

AD-A175 286

CHANGES IN PHYSIOLOGIC VARIABLES IN RESPONSE TO  
PHYSICAL RESTRAINT DURING (U) LETTERMAN ARMY INST OF  
RESEARCH PRESIDIO OF SAN FRANCISCO CA  
G D BONNER ET AL. AUG 86 LAIR-87-64

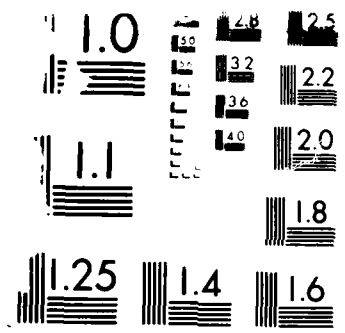
1/1

UNCLASSIFIED

F/G 6/5

NL





Resolution Test Chart

2



AD-A175 286

LABORATORY NOTE NO. 87-64

CHANGES IN PHYSIOLOGIC VARIABLES IN RESPONSE TO  
PHYSICAL RESTRAINT DURING HEMORRHAGE  
IN CONSCIOUS SWINE

GARY D. BONNER, SGT  
CAROL A. BOSSONE, SGT  
BERRY F. WILLIAMS, SGT  
DIANE S. TRAIL, SSG  
CHARLES E. WADE, Ph.D.  
and  
JOHN P. HANNON, Ph.D.

JTIC FILE COPY

DIVISION OF MILITARY TRAUMA RESEARCH

AUGUST 1986

LETTERMAN ARMY INSTITUTE OF RESEARCH  
PRESIDIO OF SAN FRANCISCO, CALIFORNIA 94129

DTIC  
ELECTE  
DEC 23 1986  
S D  
E

86 12 22 007

REPORT DOCUMENTATION PAGE		READ INSTRUCTIONS BEFORE COMPLETING FORM
1. REPORT NUMBER LAIR Laboratory Note No. 87-64	2. GOVT ACCESSION NO.	3. RECIPIENT'S CATALOG NUMBER
4. TITLE (and Subtitle) Changes in Physiologic Variables in Response to Physical Restraint During Hemorrhage in Conscious Swine		5. TYPE OF REPORT & PERIOD COVERED
		6. PERFORMING ORG. REPORT NUMBER
7. AUTHOR(s) G.D. Bonner, C.A. Bossone, B.F. Williams, D.S. Trail, C.E. Wade and J.P. Hannon		8. CONTRACT OR GRANT NUMBER(s)
9. PERFORMING ORGANIZATION NAME AND ADDRESS Division of Military Trauma Research Letterman Army Institute of Research Presidio of San Francisco, CA 94129-6800		10. PROGRAM ELEMENT, PROJECT, TASK AREA & WORK UNIT NUMBERS 3S162772A874 AB 241
11. CONTROLLING OFFICE NAME AND ADDRESS Division of Military Trauma Research Letterman Army Institute of Research Presidio of San Francisco, CA 94129-6800		12. REPORT DATE August 1986
14. MONITORING AGENCY NAME & ADDRESS (if different from Controlling Office)		13. NUMBER OF PAGES 8
		15. SECURITY CLASS. (of this report)  Unclassified
15a. DECLASSIFICATION/DOWNGRADING SCHEDULE		
16. DISTRIBUTION STATEMENT (of this Report)  Approved for public release and sale; distribution is unlimited		
17. DISTRIBUTION STATEMENT (of the abstract entered in Block 20, if different from Report)		
18. SUPPLEMENTARY NOTES		
19. KEY WORDS (Continue on reverse side if necessary and identify by block number)  Immobilization; Swine		
20. ABSTRACT (Continue on reverse side if necessary and identify by block number)  *see reverse		

