## SUPPLEMENTAL MATERIAL

for

## IcyHot analgesic cream triggers remote cardioprotection in rodents

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### Supplemental Tables:

Supplemental Table 1. Hemodynamics for analgesic cream infarct size studies. Hemodynamic results for *in vivo* experimental groups measured at baseline, at 15 minutes of ischemia, and at 2 hours of reperfusion. No significant changes were noted between groups for each phase relative to control. Statistical differences were noted when comparing within groups reperfusion values to baseline values ( $^{+}P<0.01$ , relative to baseline value,  $^{*}P<0.01$ , relative to control for that time point). Abbreviations for the table are as follows: HR = heart rate (beats/min), MAP = mean arterial pressure (mmHg), RPP = rate pressure product (mmHg/1000).

Supplemental Table 2. Hemodynamics for TRPA1 inhibitor infarct size studies. Hemodynamic results for *in vivo* experimental groups measured at baseline, at 15 minutes of ischemia, and at 2 hours of reperfusion. No significant changes were noted between groups for each phase relative to control. Statistical differences were noted when comparing within groups reperfusion values to baseline values ( $^+P<0.01$ , relative to baseline value). Abbreviations for the table are as follows: HR = heart rate (beats/min), MAP = mean arterial pressure (mmHg), RPP = rate pressure product (mmHg/1000).

Supplemental Table 3. Hemodynamics for timing of lcyHot infarct size studies. Hemodynamic results for *in vivo* experimental groups measured at baseline, at 15 minutes of ischemia, and at 2 hours of reperfusion. No significant changes were noted between groups for each phase relative to control. Statistical differences were noted when comparing within groups reperfusion values to baseline values ( $^{+}P<0.01$ , relative to baseline value). Abbreviations for the table are as follows: HR = heart rate (beats/min), MAP = mean arterial pressure (mmHg), RPP = rate pressure product (mmHg/1000).

#### Supplemental Figures:

Supplemental Figure 1. Area at risk per left ventricle for analgesic cream infarct size studies. Area at risk per left ventricle percentage (AAR/LV%) for each experimental group. No significant differences were seen between groups.

Supplemental Figure 2. Dose-response for methyl salicylate in isolated cardiac myocytes subjected to hypoxia-reoxygenation. Percentage of trypan blue positive cells after isolated adult cardiac myocytes are subjected to 2 hours hypoxia followed by 4 hours reoxygenation. Methyl salicylate dose-dependently reduced the number of trypan blue positive cells after reoxygenation. \*P<0.01 versus DMSO treated control.

**Supplemental Figure 3. Area at risk per left ventricle for TRPA1 inhibitor infarct size studies.** Area at risk per left ventricle percentage (AAR/LV%) for each experimental group. No significant differences were seen between groups.

**Supplemental Figure 4. Area at risk per left ventricle for timing of lcyHot infarct size studies.** Area at risk per left ventricle percentage (AAR/LV%) for each experimental group. No significant differences were seen between groups.

# Supplemental Table 1.

		Baseline			15min Ischemia			2hr Reperfusion		
	n	HR	MAP	RPP	HR	MAP	RPP	HR	MAP	RPP
Control	6	414±8	103±10	48±5	408±8	101±7	46±4	374±10	79±4	37±2
lcyHot	6	440±17	112±8	57±5	437±14	92±11	47±4	412±18	79±6+	40±3+
Preparation H	6	418±11	119±6	59±3	441±12	108±8	58±4	425±12	91±3+	52±3*
Aspercreme	6	422±18	121±8	60±4	421±21	107±10	54±5	386±8	87±3+	44±2+
Tiger Balm	6	387±9	109±3	49±1	397±12	94±11	44±4	380±10	91±4	42±3

# Supplemental Table 2.

		E	Baseline		15m	nin Ischei	mia	2hr	Reperfus	sion
	n	HR	MAP	RPP	HR	MAP	RPP	HR	MAP	RPP
TCS+IcyHot	6	411±15	105±4	51±3	419±14	93±10	45±5	392±6	77±5+	38±2
AP+lcyHot	6	421±14	117±3	55±3	423±10	109±9	50±4	383±14	82±5+	38±3+
TCS	6	400±23	111±8	53±5	418±15	105±9	51±4	367±8	76±4+	41±3
AP	6	404±17	122±6	58±4	421±10	124±5	58±3	375±9	79±6+	41±3+

		Baseline				15min Ischemia				2hr Reperfusion			
_	n	HR	MAP	RPP		HR	MAP	RPP		HR	MAP	RPP	
Control Ischemia	6	406±11	104±2	50±2		422±11	86±5	42±2		384±16	78±4+	38±3	
IcyHot Ischemia	6	396±22	106±6	50±5		397±17	94±5	45±4		365±12	79±6+	36±3+	
Control 24 hour	6	409±12	113±7	54±4		448±12	107±5	55±4		390±20	82±4+	41±4+	
lcyHot 24 hour	6	417±9	132±4	65±2		411±14	116±10	58±6		357±5⁺	98±6+	44±3+	

Supplemental Figure 1.





Supplemental Figure 3.

Supplemental Figure 4.

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