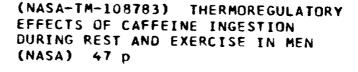


# Thermoregulatory Effects of Caffeine Ingestion During Rest and Exercise in Men

Nancy Dunagan, J. E. Greenleaf, and C. J. Cisar

January 1994



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Ames Research Center Moffett Field, California 94035-1000

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#### INTRODUCTION

# Acknowledgments

I would like to thank Dr. Craig Cisar of San Jose State University for the time he donated during the conception and execution of this research. In addition, Dr. John Greenleaf, Ames Reasearch Center, provided valuable guidance and necessary equipment for this research.

#### **Background**

Caffeine has long been used for its alleged capability to uplift attitude, decrease fatigue, and increase work capacity (Rall, 1985). Only 50 mg of caffeine evokes physiological responses involving the digestive system, the central nervous system, and the cardiovascular system (Somani and Gupta, 1988). Because of caffeine's possible ergogenic effects on athletic competition, the International Olympic Committee has limited competition to those with plasma caffeine concentrations of less than 15 ug/ml (Rosenbloom and Sutton, 1985).

Caffeine has been reported to either improve endurance capabilities (Chad and Quigley, 1989; Costill, Dalsky, and Fink, 1978; McNaughton, 1986; Sasaki, Maeda, Usui, and Ishiko, 1987) or to have no effect on aerobic performance (Aragyasami, Yang, and Winder, 1989; Bond, Adams, Balkisson, McRae, Knight, Ribbins, and Banks, 1987; Falk et al. 1989; Tarnopolsky, Atkinson, MacDougall, Sale, and Sutton, 1989; Winder, 1986). Athletes continue to take caffeine before competition although data concerning caffeine's effectiveness as an ergogenic aid has been disparate. Because caffeine is so commonly ingested in situations which tax the thermoregulatory system, caffeine's effect on thermoregulation should be addressed.

There are several pharmacological effects of caffeine which could possibly cause shifts in temperature regulation. Caffeine increases urinary water and sodium excretion (O'Neil, Hynak-Hanbison, and Gordan, 1986). Thus caffeine may decrease the available fluid and electrolytes which are necessary for maintenance of body fluid volumes and evaporative and convective cooling. Caffeine also stimulates metabolic rate (Dullo and Miller, 1986; Dullo, Geissler, Horton, Collins, and Miller, 1989; Jequier, 1987; and LeBlanc, Jobin Cote, Samson, and Labrie, 1985). If heat generated from higher metabolic rate is not released through thermoregulatory avenues (convection, radiation, conduction, or evaporation), heat storage (and therefore body temperature) would ultimately be increased.

Caffeine's effect on the cutaneous vascular system is unclear. Several authors (Beckman, 1961; Govoni and Hayes, 1978; Kastrup, 1984) have attributed peripheral vasodilation to caffeine consumption. Cutaneous vasodilation would enhance internal heat dissipation via conduction (Sawka, 1988). However, Robertson and Curatolo (1984) indicated that caffeine induces a pressor response by potentiating the effect of catecholamines on smooth muscle and actually increasing catacholamine release in most individuals. Epinephrine and norepinephrine constrict many arterioles, especially those in the skin (Schottelius and Schottelius, 1978), which would result in decreased heat dissipation by lowering conduction and possibly evaporative heat loss. The stimulating effect of caffeine on the sympathetic nervous system and cardiovascular system may also play an important role in human temperature regulation: "Drugs that affect the cardiovascular and central nervous systems, either directly or via the autonomic nervous system, can have important implications for the maintenance of normal thermoregulation" (Rosenbloom and Sutton, 1985, p. 177).

Caffeine's effect on temperature regulation is beginning to receive scientific attention. Lin, Chandron, and Liu (1980) and Schlosberg (1983) found that caffeine produced hyperthermic responses in rats placed in normal and hot environments. Human subjects experienced less of a temperature decrease in a cold environment when given an ephedrine (1 mg/kg) and caffeine (2.5 mg/kg) mixture, but caffeine's exclusive effect on temperature regulation was unclear (Valerand, Jacob, and Kavanagh, 1989).

Caffeine may have a positive effect, negative effect, or no effect on temperature regulation. Starkenstein (1927) found a xanthine derivative (Diuretin) to increase sweating in a steam environment. This additional water loss could give short-term beneficial evaporative cooling to the body. But xanthines may produce dehydration under long-term heat exposures, ultimately putting stress on the thermoregulatory system. During sitting in heated conditions, Biren, Al-Refaee, Finck, and Paolone (1992) found that skin temperatures and oxygen uptake increased with caffeine ingestion, which would normally result in heat storage (Pandolf, Sawka, and Gonzalez 1988). However, rectal temperature remained constant, most likely due to skin vasodilation, causing higher skin temperatures, thereby increasing heat conductance from the core to the skin.

Caffeine appears to have no effect on thermoregulation during exercise in cold environments (Doubt and Hsieh, 1991; Graham, Sathasivan, and MacNaughton, 1991). Graham et al. (1991) found a dose of 5 mg caffeine/kg body weight produced a significant increase in rectal

temperature when subjects performed sub-maximal exercise during head-out water (28°C) immersion.

Falk et al. (1990) found ingested caffeine produced tendencies toward greater sweat rate (p < 0.07) and higher final rectal temperature (p < 0.068) during treadmill running to exhaustion under normal ambient conditions. Five ml caffeine/kg caused no significant difference in rectal temperature at the completion of a two hour run (Gordon et al., 1982), but 2 of the 10 subjects reached rectal temperatures >40°C.

Athletes often ingest caffeine prior to exercise, either inadvertently or specifically for its alleged ergogenic effects. Since many endurance competitions are performed in environments that tax the body's temperature regulating capabilities, the effect of caffeine on thermoregulation should be studied further.

#### **Purpose**

The purpose of this study was to determine if caffeine ingestion affects body temperatures and thermoregulatory factors such as, skin heat conductance and sweat loss in active men at rest, and when preforming intensive submaximal aerobic exercise under normal ambient conditions.

# **Approach**

After pre-test measurements of height, weight, percentage body fat, and maximal oxygen uptake rate were collected, 12 aerobically conditioned males (20 to 41 years of age) underwent a non-caffeine control trial (NCT) and a caffeine trial (CT) (10 mg caffeine per kg body weight) in normal ambient conditions. Rectal temperature, skin temperatures, oxygen uptake, and heart rate where recorded during 30 minutes of rest followed by 70 minutes of sitting cycle ergometry exercise at approximately 68% of VO<sub>2</sub> max. Total weight loss was determined from the subjects pre-drink weight (plus weight of ingested liquid) and their weight taken directly after the post-exercise cool-down period. Skin heat conductance, mean body temperature, mean skin temperature, and total sweat loss were calculated. Mean 10-minute values were analyzed using repeated measures two- way analyses of covariance. Weight loss and sweat loss were analyzed using dependent t-tests. The null hypothesis was rejected when p < 0.05.

#### **Null Hypotheses**

There will be no significant differences found between the caffeine and non-caffeine trials in rectal temperature, mean skin temperature, skin heat conductance, or total sweat loss.

#### **Delimitations**

This study was limited to 12 healthy male volunteers from 20 to 41 years. The subjects had less than 18% body fat and all had been exercising aerobically at least three times per week for 30 minutes per session prior to testing.

#### Limitations

Factors in this study that could not be controlled included: (1) variations in the genetic makeup of the subjects and their reaction to caffeine; (2) slight temperature and humidity variations within the testing room; and (3) changes in the fitness level of the subjects during the testing period.

#### **Assumptions**

Assumptions made for this research included: (1) subjects gave their best effort during the maximum oxygen uptake test and during the two test trials; (2) subjects complied with dietary restrictions; (3) subjects maintained their fitness level throughout the two week experimental period; and (4) subject's heat acclimation status did not change during the study.

#### **Definitions**

The following definitions will be used:

Catecholamines: The class of chemicals which includes epinephrine and norepinephrine (Astrand and Rodahl, 1986).

Conduction: Heat exchange between two solid surfaces that are in direct contact. The rate of conductance is directly related to the temperature differences between the two surfaces and the conductivity of the materials, and inversely related to the distance the heat must travel (Santee and Gonzalez, 1988).

Convection: Heat exchange that occurs between a surface and a fluid (Santee and Gonzales, 1988). Skin and air are involved in convection in normal human environmental conditions.

**Diaphoretic:** An adjective meaning "causing perspiration" (Stein, 1982).

**Epinephrine**: A hormonal substance released from the adrenal medulla when the body experiences pain, cold, hypotension, hypoglycemia, or hypoxia. Epinephrine

causes constriction of many arterioles especially in the skin, mucous membranes, and kidneys; but causes dilation of the coronary system, skeletal muscles, and lungs (Schottelius and Schottelius, 1978).