## ABSTRACT

Responses to hemorrhage (38.5 ml/kg/60 min) were evaluated in chronically catheterized conscious pigs (20-25 kg) restrained in a portable holding cage (n=6), a squeeze cage (n=8), or a Pavlov sling (n=8). Before hemorrhage, heart rates of squeeze cage animals ( $146 \pm 4$  b/m,  $\bar{x} \pm \text{SEM}$ ) and sling animals ( $129 \pm 9$  b/m) were significantly higher ( $p < 0.05$ ) than heart rates of holding cage pigs ( $107 \pm 5$  b/m). Hemorrhage caused a further increase in heart rate ( $216 \pm 15$  b/m) in sling animals, but not in holding cage animals ( $104 \pm 7$  b/m) or squeeze cage animals ( $157 \pm 20$  b/m). Initially animals in the squeeze cage had a mean arterial pressure ( $90 \pm 4$  mmHg), lower than in the holding cage animals ( $105 \pm 2$  mmHg). However, these differences were not significant. Following hemorrhage, mean arterial pressures were similar in holding cage ( $46 \pm 4$  mmHg) and squeeze cage animals ( $43 \pm 4$  mmHg) while the mean arterial pressure in sling animals was elevated ( $58 \pm 8$  mmHg). There were no differences in hematocrit values between groups either before or after hemorrhage. Plasma glucose and lactate levels were lower in sling animals before hemorrhage ( $82 \pm 4$  mg/dl and  $8.1 \pm 0.05$  mg/dl) and after hemorrhage ( $161 \pm 31$  mg/dl and  $85.2 \pm 6.9$  mg/dl) than in holding cage animals. Arterial  $\text{PO}_2$  values were elevated in squeeze cage and sling animals throughout the experiment compared to holding cage animals. Also their  $\text{PCO}_2$  values were lower than the holding cage animals. The evidence suggest that cardiovascular and metabolic responses are altered qualitatively and quantitatively by method of restraint during hemorrhage in conscious swine.

ABSTRACT

Responses to hemorrhage (38.5 ml/kg/60 min) were evaluated in chronically catheterized conscious pigs (20-25 kg) restrained in a portable holding cage (n=6), a squeeze cage (n=8), or a Pavlov sling (n=8). Before hemorrhage, heart rates of squeeze cage animals (145±4 b/m, x±SEM) and sling animals (129±9 b/m) were significantly higher (p<0.05) than heart rates of holding cage pigs (107±5 b/m). Hemorrhage caused a further increase in heart rate (216±15 b/m) in sling animals, but not in holding cage animals (104±7 b/m) or squeeze cage animals (157±20 b/m). Initially animals in the squeeze cage had a mean arterial pressure (99±4 mmHg), lower than in the holding cage animals (105±2 mmHg). However, these differences were not significant. Following hemorrhage, mean arterial pressures were similar in holding cage (46±4 mmHg) and squeeze cage animals (43±4 mmHg) while the mean arterial pressure in sling animals was elevated (58±8 mmHg). There were no differences in hematocrit values between groups either before or after hemorrhage. Plasma glucose and lactate levels were lower in sling animals before hemorrhage (82±4 mg/dl and 8.1±0.05 mg/dl) and after hemorrhage (161±31 mg/dl and 85.2±6.9 mg/dl) than in holding cage animals. Arterial PO<sub>2</sub> values were elevated in squeeze cage and sling animals throughout the experiment compared to holding cage animals. Also their PCO<sub>2</sub> values were lower than the holding cage animals. The evidence suggest that cardiovascular and metabolic responses are altered qualitatively and quantitatively by method of restraint during hemorrhage in conscious swine.

<b>Accession For</b>	
NTIS GRA&I	<input checked="" type="checkbox"/>
DTIC TAB	<input type="checkbox"/>
Unannounced	<input type="checkbox"/>
Justification	
By _____	
Distribution/	
Availability Codes	
Dist	Avail and/or Special
A-1	



## CHANGES IN PHYSIOLOGIC VARIABLES IN RESPONSE TO PHYSICAL RESTRAINT DURING HEMORRHAGE IN CONSCIOUS SWINE

Responses of cardiovascular, endocrinological, and biochemical variables may be altered both quantitatively and qualitatively by different methods of handling swine (1-3). The physiologic responses to hemorrhage were assessed in conscious swine restrained by three methods. The evidence shows that the responses varied in the three groups.

