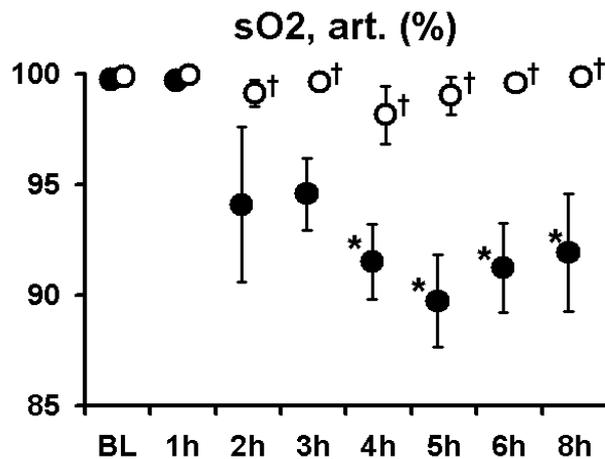
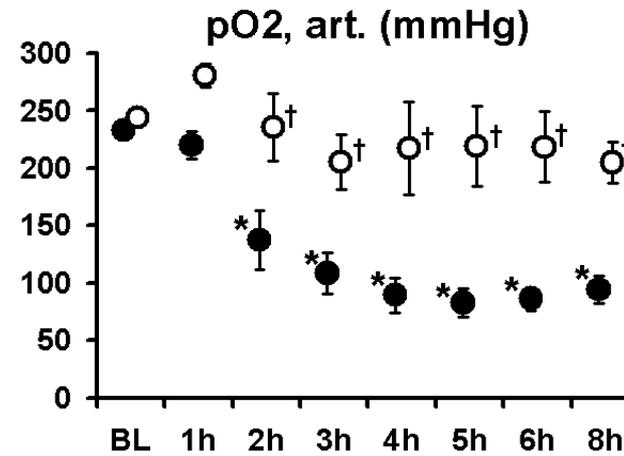
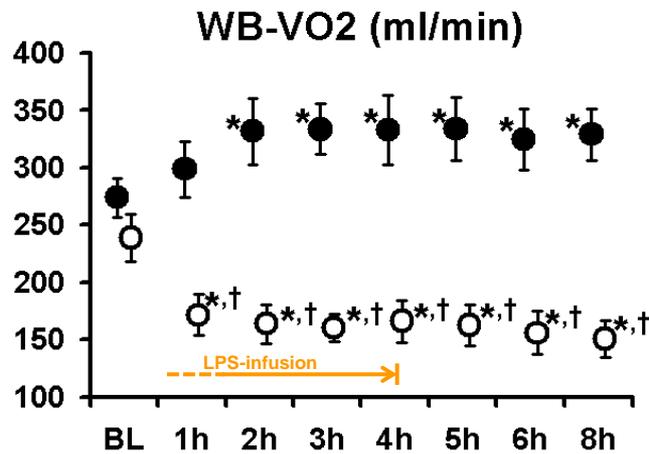


● : 38 °C (NT, n=7)  
○ : 33 °C (MH, n=6)  
\* : p<0.05 vs. baseline (BL)  
† : p<0.05 vs. MH