Ergogenic: Inclined to increase work output (Astrand and Rodahl, 1986).

Evaporative heat loss: Moist heat exchange which requires evaporation of water. The physical determinant of evaporative heat loss is the water concentration gradient between the body surface and the environment. The physiological limits depend on sweating rate and the body hydration level (Santee and Gonzalez, 1988).

Mean body temperature: The average body temperature calculated as the sum of 80% of the rectal temperature and 20% of the mean skin temperature (Greenleaf and Reese, 1980).

Metabolic rate: Magnitude of heat production for living animals. Metabolic rate is calculated from oxygen uptake. One liter of oxygen consumed corresponds to approximately 20 kJ (4.8 kcal) (Astrand and Rodahl, 1986).

Norepinephrine: The principal chemical transmitter at peripheral sympathetic nerve terminals. Norepinephrine has a vasoconstrictive effect and raises both systolic and diastolic blood pressure (Schottelius and Schottelius, 1976).

Oxygen uptake: The volume of oxygen extracted by the body from the inspired air, and is usually expressed in liters of oxygen per minute (l/min) or milliliters of oxygen per kilogram of body weight per minute (ml/kg/min) (at 0°C, 760mm Hg, dry standard temperature and pressure). If the oxygen content of the body remains constant during the measurements, oxygen uptake is equivalent to the volume of oxygen utilized from the metabolic oxidation of foodstuffs. One liter of oxygen corresponds to 4.7 to 5.05 kcal of energy liberated depending on the proportion of fat and carbohydrate metabolized (Astrand and Rodahl, 1986).

**Pressor**: Causing an increase in blood pressure via vaso-constriction (Stein, Hauck, and Su, 1982).

Radiation: Energy released by one body, transmitted through an intervening medium, and absorbed by another body (Stein et al., 1982). The net radiation balance is determined by calculating the incoming radiation and subtracting the outgoing or emitted radiation (Pandolf et al., 1988).

Rectal temperature: The temperature recorded from the rectum (Pandolf et al., 1988). A probe depth of 16 centimeters was used in this study.

Skin heat conductance: The ratio between the average heat flow to the skin surface and the temperature gradient from the core to the skin surface (Greenleaf and Reese, 1980).

Skin temperature: Mean skin temperature was calculated relative to body surface area using coefficients and temperatures obtained from six skin sites: the upper arm, forearm, chest, back, thigh, and leg. Mean skin temperature ( $\overline{T}_{sk}$ ) was calculated as: 0.06 ( $T_{arm}$ ) + 0.13 ( $T_{forearm}$ ) + 0.19 ( $T_{chest}$ ) + 0.20 ( $T_{back}$ ) + 0.21 ( $T_{thigh}$ ) + 0.21 ( $T_{leg}$ ) (Greenleaf and Castle, 1972).

Vasodilation: Dilation of the blood vessels (Stein et al., 1982).

**Vasoconstriction:** Constriction of the blood vessels (Stein et al., 1982).

# Significance of the Problem

Athletes often ingest caffeine prior to exercise, either inadvertently or specifically for its ergogenic effects. Since many endurance competitions are performed in hot environments that tax an athlete's temperature regulating capabilities, the effect of caffeine on thermoregulation should be understood. The purpose for this study was to determine if caffeine affects rectal and skin temperatures, and thermoregulatory factors such as skin heat conductance and sweat rate in active men at rest and when preforming intensive, sub-maximal aerobic exercise within normal ambient environmental conditions.

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# REVIEW OF LITERATURE

# **Consumption of Caffeine**

Caffeine is ingested freely in today's society. In an excellent review Somani and Gupta (1988) identified many caffeine sources: tea, cocoa, chocolate, cola drinks, approximately 2,000 nonprescription drugs, and over 1,000 prescription drugs. Coffee contains approximately 100 to 150 mg of caffeine per eight ounce cup. Americans average 1.52 cups of caffeine-containing beverages per day; however, up to 10% of America's adult population are believed to consume more than 1,000 mg of caffeine/day.

# Physiological Effects of Ingested Caffeine

Caffeine, one of three known methelated xanthine compounds (Power and Dodd, 1985), is an alkaloid identified structurally as 1,3,7-trimethylxanthine (Jackobson and Kulling, 1989). Methylxanthines can be absorbed via oral, rectal, or parteneral administration (Powers and Dodd, 1985). Only 50 mg of caffeine can evoke gastric secretion and activation of the central nervous and cardiovascular systems (Somani and Gupta, 1988). One gram of caffeine (about 6 to 7 eight-ounce cups of coffee) can increase respiration and cause insomnia, tachycardia, excitement, ringing in the ears, restlessness, and extra systoles (O'Neil et al., 1986). The acute lethal dose of caffeine for adults is 5 to 10 grams (Rall, 1985).

Orally ingested caffeine is rapidly absorbed through the gastrointestinal tract. Elevated 1,3,7-trimethylxanthine levels can be found in the blood 15 minutes after ingestion and are at peak concentrations 30 to 60 minutes later (Jackobson and Kulling, 1989). The liver metabolizes caffeine, leaving about 1% of the caffeine to be recovered in urine (Rall, 1985). The half life of caffeine has been estimated to be 3.5 hours (Kastrup, 1984), 2 to 10 hours (Powers and Dodd, 1985), and 4 to 48 hours (Jackobson and Kulling, 1989). Varying metabolism with age may explain the large variation.

Caffeine crosses the blood-brain barrier freely and may act directly on the vagal, medullary, and vaso-motor centers (Jackobson and Kulling, 1989). Every system of the body controlled by the central nervous system may be affected by caffeine (Somani and Gupta, 1988). Acute pharmacological actions of caffeine include: (1) stimulation of the sympathetic nervous system (Rosenbloom and Sutton, 1985); (2) stimulation of the resting metabolic rate (Jequier, 1987); (3) elevation of plasma epinephrine levels (Bangsbo, Jacobsen, Nordberg, Christensen, and Graham, 1992; Graham and Spriet, 1991); (4) increasing

jejunal water and sodium secretion (O'Neil et al., 1986); (5) stimulation of cardiac and perhaps skeletal muscle function (Powers and Dodd, 1985); (6) elevation of systolic (Robertson and Curatolo, 1984) and diastolic blood pressures (Pincomb, Wilson, Hee Sung, Passey, and Lovallo, 1991); (7) increasing gastric acid secretion (O'Neil et al., 1986); (8) relaxation of smooth muscle (Powers and Dodd, 1985); and (9) mobilization of free fatty acids (Powers and Dodd, 1985).

Three cellular mechanisms have been identified to explain caffeine's pharmacological effects including: (1) the translocation of intracellular calcium; (2) the inhibition of the phosphodiesterase enzyme; and (3) the blocking of adenosine receptors (Powers and Dodd, 1985). Calcium permeability may be altered by caffeine at the muscle's sarcoplasmic reticulum, resulting in increased Ca2+ release. Phosphodiesterase is a catalyst for the conversion of cyclic AMP to 5'AMP. By decreasing phosphodiesterase, caffeine would indirectly elevate cyclic adenosine monophosphate (cyclic AMP) levels which would result in an increased release of glycerol and free fatty acids from triglyceride stores. Adenosine, a component of ATP and nucleic acids, can affect organs via receptors on the cell's surface (Snyder, 1984). Peripherally administered adenosine will constrict bronchi, inhibit platelet aggregation, dilate blood vessels, reduce blood pressure, produce hypothermia, and inhibit the firing of most neurons (Borstel and Wurtman, 1984; Snyder, 1984). By suppressing adenosine, caffeine often causes increases in blood pressure, renin release, catecholamines, and urine output, while simultaneously decreasing bronchial tone (Somani and Gupta, 1988). Because the concentration of caffeine required to inhibit phosphodiesterase is roughly 100 times that of normal caffeine ingestion, it is more likely that caffeine's molecular mechanism is related to the inhibition of adenosine receptors (Borstel and Wurtman, 1984; Fernstrom and Fernstrom, 1984; Snyder, 1984).