### MATERIALS AND METHODS

Twenty-two immature Yorkshire swine, body weight 20-25 kg, were studied. The animals were prepared surgically with a carotid artery catheter (1) or an aortic side port catheter (4). The pigs were allowed 5 to 7 days to recover from the operation. Before the experiment, animals were fasted overnight and moved to the laboratory in portable transfer cages. The animals were divided into three groups:

- Group I. Restrained in a portable holding cage (105x60x75 cm), n=6.
- Group II. Restrained in a squeeze cage (105x30x75 cm), n=8.
- Group III. Restrained in a modified Pavlov sling, n=8.

Heart rate and mean arterial pressure were measured by a cardiovascular monitoring system (5) comprised of an arterial catheter connected to a 1 ft (30.5 cm) pressure monitoring/injection line. The line was fitted with a three-way plastic stopcock and filled with heparinized saline (500 units/ml). Residual blood and heparinized saline were then aspirated from the catheter and the entire system was filled with fresh heparinized saline (10 units/ml). Thereafter, the stopcock was connected to the monitoring/injection line, filled with heparinized saline and connected to a Statham P23Db pressure transducer. The transducer was mounted to a ring stand positioned at the level of the animal's heart.

Heart rate, mean arterial pressure, and pulse pressure were measured by Gould 2200S recorder before and after hemorrhage. Over a period of 60 minutes the pigs were hemorrhaged 50 percent of their total blood volume (38.5 ml/kg) as estimated by the vonEngelhardt regression equation (2). Blood samples were drawn before and after hemorrhage for analysis by a ILS System 1303 pH/blood gas analyzer, and hematocrit determined with a Clay-Adams microhematocrit centrifuge. Additionally, plasma glucose levels were determined by the Smith-Kline Spichem reagent method and plasma lactate according the method described in Sigma Technical Bulletin No. 7261-UV. Difference between groups of animals were determined by using a two-way ANOVA and Newman-Keuls test. Significance was determined at  $P < 0.05$ , and values shown as mean  $\pm$ SEM.

## RESULTS

Before hemorrhage the mean heart rate of squeeze cage (Group II) animals was  $145 \pm 4$  b/m ( $\bar{x} \pm$ SEM for all values), and sling (Group III) was  $129 \pm 9$  b/m. The control (Group I) animals had a mean heart rate of  $107 \pm 5$  b/m (Figure 1). After hemorrhage, the mean heart rate of Group III animals was increased to  $216 \pm 15$  b/m. In response to hemorrhage, animals in Groups I and II showed no change in heart rate. Immediately before hemorrhage, animals in the squeeze cage (Group II) had a mean arterial pressure of  $99 \pm 4$  mmHg, and holding cage (Group I) animals had a mean arterial pressure of  $105 \pm 2$  mmHg. The mean arterial pressure of Group III animals was elevated to  $111 \pm 7$  mmHg, an increase of approximately 6 percent. These values were not significantly different (Figure 1). After hemorrhage, the mean arterial pressure of all groups declined.

Before hemorrhage, the mean pulse pressure of control (Group I) animals was  $42 \pm 2$  mmHg while the mean pulse pressure of Group II was  $53 \pm 5$  mmHg, and Group III was  $59 \pm 4$  mmHg. After hemorrhage, the mean pulse pressure of Group III animals in the Pavlov sling was  $39 \pm 7$  mmHg (33 percent less than the mean pulse pressure of control animals).

There were no differences in hematocrits between groups of animals either before or after hemorrhage (Figure 2). Before hemorrhage, the mean plasma glucose levels were lower in both Group II ( $74 \pm 12$  mg/dl) and Group III ( $82 \pm 4$  mg/dl) than in control (Group I) animals, ( $87 \pm 3$  mg/dl).