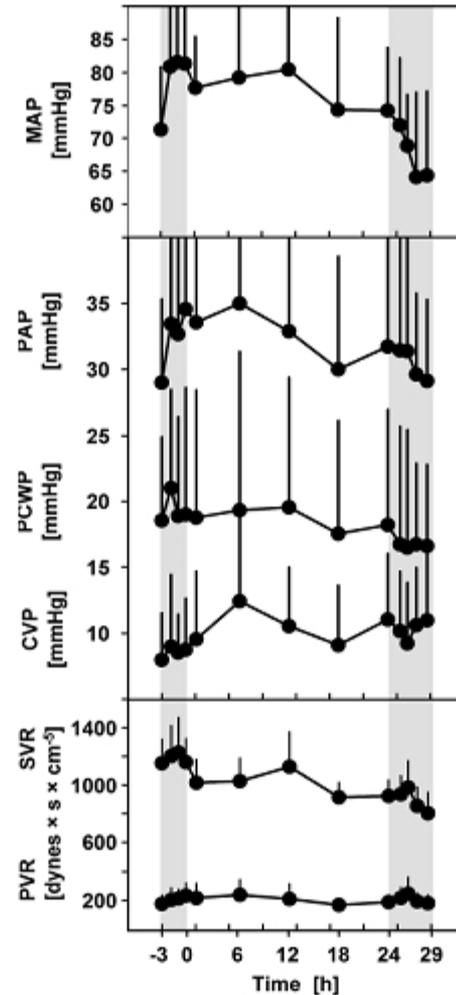
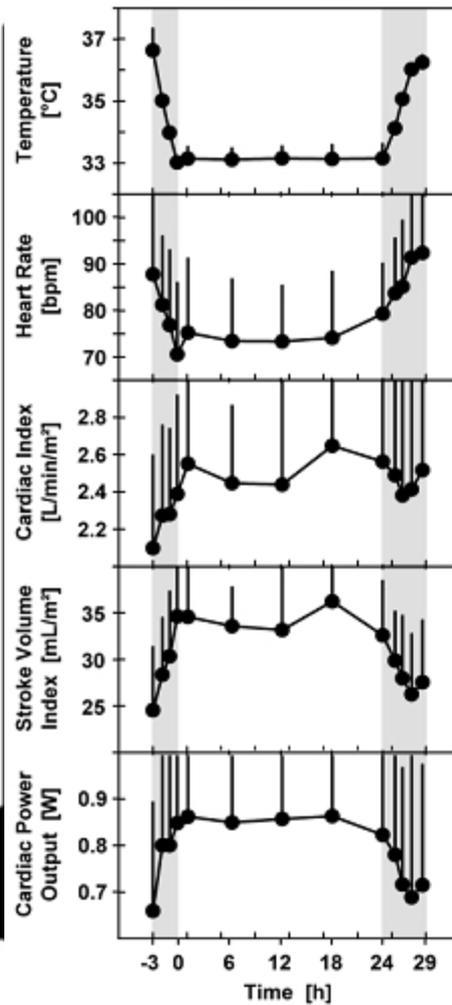
# LPS + MH: preserved systemic oxygen supply/demand balance



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Patients (n=14): MH during cardiogenic shock





## Patienten:

- MH vor Reperfusion reduziert die Infarktgröße bei STEMI
  - multizentrische Studie Chill-MI
- Der kardiogene Schock ist keine Kontraindikation für MH
  - wahrscheinlich protektiv → Studien

## Experimentell:

- MH ist ohne sympathische Aktivierung positiv inotrop
  - MH induziert eine diastolische Dysfunktion
    - durch spontane Bradykardie kompensiert
  - MH erhöht den systemischen Gefäßwiderstand
  - MH verbessert für einen gegebenen CO die SvO<sub>2</sub>
  - MH ist (kurzfristig) lungenprotektiv
-



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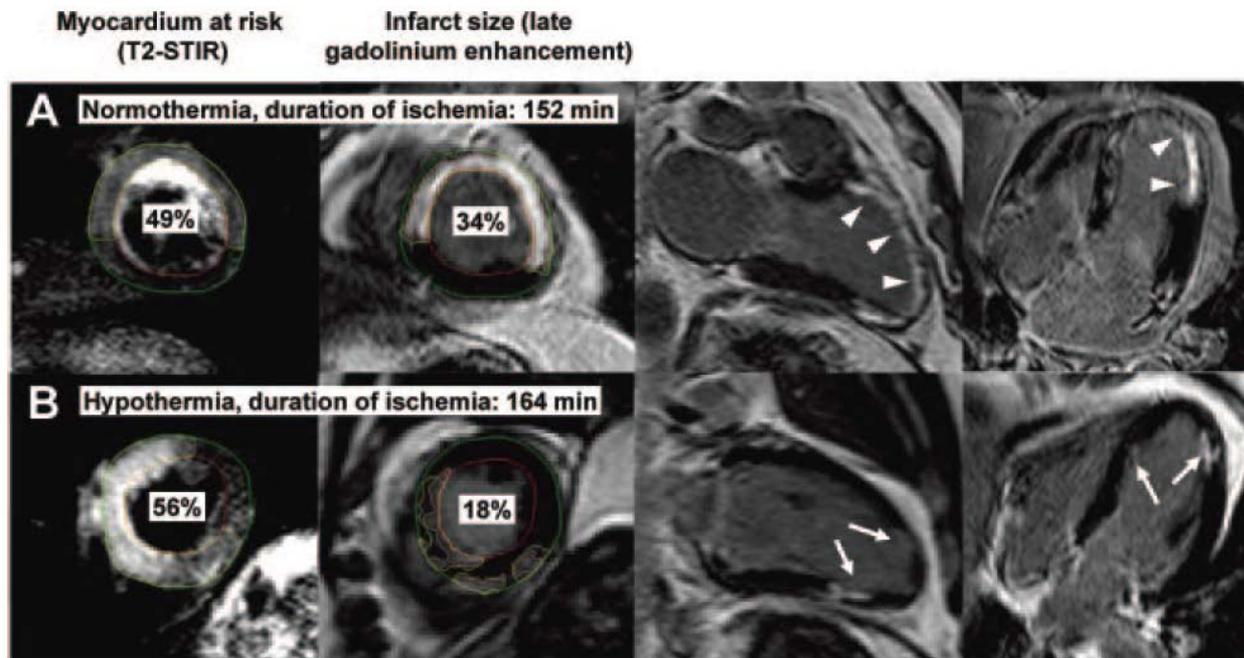
**Vielen Dank!**