# General Factors Involved in Thermoregulation

There are many factors that contribute to human thermoregulation. The First Law of Thermodynamics has been used by Santee and Gonzalez (1988) as a basis for measuring the thermal effects of the environment on living organisms. Heat storage (S) involves a balance between internal heat production from metabolism (M), heat exchange that occurs via evaporation (E), conduction (K), convection (C), radiation (R), and work (W) either leaving the system (–) or by eccentric work (+) done on the system. The heat balance equation (Santee and Gonzalez, 1988) is as follows:

## $S = M - (\pm W) \pm E \pm R \pm C \pm K [W/m^{2}]$

If the factors on the right side of the equation result in a positive sum for heat storage, the body temperature will increase. Alternatively, if the sum for heat storage is negative, body temperature will decrease. Insight into caffeine's effect on thermoregulation may be enhanced by indentifing its specific effect on any of the heat balance factors. For additional information concerning the mechanisms of thermoregulation, readings from Grayson and Kuehn (1977) or Santee and Gonzalez (1988) are recommended.

#### **Effects of Caffeine on Thermoregulation**

Until recently, little attention has been given to caffeine's effects on the thermoregulatory system. Starkenstein (1927) observed that a xanthine derivative (Diuretin) increased diuretic and diaphoretic (sweating) responses. At room temperature and in a steam bath total water loss (renal and extrarenal) were measured at the end of four hours under the following conditions: (1) without water ingestion; (2) with 1 L tap water ingestion; (3) with 1 L of 0.9% NaCl solution ingestion; (4) with 2 g of Diuretin ingestion; and (5) with 1 L of 0.9% NaCl solution + 2 g Diuretin ingestion. Starkenstein concluded that Diuretin functioned as a weak diuretic at normal room temperature, but Diuretin acted as a diaphoretic in the steam bath.

Rectal temperature was monitored in resting rats during exposure to different ambient temperatures (Lin et al., 1980). Adult male rats were placed in either an 8, 22, or 30°C environment and given 20 minutes to attain thermal balance before being injected intraperitoneal or intraventricular with 10 to 20 mg caffeine/kg. Metabolic rate (calculated from oxygen uptake), respiratory evaporative heat loss (change in water vapor tension of the expired air), and body temperatures (rectal, back skin, foot skin, and tail skin temperatures) were measured each minute for 120 minutes after caffeine injection. In the second phase the rats were pre-treatment with either 6-hydroxydopamine (a central catecholaminergic nerve fiber destroyer), phentolamine (blocks alpha-adrenergic receptors), haloperidol (blocks dopaminergic receptors), or propranolol (a B-adrenergic blocker) before caffeine injections. In all environments caffeine caused a dosedependent increase in rectal temperature, which was attributed solely to increased metabolic heat production. Caffeine injections also increased behavioral excitation, cutaneous vasodilation (estimated from higher foot and tail skin temperatures), and diuresis. The hyperthermia due to caffeine injections was attenuated when the rats were pre-treated with 6-hydroxydopamine, phentolamine, and haloperidol. In addition to the blockade of alphaadrenergic and dopaminergic receptors, destruction of central catecholaminergic nerve fibers nullified caffeine's hyperthermic effect. Lin et al. (1980) suggested that "xanthines elicit a central activation of both adrenergic and dopaminergic receptors via release of endogenous noradrenaline and dopamine and lead to behavioral excitation and hyperthermia." The hyperthermic response to caffeine was therefore attributed to increased motor activity, which would result in increased metabolic heat production.

Schlosberg (1983) found that the thermoregulatory response of rats to caffeine was dose dependent and that tolerance to caffeine was rapidly developed. Adult male rats acclimated to an ambient temperature of 22°C were used in three experiments. Each protocol used injections of either water or caffeine (12.5, 25, 50, and 100 mg/kg). Experiments one and three were conducted in room temperatures of 21-23°C; experiment two was conducted in an environmental chamber with temperatures of either 4 or 32°C. In both experiments one and two the acute effects of caffeine on thermoregulation were measured, and in experiment three the effects of daily caffeine administration on thermoregulation over 28 days were measured. Results from experiment one indicated acute caffeine administration produced dose dependent changes in body temperature. The lower caffeine doses (12.5 and 25 mg/kg) were followed by hyperthermia. Peak significant temperature increases of ~1°C occurred 30 to 60 minutes after caffeine injection, and were no longer significantly different after 2 to 2.5 hours. Conversely, both higher caffeine doses (50 and 100 mg/kg) induced hypothermia: Maximal decreases in temperature between 1 to 1.5°C occurred two hours after caffeine injection and 3.5 hours after injection a 1°C deficit was still observed. When the rats were put in different environmental conditions their body temperature responses to caffeine were altered. In the 32°C environment there was no hypothermic response to 50 and 100 mg/kg of caffeine. Although the hyperthermic reactions to 12.5 and 25 mg/kg of caffeine were less pronounced, there were still significant temperature elevations when compared to the control group 60 and 90 minutes after injection. When placed in the 4°C environment the rats injected with 50 and 100 mg/kg caffeine continued to have significantly lower body temperatures than the control group. While in the cold environment the 50 mg/kg dose produced a similar declining body temperature pattern, as was observed in the normal ambient environment; the 100 mg/kg dose produced a drop of almost 4.5°C in body temperature after 3.5 hours. Neither of the lower doses of caffeine produced significant changes in body temperature when the rats were exposed to the cold environment. Caffeinedependent alterations in body temperature changed with

duration of exposure to the drug. Tolerance to the hypothermic reaction to caffeine developed rapidly: By the fourth day the high caffeine groups displayed body temperatures no different from the control group, and after seven days the 50 mg/kg group showed a hyperthermic effect which continued for the remaining 28 days. Rats receiving both of the lower caffeine doses continued to display a hyperthermic response throughout the duration of the experiment. These results were attributed to: (1) possible direct and/or indirect effects on the central and/or peripheral mechanisms involved in thermoregulation; or (2) a secondary effect of physiological and behavioral processes that is unrelated to temperature regulation. This well designed study provided strong evidence that the thermoregulatory response of rats to caffeine is dose and tolerance dependent.

The effect of caffeine ingestion on thermoregulatory response during endurance running was observed by Gordon et al. (1982) in 10 male subjects who had trained together for 17 days. Caffeine consumption was prohibited for 24 hours before the test. The morning of the test all subjects ingested the same breakfast three hours before the run and, later, five subjects were given 5 mg of caffeine/kg orally one hour before the run. Rectal temperature was taken before and within two minutes of completion of the two-hour run. The environmental conditions, obtained from a nearby weather station, included dry bulb temperatures ranging from 24.5-28.9°C, wind speed between 17-22 km/h, and relative humidity of 41-54%. All subjects were given 200 ml of cold water every 20 minutes during the run. Oral caffeine had no significant effect on rectal temperature, plasma volume, or electrolyte levels. Upon completion of the run two of the 10 subjects had reached rectal temperatures at the heat stroke threshold (40.6°C). The rate of temperature elevation was not available since rectal temperature was not monitored throughout the run. Environmental temperature, exercise intensity, fitness level, and caffeine tolerance were additional variables not closely monitored in this study.

At the other end of the temperature spectrum the effect of an ephedrine-caffeine mixture on humans exposed to a cold environment was investigated by Valerand et al. (1989). Nine adult male subjects underwent two cold exposure tests (10°C for three hours) directly after ingesting either a placebo or an ephedrine (1 mg/kg) plus caffeine (2.5 mg/kg) mixture. The subjects fasted 12 hours and refrained from alcohol use at least 48 hours prior to the test. Oxygen consumption and CO<sub>2</sub> production were measured during cold exposure in the last 10 minutes of each 30 minute interval. The decreases in core, mean skin, and mean body temperatures were significantly less (p < 0.01) during the ephedrine/caffeine trial when com-

pared with those during the placebo trial. In addition, subjects who received ephedrine/caffeine exhibited greater energy expenditure (18.6%) and greater carbohydrate oxidation (41.5%) compared with placebo results. Heat loss during both trials was similar. Valerand et al. suggested the beneficial effects of the ephedrine/caffeine mixture were due exclusively to greater energy production via enhanced carbohydrate utilization. The individual effects of caffeine or ephedrine on cold responses have not been established.