The squeeze cage (Group II) animals had a mean plasma glucose level significantly lower than control animals (Figure 2).

Before hemorrhage, the mean lactate level in Group III animals was  $8.1 \pm 0.5$  mg/dl, and significantly lower than control (Group I) animals ( $10.2 \pm 1.1$  mg/dl). After hemorrhage the mean lactate levels were elevated in all groups. The mean lactate level in sling animals was significantly lower than control animals (Figure 2).

The arterial pH between groups of animals before and after hemorrhage showed no significant change (Figure 3).

Before hemorrhage, the mean arterial PO<sub>2</sub> values were elevated significantly in Group I animals ( $94.5 \pm 3.5$  torr) and Group II animals ( $92.9 \pm 3.1$  torr). After hemorrhage, the mean arterial PO<sub>2</sub> values were once again elevated significantly in Group I animals ( $126.5 \pm 6.0$  torr) and Group II animals ( $115 \pm 6.0$  torr), (Figure 3).

Before hemorrhage, the mean arterial PCO<sub>2</sub> values were reduced in squeeze cage ( $37.1 \pm 0.9$  torr) and sling animals ( $38.3 \pm 0.8$  torr) compared to control animals ( $41.0 \pm 1.1$  torr). This reduction was significant in squeeze cage (Group II) animals. After hemorrhage, the mean arterial PCO<sub>2</sub> values were reduced in all groups. However, these values were lower in squeeze cage (Group II) animals ( $26.7 \pm 1.4$  torr) compared to control (Group I) animals ( $28.4 \pm 1.9$  torr), (Figure 3).

#### COMMENT

Studies indicate that physiologic responses, e.g., cardiovascular, biochemical, and endocrinologic can be altered by handling and restraint of laboratory animals (1-3,5). Hannon (1) showed that swine resting in a recumbent position for 30 minutes have relatively stable vital signs. VonEngelhardt (2) and Wade, et al (3) showed that animals excited by handling have variable heart rates, blood pressures and catecholamine levels. Our study assessed the changes in physiologic responses produced by different environmental factors and in response to hemorrhage in three groups of conscious swine, i.e., a holding cage that allowed the swine to assume a recumbent position (Group I); a squeeze cage that allowed the pig to lay down but not move about (Group II); a modified Pavlov sling that

completely restrained the animal (Group III). An evaluation of the data suggests that all physiologic responses were consistent with results from similar studies (1-3,5). In most instances the greater the degree of restraint (modified Pavlov sling >squeeze cage >holding cage) the greater the variability in the measured variables and in the response to hemorrhage.

#### CONCLUSION

Cardiovascular and metabolic responses are altered qualitatively and quantitatively by the method of restraint during hemorrhage in conscious swine.

#### RECOMMENDATION

Our findings suggest that it is important to evaluate the physiologic responses of swine in different types of restraint before gathering data on cardiovascular and metabolic responses.