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# Prä-Reperfusion MH reduziert die Infarktgröße bei STEMI



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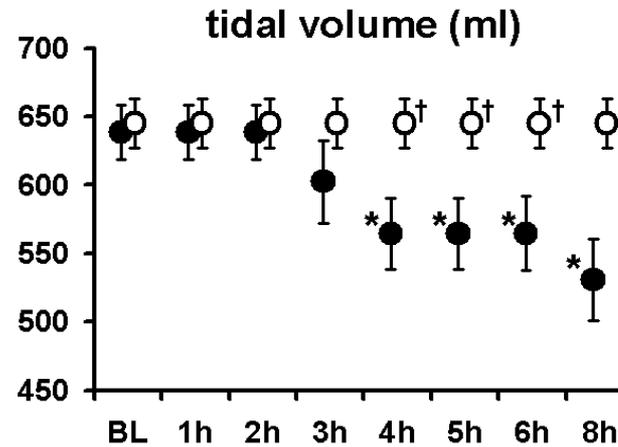
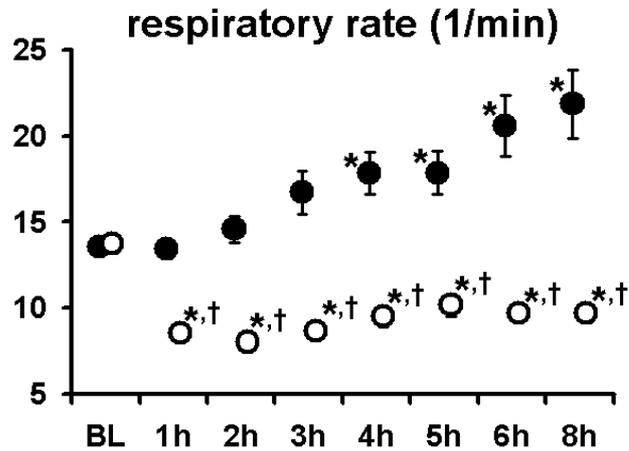


Götberg, Circ CI 2010

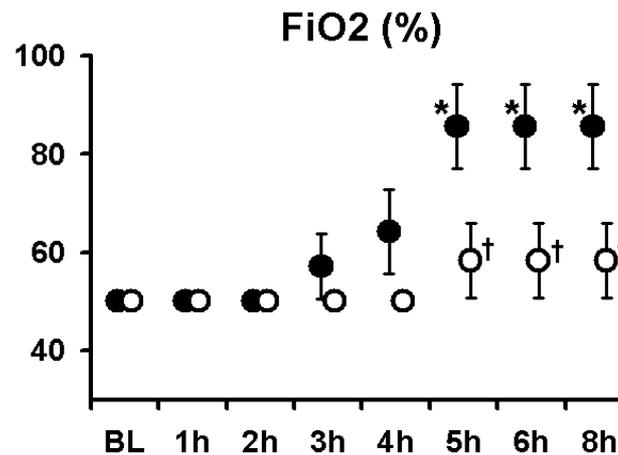
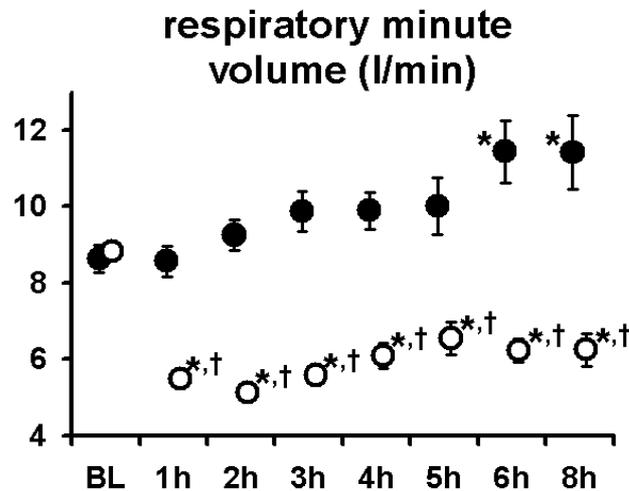
# Ventilation parameters



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●: 38 °C (NT, n=7)  
○: 33 °C (MH, n=6)  
\*: p<0.05 vs. baseline (BL)  
†: p<0.05 vs. NT



# Pulmonary mechanics



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