Several investigators have addressed caffeine's effect on thermoregulation recently. Falk et al. (1990) investigated caffeine's effect on body fluid balance and thermoregulation during treadmill exercise (70-75% VO2 max) performed in a neutral environment (25°C, 50% RH). Seven male subjects underwent a placebo trial and a caffeine trial. A dosage of 5 mg caffeine/kg body weight was given 120 minutes prior to exercise, plus an additional 2 mg caffeine/kg body weight was given 30 minutes prior to exercise. Serum caffeine levels during the caffeine trial were  $10.74 \pm 0.98$  ug/ml just prior to exercise,  $13.07 \pm 0.98$  ug/ml after 15 minutes of exercise, and remained at that level throughout the remainder of the test. Time to exhaustion was not significantly different between the placebo and caffeine trials  $(61.60 \pm 9.99)$  and  $63.91 \pm 6.42$  min, respectively). Mean total water loss  $(1,376 \pm 154 \text{ vs } 1,141 \pm 158 \text{ ml, respectively})$ , sweat rate  $(12.4 \pm 1.1 \text{ vs } 10 \pm 0.7 \text{ g/m}^2/\text{min}, \text{ respectively}), \text{ rise in}$ rectal temperature (2.1  $\pm$  0.3 versus 1.5  $\pm$  0.4 °C, respectively), and calculated heat storage (134.4  $\pm$  17.7 versus  $93.5 \pm 22.5$  W, respectively) were not significantly different. Final rectal temperature, although not significantly different (p = 0.068), tended to be higher in the caffeine trial. Caffeine may hinder thermoregulation under higher dosages or more stressful conditions because of the observed tendency toward greater water loss with caffeine.

The influence of caffeine and exercise on catacholamines and metabolism was observed in six men during two hours of 5°C cold air exposure by Graham, Sathasivam, and MacNaughton (1991). Either a placebo or caffeine (5 mg/kg body weight) capsule was ingested 30 minutes before subjects were exposed to exercise (60 W) in the cold or exposed to sitting at rest in the cold. Rectal temperature, skin temperatures, oxygen consumption, carbon dioxide production, and venous blood samples were taken prior to caffeine or placebo ingestion, and at 5, 30, 60, 90, 120 minutes into cold exposure. Caffeine caused a significantly greater increase (by 86%) in plasma epinephrine over the two hour period when compared to the placebo trials; rectal and skin temperatures were not significantly affected by caffeine. Because there was no decrease in resting core temperature, environmental conditions may

not have been cold enough to produce observable temperature differences.

Another study concerning caffeine's effect on thermoregulation in a cold environment was carried out by Doubt and Hsieh (1991). Four trials were completed using caffeine (5 mg/kg body weight) or a placebo in both cold water (18°C) and warmer water (28°C) to observe the possible additive physiological effects of caffeine and cold water during head out immersion ergometer exercise (1.5 W/kg body weight). Ten male subjects ingested either caffeine or the placebo 90 minutes prior to exercise and immersion. Data were collected starting with immersion and continuing throughout 5 minutes of rest, 5 minutes of warm up, 55 minutes of exercise, and 15 minutes of recovery. Caffeine increased plasma free fatty acids (by  $52 \pm 18\%$ ), glycerol (by  $14 \pm 8\%$ ), lactate (by  $28 \pm 10\%$ ), as well as oxygen uptake (by  $9 \pm 3\%$ ), minute ventilation (by  $7 \pm 5\%$ ), heart rate (by  $4 \pm 1\%$ ), and rectal temperature (by  $2.0 \pm 0.4\%$ ), but had no effect on respiratory exchange ratio in the 28°C environment. The increase in rectal temperature was greater (p < 0.01) with caffeine ( $\pm 0.78 \pm 0.09$ °C) than the placebo  $(+0.55 \pm 0.15$ °C) trials during the 28°C exposure. Rectal temperature did not vary significantly during either the caffeine or placebo exposure during the 18°C exposure. Mean skin temperature tended to increase with caffeine by 0.1 to 0.2°C during both environmental conditions. Caffeine and cold did not have significant additive effects on thermoregulation; but, as stated earlier, caffeine stimulated an increase in rectal temperature during the 28° exposure. The gradient for heat loss was low in the 28°C environment compared with heat production, and vasodilation was insufficient to prevent the increase in core temperature.

Caffeine's effect on body temperature during exposure to a heated (40°C, 70% rh) and neutral (23°C, 40% rh) environment studied in eight male volunteers who took either a placebo, a 4 mg/kg dose of caffeine, or an 8 mg/kg dose of caffeine 60 minutes prior to the experimental period which was 60 minutes sitting at rest in the neutral or hot air environments (Biren et al., 1992). Oxygen uptake was significantly higher (p < 0.05) for both caffeine trials when compared to the placebo trial, with environment having no effect. Mean skin temperature was significantly higher (p < 0.05) for both the 4 mg/kg caffeine trial

 $(34.18 \pm 0.40^{\circ}\text{C})$  and the 8 mg/kg caffeine trial  $(34.08 \pm 0.41^{\circ}\text{C})$  when compared to the placebo trial  $(33.84 \pm 0.28^{\circ}\text{C})$ . Body core and mean body temperatures were unaffected by caffeine. It was concluded that "the caffeine induced increase in skin temperature reflected an increased conduction of heat from core to shell allowing rectal temperature to be maintained in spite of the drug induced metabolic increase" (Biren et al., 1992).

#### **Summary**

A summary table of selected caffeine studies is found in appendix H.

There are several pharmacological effects of caffeine on the sympathetic nervous and cardiovascular systems (Rosenbloom and Sutton, 1985) which could cause shifts in temperature regulation; caffeine increases water and sodium excretion O'Neil et al. (1986) and may cause dehydration and compromise evaporative and convective cooling. Caffeine can stimulate resting metabolic rate (Dullo and Miller, 1986; Dullo et al., 1989; Jequier, 1987; LeBlanc et al., 1985) which, if not released through increased convection, radiation, conduction, or evaporation, the body heat storage (and therefore body temperature) would ultimately increase.

Caffeine increases resting metabolic rate (Dullo and Miller, 1986; Dullo et al., 1989; LeBlanc et al., 1985) and produces dose dependent temperature changes in rats (Lin et al., 1980; Scholsberg, 1983). In human subjects xanthines: (1) increase sweating in steam bath conditions (Starkenstein, 1927); (2) increase skin temperature during heat exposure (Biren et al., 1992); (3) increase rectal temperature during immersion in 28°C water (Graham et al., (1991); (4) have no effect on thermoregulation during cold exposure (Doubt et al., 1991; Graham et al., (1991); (5) have no thermoregulatory effect in normal or hot ambient environments during rest (Biren et al., 1992); and (6) apparently have no thermoregulatory effects during exercise (Falk et al., 1990; Gordon et al., 1982).

Studies on humans incorporating adequate controls in a normal environment, and when the thermoregulatory system is stressed (as during exercise), are necessary to determine possible effects of caffeine on human temperature regulation.

#### **METHODS**

### **Subjects**

The experimental protocol was approved by the San Jose State University Human Subjects Institutional Review Board. Subjects were solicited verbally and through posted advertisement (appendix C). Twelve male volunteers were tested, but data from one of the volunteers were not included due to difficulties with temperature measurements. Each subject was screened via questionnaire (appendix A) for age, sex, medical history, family health history, addictive habits, and exercise- training routines. Only those considered "apparently healthy" by the standards found in the American College of Sports Medicine's (ACSM) Guidelines for Exercise Testing and Prescription (4th edition, 1991) were used in this study. Since fat provides thermal resistance (Toner and McArdle, 1988), subject's fat content was required to be below 18% fat (as measured by underwater weighing). All subjects were non-smokers. Six of the subjects consumed less than 100 mg caffeine per day; three subjects consumed 100-200 mg caffeine per day; and two subjects consumed 200-450 mg caffeine per day.

Additionally, only active volunteers (who exercised aerobically for a minimum of 30 minutes at least three times per week) were selected, since fitness level alters caffeine's effect on resting metabolic rate (LeBlanc et al., 1985). Volunteers were informed of the procedures and the possible risks of participating in this research, and informed consent (appendix B) was obtained before initiation of testing.

#### **Procedure and Methods**

Eleven of 12 subjects completed three testing sessions. First, in the pre-test session the subject's weight, height, body fat percentage, and maximum oxygen uptake were measured. Using a single-blind method, the subjects then per formed caffeine and non-caffeine exercise sessions. Both exercise sessions were completed within two weeks of the pre-test, and the first exercise session took place at least two days after the pre-test, and the second exercise session took place a minimum of two days after the non-caffeine exercise session or a minimum of four days after the caffeine exercise session.

#### Pre-test

Subjects were instructed not to eat or drink three hours prior to testing (ACSM, 1991), and to not exercise for 12 hours before the pre-test. Data were recorded on a

pre-test form (appendix C) from measurements taken in the following order: height, weight, residual lung volume, body composition (relative fat, fat weight, and fat free weight), and maximum oxygen uptake.

#### Height

A stadiometer was used to measure the subjects' height. The bare-footed subject stood (heels together) with his heel, buttocks, scapula, and cranium against a vertical pre-measured surface. With head held upright, arms to the side, and weight distributed evenly on both feet, the subject inhaled deeply and maintained an upright position as the technician brought the headboard to the most superior part of the subject's scalp. The subject then stepped away as the technician held the headboard in position and measured the subject's height to the nearest 0.1 cm (Christensen, 1990).