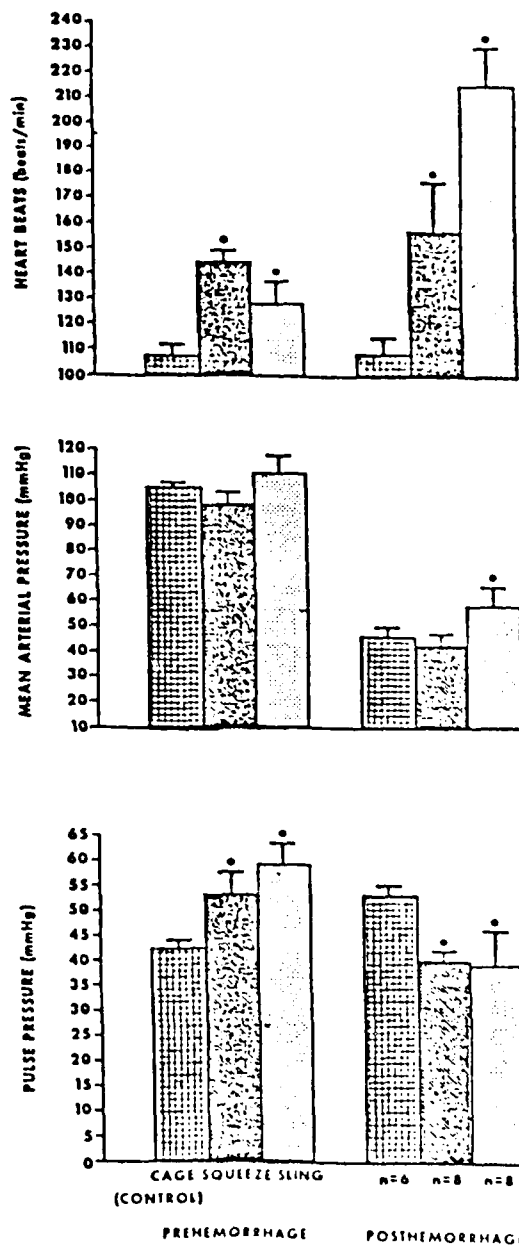


FIGURE 1: Heart rate (beats/min), mean arterial pressure (mmHg), and pulse pressure (mmHg) of holding cage (control) animals (n=6), squeeze cage animals (n=8), and sling animals (n=8), before and after hemorrhage. (\*) indicates significant difference between groups of animals as compared to control animals. Note, (\*) does not indicate significance across groups of animals, i.e., groups of animals before hemorrhage are not compared to groups of animals after hemorrhage.

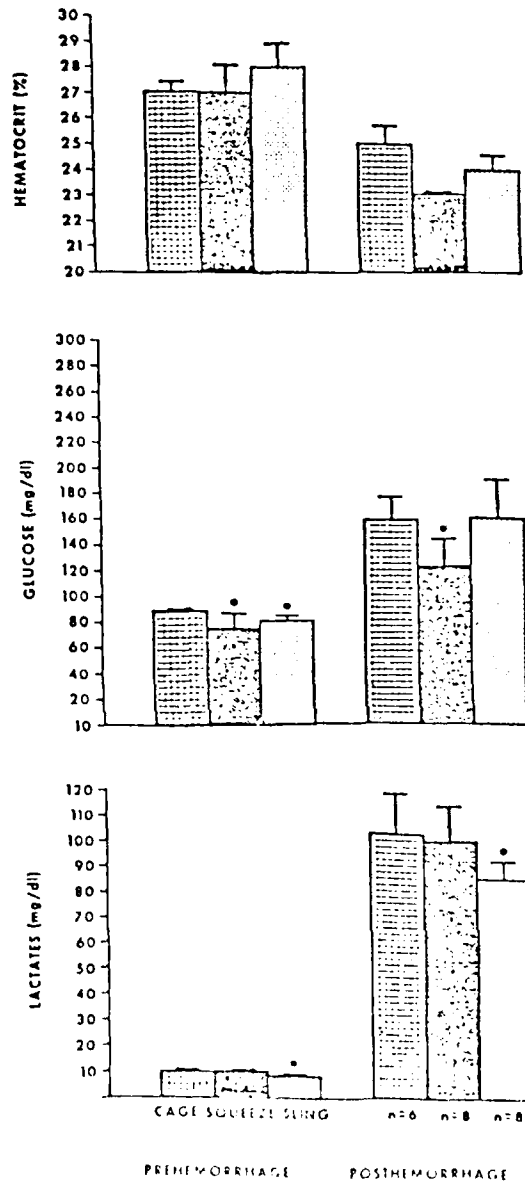


Figure 2: Hematocrit (%), glucose (mg/dl), and lactate (mg/dl) for holding cage (control) animals, squeeze cage animals, and sling animals before and after hemorrhage. (\*) indicates significant difference between groups of animals as compared to control animals. Note, (\*) does not indicate significance across groups of animals, i.e., groups of animals before hemorrhage are not compared to groups of animals after hemorrhage.