#### Weight

Subjects wore only shorts as they were weighed on a calibrated Accu-Weight Scale (San Francisco, California). With weight evenly distributed on both feet the subject stood in the center of the platform and the technician added or removed weights, and then moved the slide of the scale until the beam was centered. Weight was recorded to the nearest 0.05 kg (Christensen, 1990).

#### **Maximal Oxygen Uptake**

Maximal oxygen uptake was determined using a continuous progressive exercise method on a Monark 818 cycle ergometer at a pedal cadence of 60 revolutions per minutes (rpm). The ergometer seat was adjusted to allow the subject near full leg extension, and the handle bar was adjusted to a comfortable level. Before the test began the subject was informed about the testing procedure, given an opportunity to ask questions, and a 10-second resting electrocardiogram strip was obtained. Throughout the test respiratory parameters and heart rate were monitored continuously.

After a five-minute warm-up at a power output of 180 kilopond meters per minute (kpm/min), the resistance was increased by 180 kpm/min each minute until the subject was unable to maintain 60 rpm. The test was terminated if: (1) the subject did not elect to or could not continue; (2) the equipment failed; (3) the ECG showed abnormalities; and/or (4) signs of distress were indicated by loss of coordination (ACSM, 1991). Upon completion of the maximal test the subject was given a three to five minute exercise cool-down period in which the heart rate

declined to below 120 beats per minute. Before removing the electrodes, a final ECG strip was obtained.

The subject's heart rate was measured using a standard V5 electrode lead placement. Electrodes were connected to a type 410 physiological monitor (Tektronix Incorporated; Portland, OR) interfaced to a model 462 strip chart recorder (Thornton Associates Incorporated; Waltham, MA). An electrocardiogram (ECG) was printed (paper speed of 25 mm per minute) for the last 10 seconds of each minute during the test.

A headpiece attached to a Hans-Rudolph (Kansas City, MO) respiratory valve and mouthpiece was fitted to the subject and a nose clip was attached. Inhaled air entered through a Parkinson-Cowan CD-4 dry test meter (New York, NY) and through the respiratory valve. A potentiometer, interfaced with an Apple II computer, recorded the volume of inspired air. Expired air passed through the Hans-Rudolph valve into a mixing chamber, then through a Wilmore-Costill spinner Valve System (WCVS) into two gas analyzers. The oxygen (FeO2) and carbon dioxide (FeCO<sub>2</sub>) concentrations of expired air were measured with a Beckman LB-2 Medical Gas analyzer (Fullerton, CA) and an Applied Electrochemistry 5-3A analyzer (Sunnyvale, CA), respectively. The gas analyzers were calibrated before and periodically during each test by the same technician. Both analyzers were connected to an Apple II computer. Data from inspired air and expired FeO2 and FeCO2 concentrations were calculated from one minute samples. Expired ventilation (VE) was calculated from the inspired ventilation rate (V<sub>1</sub>), oxygen uptake (VO<sub>2</sub>), and carbon-dioxide production (VCO2); respiratory exchange ration (RE) was calculated from  $\dot{V}$ CO<sub>2</sub>/ $\dot{V}$ O<sub>2</sub>.

#### **Body Composition**

Body composition was determined by underwater weighing with correction for residual lung volume. Using the oxygen dilution method, residual lung volume was determined by averaging the two closest measurements (within ±2%) collected from a model 505 Nitralyzer nitrogen gas meter (Med Science, St. Louis, MO). The same technician calibrated the Nitralyzer prior to each test.

To measure weight underwater the subjects wore a weighted belt and sat in a webbed sling suspended from a Chatillon scale (New York, NY). Six to 10 trials were given where the subject exhaled as much air as possible when completely immersed. The average of three trials (within ±0.05 kg) were used for the underwater weight (Cisar et al., 1987). Body composition was calculated using the equation developed by Brozek, Grande, Anderson, and Keys (1963).

#### Caffeine/Non-Caffeine Sessions

The two exercise sessions were performed similarly except the subjects were given either a non-caffeinated or a caffeinated drink. To control for possible effects of circadian rhythms on body temperature, both caffeine/non-caffeine sessions were conducted in the morning at approximately the same time (Stephenson and Kolka, 1988). Subjects wore brief shorts, shoes, and ankle socks. Back-up data were recorded on separate data collection sheets (appendix F).

The following paragraphs will present information on: pre-exercise test requirements; caffeine administration; environmental conditions; pre- and post-test weights; resting rectal temperature, mean skin temperatures, heart rate and oxygen uptake; exercise procedures; and exercise rectal temperature, mean skin temperatures, heart rate and oxygen uptake. Body surface area, mean skin temperature, mean body temperature, respiratory water loss, sweat loss, and skin heat conductance formulas are included within the temperature sections.

#### **Pre-Exercise Session Requirements**

To eliminate any previously developed caffeine tolerance, before both exercise sessions the subjects were given a list of caffeine containing substances (appendix D) and asked not to eat or drink any caffeine containing substances for at least four days prior to their tests (Fisher, McMurray, Berry, Mar, and Forsythe, 1986). To help normalize hydration levels, the subjects were asked to refrain from alcohol 24 hours prior to testing, to not eat or drink eight hours prior to testing, and to abstain from strenuous exercise 24 hours before each exercise session.

#### **Caffeine Administration**

Both exercise sessions were performed in an identical manner except subjects were given either a control mixture or a caffeine mixture. The control mixture contained Sugar Free Crystal Light (Kraft General Foods, Inc., White Plains, NY) dissolved in 237 ml (8 ounces) of water (37°C). The caffeine test used the same mixture with caffeine (Chemicals for Research and Industry, Martinez, CA) at a dosage of 10 mg/kg body weight. To control for psychosomatic effects related to learning or expectation, the subjects were not told which drink they consumed, but several commented on the bitter taste of caffeine. Five of the 11 subjects completed the caffeine session first.

#### **Environmental Conditions**

The environmental conditions, including dry bulb and wet bulb temperatures and barometric pressure, were recorded prior to and after the caffeine/non-caffeine sessions. Rectal temperature levels during exercise are independent of ambient air temperatures between 5° and 30°C (41° and 86°F) (Gonzalez, Berglund and Gagge, 1978; Lind, 1963; Nielsen, 1970; Stolwijk, Saltin and Gagge, 1968). Ambient air temperatures during this study  $20.7 \pm 0.8$ °C. A fan was turned on in front of the subjects during exercise to enhance sweat evaporation and to minimize dripping. Wind speed was measured with a hot wire anemometer and recorded three times during each session.

#### Pre- and Post-Session Weight

The same technician weighed the nude subject before he drank either mixture. The same equipment and procedures were used as in the pre-test session. The weight of the drink was added to the pre-exercise body weight. Then, immediately after the cool-down period, all equipment was removed from the subject and the remaining sweat wiped off before the final weight was taken.

#### **Resting Measurements**

Equipment required for data collection were attached to the subject after weighing. Baseline heart rates were recorded prior to drinking. Electrode and thermistor placement are shown in figures 1 and 2. The experimental set up is shown in figure 3. Drinks were ingested within one minute, then session time started. Rectal temperature and skin temperatures were recorded every minute, heart rate,  $\dot{V}O_2$ , and  $\dot{V}CO_2$  were recorded the first, second, and third minutes, and every eighth, ninth, and tenth minute of each 10-minute cycle thereafter.

#### **Rectal Temperature**

After being weighed the subject inserted a rectal probe (Yellow Springs Instrument Company; Yellow Springs, OH) 16 cm as indicated by a mark on the probe. The rectal probe was connected to a Squirrel Meter-Logger (Science Electronics Inc., Miamisburg, OH). Data collected on the Squirrel were downloaded and processed on a IBM compatible Leading Edge model "D" computer (Leading Edge Hardware Pro., Inc., Canton, MA). Rectal

temperature ( $T_{re}$ ) was recorded to the nearest 0.01°C every minute.

#### Skin Temperatures

Thermistors were attached to the skin with a prototype model of the YSI model PH09 probe holders which allowed for free air circulation. Thermistors were connected to the a Squirrel Meter-Logger (Science Electronics Inc., Miamisburg, OH). Data collected on the Squirrel were down-loaded and processed on a IBM compatible Leading Edge model "D" computer (Leading Edge Hardware Pro., Inc., Canton, MA). Attachment sites for the thermistors were on the right side in the following positions: (1) arm-over the deltoid insertion on the humerus: (2) forearm-over the belly of the extensor carpi radialis longus; (3) chest-3 cm over nipple; (4) backdirectly posterior to the chest probe; (5) thigh-over the middle of the vastus lateralis; and (6) leg-on the lateral side of the belly of the gastrocnemius (J. E. Greenleaf, per sonal communication). Mean skin temperature ( $\overline{T}_{sk}$ ) was calculated as: 0.06 (arm) + 0.13 (forearm) + 0.19(chest) + 0.20 (back) + 0.21 (thigh) + 0.21 (leg temperature) (Greenleaf and Castle, 1972).

#### Oxygen Uptake

Oxygen uptake, carbon-dioxide production, expired ventilation, and respiratory exchange ratio were collected and measured using the same techniques and equipment as during the maximal oxygen uptake rate. They were recorded the first, second and third minutes, and the eighth, ninth, and tenth minutes of each 10-minute cycle thereafter.

#### **Body Surface Area**

Body surface area (SA in m<sup>2</sup>) was calculated with the DuBois formula (Greenleaf and Reese, 1980):

$$SA = 71.84 [(wt)^{0.425} \times (ht)^{0.725})] \times 10^{-4}$$
 (1)

#### Mean Body Temperature

Mean body temperature ( $\overline{T}_{mb}$  in °C) was calculated (Greenleaf and Reese, 1980):

$$\overline{T}_{mb} = 0.8 (T_{re}) + 0.2 (\overline{T}_{sk})$$
 (2)

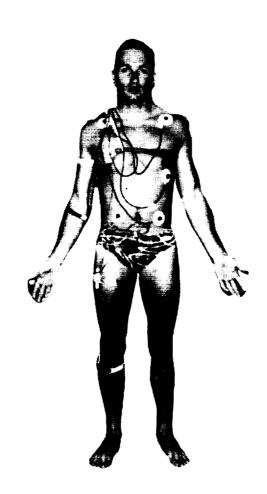


Figure 1. Electrode and thermistor placement.

Figure 2. Thermistor placement.

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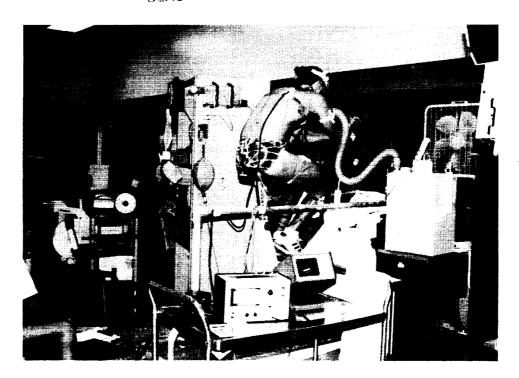


Figure 3. Experimental protocol.

#### **Total Water Loss**

Total water loss (H<sub>2</sub>O<sub>tot</sub> in g/m<sup>2</sup>/hr) was calculated (Ekblom, Greenleaf, Greenleaf, and Hermansen, 1971):

$$H_2O_{tot} = \{(total \text{ wt loss/SA}) - [118.614 (\dot{V}_{CO2}) - 85.742 (\dot{V}_{CO})]\} \times Vm^2$$
 (3)

#### Respiratory Water Loss

Respiratory water loss ( $H_2O_{res}$  in  $g/m^2/hr$ ) was calculated (Ekblom et al., 1971):

$$H_2O_{res} = \dot{V}E_{BTPS} [(H_2O_{do} - H_2O_{dar}) \times 0_a] \times 60/m^2$$
(4)

where:

H<sub>2</sub>O<sub>do</sub> = water density at mean oral temperature

determined prior to liquid consumption and

just after exercise completion

H<sub>2</sub>O<sub>dar</sub> = water density at ambient room temperature

0<sub>a</sub> = relative humidity at ambient room

temperature

#### **Sweat Loss**

Sweat loss ( $\dot{M}_{sw}$  in g/m<sup>2</sup>/hr) was calculated (Ekblom et al., 1971):

$$\dot{M}_{sw} = (H_2O_{tl} - H_2O_{rl}) - H_2O_{il}$$
 (5)

where:

H<sub>2</sub>O<sub>tl</sub> = total weight loss (in grams)

 $H_2O_{rl}$  = respiratory water loss

 $H_2O_{il}$  = insensible water loss (18g/m<sup>2</sup>/hr)

#### **Skin Heat Conductance**

Mean skin heat conductance ( $\overline{H}_{sk}$  in kcal//m<sup>2</sup>/h/°C) was calculated (Greenleaf and Reese, 1980):

$$\overline{H}_{sk} = \frac{M_{gross} - E_{res} - C_{res} - W - (+S)}{SA (\overline{X} T_{re60 - 70} - \overline{X} \overline{T}_{sk60 - 70})}$$
(6)

where:

M<sub>gross</sub> = total heat production from oxygen uptake (kcal/h)

E<sub>res</sub> = evaporative heat loss from the respiratory tract (kcal/h)

= 0.58 (respiratory water loss)

C<sub>res</sub> = conductive and convective heat loss from the respiratory tract (kcal/h),

assumed to be zero

W = heat loss due to external work (kcal/h)

 $= 0.86 \times (work load in watts)$ 

 $\pm S = \text{change in body heat storage (kcal/h)}$   $= 0.83 \times \text{wt } [0.8(T_{re70-60}) + 0.2 \\ (\overline{T}_{sk70-60})] \times 6 \\ \text{weight (wt) in kilograms}$   $SA = \text{body surface area } (m^2)$   $= 71.84 \times [\text{wt}^{(0.425)} \times \text{ht}^{(0.725)}] \times 10^{-4} \\ \text{height (ht) in centimeters}$   $T_{re60} = \text{rectal temperature after } 60 \text{ minutes } (^{\circ}\text{C})$ 

 $T_{re60}$  = rectal temperature after 60 minutes (  $\overline{X} T_{re60-70}$  = mean of  $T_{re}$  measured at 60, 65, and 70 minutes

 $T_{re70-60}$  = the difference between  $T_{re70}$  and  $T_{re60}$ 

#### **Heart Rate**

Heart rate was measured using a standard V5 electrode lead placement connected to a model 410 physiological monitor (Tektronix Incorporated, Portland, OR) connected to a type 462 strip chart recorder (Thornton Associates Incorporated; Waltham, MA). An electrocardiogram (ECG) was printed (paper speed of 25 mm/min) the last 10 seconds of the first, second, and third minutes, and the last 10 seconds of every eighth, ninth, and tenth minute of each 10 minute cycle thereafter for the duration of the session.

#### **Exercise Procedures**

After a 30-minute rest period the subject, seated on the cycle ergometer, exercised for 70 minutes. Caffeine blood level should have peaked during exercise since caffeine concentrations peak 30 to 60 minutes after ingestion (Jackobson and Kulling, 1989). Since the rise rectal temperature is related to the relative exercise load (Saltin and Hermanssen, 1966), the subject cycled using the same absolute load equivalent to approximately 65% of his previously determined  $\dot{V}$  O<sub>2</sub> max. The exercise load (in kgm/min) was determined from the following equation (Lang, Latin, Berg, and Mellion, 1992):

Load = {  $\dot{V}O_2$  (ml/min) – [(3.5 ml/kg/min  $\times$  kg body weight) + 260 ml/min]}/1.9 ml/min.

A fan was turned on at the start of exercise. To reduce sweat loss via dripping and enhance evaporative heat loss, sweat was wiped to drier areas of the body. Verbal encouragement was given throughout the exercise session.

The test protocol would be terminated early if: (1) the subject did not want to (or could not) continue; (2) the equipment failed; (3) the ECG showed significant

abnormalities indicating signs of distress; (4) if there were signs of distress indicated by loss of coordination (ACSM, 1991); or (5) if rectal temperature reached 40°C. Upon completion of the caffeine/non-caffeine sessions, the subject was given a three to five minute exercise cooldown period to allow heart rate to decline to below 120 beats per minute when a final ECG strip was obtained.

## Physiological Parameters Monitored During Exercise

Exercise rectal temperature, mean skin temperatures, heart rate, oxygen uptake and carbon dioxide production were recorded during exercise employing the same techniques and frequency periods as during the resting period. Body surface area, average skin temperature, mean body temperature, respiratory water loss, sweat rate, and skin heat conductance were calculated from the exercise data.

# Level of Significance

A probability level (p) of 0.05 or less was selected as the level of statistical significance to reject the null hypothesis.