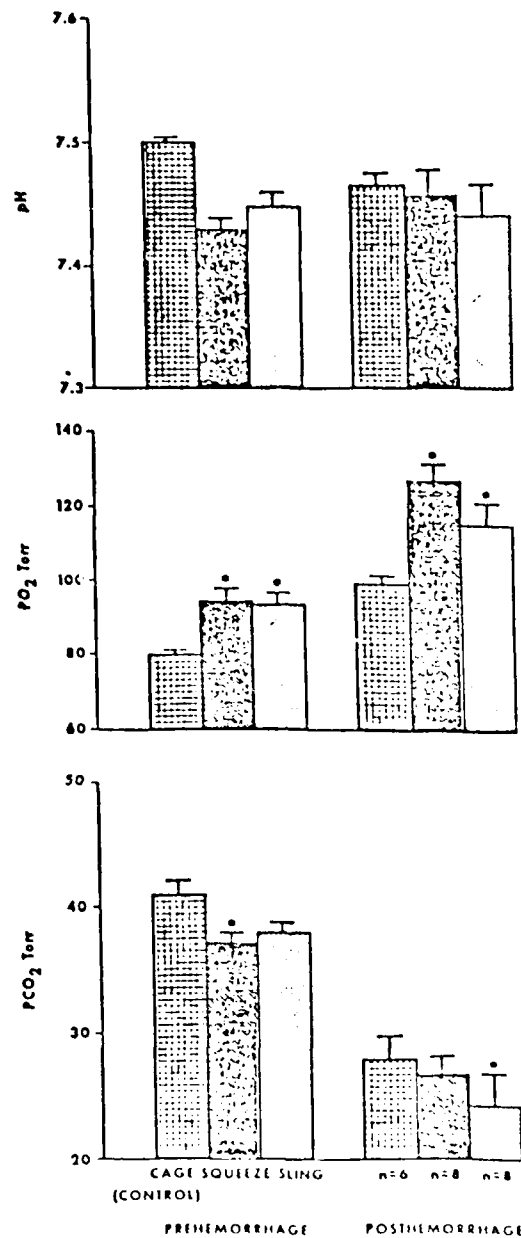


Figure 3: The pH,  $PO_2$  (torr), and  $PCO_2$  (torr) of holding cage animals, squeeze cage animals, and sling animals before and after hemorrhage. (\*) indicates significant difference between groups of animals as compared to control animals. Note, (\*) does not indicate significance across groups of animals, i.e., groups of animals before hemorrhage are not compared to groups of animals after hemorrhage.

BIBILOGRAPHY

1. Hannon, JP, Domestic swine in physiologic research, I. A biomedical model. Presidio of San Francisco, California: Letterman Army Institute of Research, 1981; Institute Report No. 91.
2. vonEngelhardt W, Swine in cardiovascular research--a review. In: Swine in Biomedical Research, edited by L.K. Bustad and R.O. McClellan. Seattle: Frayn Printing, 1966. pp 307-329.
3. Wade CE, Hannon JP, Bossone CA, Hunt MM, and Rodkey WG, Cardiovascular and hormonal responses of conscious pigs during physical restraint. In: Swine in Biomedical Research, edited by M.E. Tumbleson. New York: Plenum Press (In Press).
4. Traverso LW, An aortic sideport catheter for rapid hemorrhage in unheparinized swine. Presidio of San Francisco California: Letterman Army Institute of Research, 1984; Institute Report No. 168.
5. Hannon JP, and Bossone CA, The conscious pig as an animal model for studies of hemorrhagic hypotension. In: Swine in Biomedical Research, edited by M.E. Tumbleson. New York: Plenum Press (In Press).

END

2-87.

D TIC