#### **Statistical Analyses**

Although 12 healthy men were tested, data from only 11 were analyzed due to difficulties in collecting accurate temperature data. Mean, standard deviation, standard error of the mean, variance, standard error of the skew. range, minimum, maximum and sum were calculated for each variable to describe the data. Repeated measures two-way analysis of covariance was used to control for possible differences between the caffeine and control sessions at the ninth minute, and to determine changes in the variables measured across time and to compare differences of effects of the two treatments (control or caffeine) on the dependent variables. If significant interaction was present between the caffeine and control sessions, data were plotted to observe where the interactions occurred. Summary tables for each of the two-way analysis of covariance tests are presented in appendix G. Repeated measures one-way analysis of variance (ANOVA) and Tukey post-hoc tests were used to determine if significant main effects were found across time in the caffeine or control sessions. Dependent t-tests were used to determine if significant differences existed between the two sessions in total weight loss, weight loss in grams per hour, total water loss, and total sweat loss.

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### RESULTS

### **Findings**

Table 1 summarizes anthropometric and physiological data. Table 2 summarizes environmental conditions present during testing. Weight and sweat loss data collected during the caffeine and control sessions are summarized in table 3, and physiological data collected during the caffeine and control sessions are summarized in table 4.

There were no significant differences between the caffeine and control sessions in total weight loss (p < 0.791), weight loss in grams per hour (p < 0.788), total water loss (p < 0.465), or total sweat loss (p < 0.465).

Means and standard errors for oxygen uptake during the caffeine and control sessions are presented in figure 4. Mean oxygen uptake for the second minute was the average of minutes one, two, and three. There were no significant differences between sessions (p < 0.16), but significant differences across time (p < 0.001), and a significant interaction (p < 0.032) between caffeine and control sessions for oxygen uptake. Resting oxygen uptake was significantly lower than exercise oxygen uptake (p < 0.05), and oxygen uptake for both sessions was very similar during rest and the first 10 minutes of exercise; then the caffeine curve rises slightly above the control session and remains higher until minute 100 where the control session curve rises slightly to become relatively equal to that in the caffeine session (fig. 4).

Table 1. Anthropometric and physiological data

|      | Age | Surface<br>area (m <sup>2</sup> ) | Height (cm) | Fat (%) | Exercise load (kpm/min) | VO2max<br>(l/min) | VO₂max<br>(ml/kg/min) |
|------|-----|-----------------------------------|-------------|---------|-------------------------|-------------------|-----------------------|
| Mean | 29  | 1.96                              | 180.1       | 12.3    | 1088.2                  | 3.73              | 48.7                  |
| ±SD  | 6   | 0.12                              | 4.4         | 3.7     | 153.0                   | 0.46              | 6.2                   |

Table 2. Ambient environmental data

|      | Barometric pressure (mmHg) | Dry bulb<br>temperature<br>(°C) | Relative<br>humidity<br>(%) | Wind speed (m/min) |
|------|----------------------------|---------------------------------|-----------------------------|--------------------|
| Mean | 760                        | 20.7                            | 64.2                        | 156                |
| ±SD  | 1                          | 0.8                             | 4.0                         | 4                  |

Table 3. Weight and sweat loss data during caffeine and control sessions

|                         | Pre-Wt<br>(kg) | Post-Wt<br>(kg) | Wt loss<br>(g) | Sweat loss<br>(g/hr) |
|-------------------------|----------------|-----------------|----------------|----------------------|
| Caffeine<br>Mean<br>±SE | 76.76<br>2.53  | 75.69<br>2.50   | 1,072<br>84    | 378<br>36            |
| Control<br>Mean<br>±SE  | 76.85<br>2.40  | 75.79<br>2.36   | 1,062<br>86    | 393<br>35            |

Table 4. Physiological data during caffeine and control sessions

|          | Exercise VO <sub>2</sub> (I/min) | Exercise $\dot{V}O_2$<br>(% $\dot{V}O_2$ max) | Resting $\dot{V}O_2$ (1/m) | Exercise heart<br>rate<br>(bpm) | Resting heart<br>rate<br>(bpm) |
|----------|----------------------------------|---|----------------------------|---------------------------------|--------------------------------|
| Caffeine |                                  |   |                            |                                 |                                |
| Mean     | 2.55                             | 68.7  | 0.35                       | 145                             | 61                             |
| ±SE      | 0.09                             | 1.7   | 0.02                       | 5                               | 4                              |
| Control  |                                  |   |                            |                                 |                                |
| Mean     | 2.50                             | 67.2  | 0.34                       | 145                             | 64                             |
| ±SE      | 0.09                             | 2.0   | 0.01                       | 7                               | 4                              |

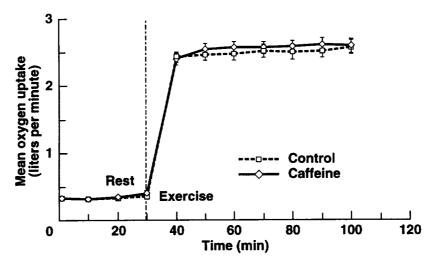


Figure 4. Mean (±SE) oxygen uptake.

Means ( $\pm$ SE) for heart rates during the caffeine and control sessions are presented in figure 5. There were no significant differences between sessions (p < 0.458), significant differences across time (p < 0.001), or significant interaction (p < 0.030) between caffeine and control sessions. Resting heart rates were significantly lower (p < 0.05) than exercise heart rates. Mean heart rate at minute 40 was also significantly lower than mean heart rate at minute 100. Interaction occurred where the caffeine session heart rates were below the control session heart rates to minute 50 (fig. 5).

Means ( $\pm$ SE) for rectal temperatures are presented in figure 6, and differences between the caffeine and control temperatures are presented in figure 7. There were no significant differences between sessions (p < 0.677), significant differences across time (p < 0.001), or non-significant interaction (p < 0.068) between caffeine and control sessions for rectal temperature. Exercise rectal temperature was significantly higher than resting rectal temperature. Rectal temperature was significantly lower during

minutes 40 and 50 than minutes 60 through 100, and rectal temperature at minute 60 was significantly lower than at minutes 70 through 100.

Means ( $\pm$ SE) for skin temperatures are presented in figure 8, and differences in mean skin temperatures for the caffeine and control sessions are presented in figure 9. There were no significant differences between sessions (p < 0.547), non-significant differences across time (p < 0.069), and non-significant interaction (p < 0.886) between caffeine and control sessions for mean skin temperatures.

Figure 10 presents means ( $\pm$ SE) for skin heat conductance. There were no significant differences between sessions (p < 0.232), but significant differences across time (p < 0.001), and non-significant interaction (p < 0.085) between caffeine and control sessions for skin heat conductance. Exercise conductance was significantly higher than resting skin heat conductance; conductance at minute 40 was significantly greater than at minutes 50 and 60.

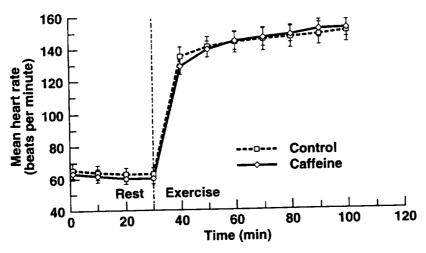


Figure 5. Mean (±SE) heart rate.

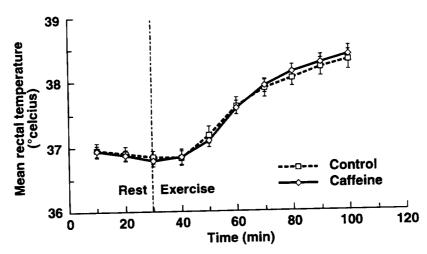


Figure 6. Mean (±SE) rectal temperature.

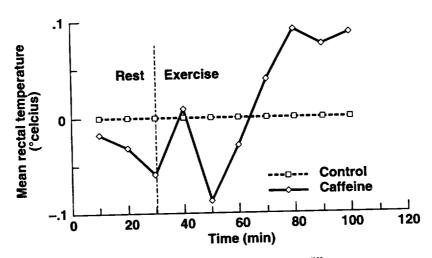


Figure 7. Mean (±SE) rectal temperature differences.

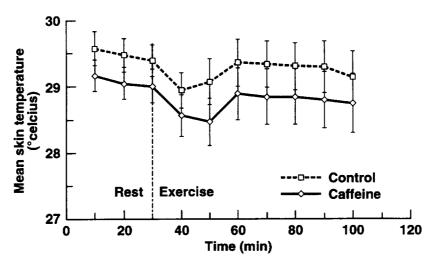


Figure 8. Mean (±SE) skin temperature.

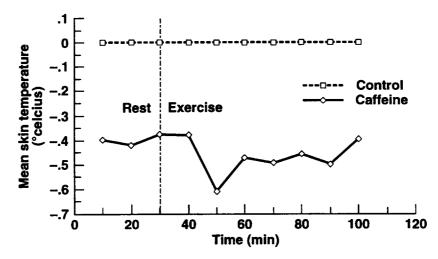


Figure 9. Mean (±SE) skin temperature differences.

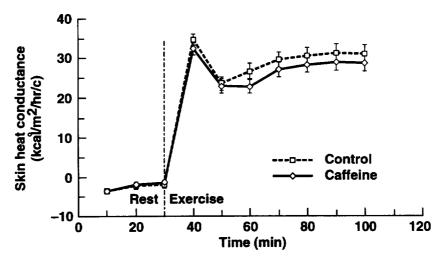


Figure 10. Mean (±SE) skin heat conductance.

#### **Summary**

There were no significant differences (paired t-test) between the caffeine and control session total weight loss (p < 0.791), weight loss in grams per hour (p < 0.788), total water loss (p < 0.465), or total sweat loss (p < 0.465).

Data for repeated measures two-way analysis of covariance for oxygen uptake, heart rate, rectal temperature, mean skin temperature, and skin heat conductance are found in appendix G. Oxygen uptake had no significant differences between sessions (p < 0.16), but significant differences across time (p < 0.001) and a significant interaction (p < 0.032) between caffeine and control sessions. Heart rate had no significant differences between sessions (p < 0.458), but significant differences across time (p < 0.001), and a significant interaction (p < 0.030) for caffeine and control sessions. Rectal temperature had no significant differences between sessions (p < 0.677), significant differences across time (p < 0.001), and a nonsignificant interaction (p < 0.068) for caffeine and control sessions. Mean skin temperature had no significant differences between sessions (p < 0.547), no significant differences across time (p < 0.069), and non-significant interaction (p < 0.886) for caffeine and control sessions. Skin heat conductance had no significant differences between sessions (p < 0.232), significant differences across time (p < 0.001), and non-significant interaction (p < 0.085) for caffeine and control sessions.

Oxygen uptake, heart rate, rectal temperature, and skin heat conductance were significantly lower during the resting phase than during the exercise phase. In addition to these differences across time produced by exercise, heart rates at minute 40 (10 minutes into exercise) were significantly lower than at minute 100 (70 minutes into exercise). Rectal temperatures were significantly lower at minutes 40 and 50 than minutes 60 through 100, and minute 60 was significantly lower than minutes 70 through 100. Skin heat conductance 10 minutes into the exercise phase (minute 40) was significantly greater than 20 and 30 minutes into the exercise phase (minutes 50 and 60).

Significant interactions between caffeine and control sessions were found for oxygen uptake and heart rate. Oxygen uptake for both sessions were similar during rest and the first 10 minutes of exercise; then the caffeine session rose slightly above the control session and remained higher until minute 100 where the control session rose slightly to become relatively equal to the caffeine session. Heart rates for the caffeine session were below control session heart rates to minute 50. At minutes 50, 60, and 70, both sessions had similar heart rates. During minutes 80 and 90 the caffeine session had slightly greater heart rates, and during minute 100 both sessions were similar.

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#### DISCUSSION

The purpose for this study was to determine if caffeine affects rectal temperature, skin temperature, skin heat conductance, or sweat loss in aerobically trained men at rest and when preforming sub-maximal aerobic exercise under normal ambient conditions. Rectal temperatures at rest and during sub-maximal exercise were not significantly affected by caffeine in agreement with results of Gordon et al. (1984), Falk et al. (1990), and Biren et al. (1992). However, it should be noted that at minutes 80, 90, and 100 of the present study, caffeine session rectal temperatures tended (NS) to be 0.08 to 0.09°C higher than those in the control session. There are several possible causes for this trend toward higher rectal temperatures. Oxygen uptake in the caffeine session, although not significantly different, was consistently 0.06 to 0.1 1/min higher during exercise. Accompanying increases in metabolic rate would increase rectal temperature unless the additional heat produced could be dissipated (Santee and Gonzalez, 1988). Skin temperatures were also consistently 0.38 to 0.50°C lower (NS) with caffeine. This trend to lower skin temperatures during the caffeine session indicates that heat was not being released primarily through increased vasodilation of the skin. Possible dehydration in the caffeine session due to increased urine formation (O'Neil et al., 1986) may also have contributed to the slight increase in rectal temperature during the final stages of exercise. Our results do not agree with Doubt and Hsieh (1991) who found that exercise rectal temperature was significantly increased during head-out immersion in 28°C water. The temperature gradient between the skin and water may not have been great enough to facilitate adequate heat transfer. The difference between evaporative heat loss in the present study and heat loss via convection from core to 28°C water plus the large diuresis in immersion are likely explanations for the contradictory results between those of Doubt and Hseih (1991) and the present study.

Mean skin temperatures tended to be lower in the caffeine session for all but the resting temperatures. This trend does not support the findings of Biren et al. (1992) where caffeine increased significantly resting skin temperatures in a normal environment. The reasons for this discrepancy in skin temperatures is not clear because caffeine dosage, environmental conditions, and exercise intensities were similar in both investigations. Because caffeine ingestion increases epinephrine levels acutely (Bangsbo et al., 1992), an enhanced vasoconstrictive effect could explain the small decrease in resting skin temperature in the present study.

Tendencies toward higher rectal temperatures during the last 30 minutes of exercise and lower skin temperatures observed in the caffeine session should have produced lower skin heat conductance. Skin heat conductance during exercise tended to be lower in the caffeine session.

Sweat rate was not affected by caffeine in agreement with Falk et al. (1990) who found that caffeine resulted in a slightly greater sweat rate while, in the present investigation, sweat rate was slightly lower. The small decrease in sweat rate could be explained by possible vasoconstriction caused by elevated epinephrine levels.

The significant differences observed over time for oxygen uptake, heart rate, rectal temperature, and skin heat conductance were normal physiological responses that occur during exercise. Additional convective and evaporative cooling produced by the fan during exercise prevented skin temperatures from increasing significantly.

#### Conclusion

Oral ingestion of 10 mg/kg of caffeine has no effect on rectal temperature, mean skin temperature, skin heat conductance, or sweat rate during moderately heavy exercise in normal ambient conditions.

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### APPENDIX A

## HEALTH AND EXERCISE HISTORY QUESTIONNAIRE

| Jame                                  |                            |   |
|---------------------------------------|----------------------------|---|
| Date                                  | Occupation                 |   |
| ork phone                             | Hom                        | ne phone                                |
|                                       |                            |   |
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| ge years                              | Weightlbs.                 | - · · · · · · · · · · · · · · · · · · · |
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| nes you could be available fo         | r tecting at SISII         |   |
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| • • • • • • • • • • • • • • • • • • • |                            |   |
| edical History                        |                            |   |
| ce a check next to any of the         | conditions that you have   | e experienced.                          |
| Heart attack                          |                            | () Coronary angioplasty                 |
| Cardiac surgery                       |                            | () Chest discomfort                     |
| Rapid heart beats                     |                            | () Heart murmurs () High blood pressure |
| Unusual cardiac finding               |                            | () Ankle swelling                       |
| Stroke                                |                            | () Diabetes                             |
| Abnormal blood lipids                 |                            | () Phlebitis (vein inflammation)        |
| Anemia                                |                            | 02                                      |
| Limp<br>Orthopedic problems, arthrit  | is                         |   |
| Asthma, emphysema or bron             | nchitis                    |   |
| Lightheadedness or fainting           | with exercise              |   |
| Shortness of breath with exe          | rcise                      |   |
| lease explain any of the abov         | e that you have experience | ced and give dates of occurrence.       |
|                                       |                            |   |
|                                       |                            |   |
|                                       |                            |   |

| 0  | Medications of any ty What kind of medicat    |                               | 41%          |   |
|----|---|-------------------------------|--------------|---|
| 0  | =   | alization, or surgical proced |              |   |
| 0  | Drug allergies Explain                        |                               |              |   |
| Fa | mily History:                                 |                               |              |   |
| 0  | Coronary disease<br>Relation, and age of d    | eath                          |              |   |
| 0  | Sudden death<br>Relation, and age of d        | eath                          |              |   |
| 0  | Lipid abnormalities<br>Relation, and age of d | eath                          |              |   |
| Ot | her Dietary Practices:                        |                               |              |   |
| 0  | Caffeine consumption                          | 1                             |              |   |
|    | Coffee  | cups/day                      | cups/week    |   |
|    | Colas   | cups/day                      | cups/week    |   |
|    | Chocolate                                     | bars/day                      | cups/week    |   |
|    | Other   |                               |              |   |
| 0  | Alcohol use Approximately how n               | nuch alcohol do you consur    | ne each day? |   |
| Sm | noking History:                               |                               |              |   |
| 0  | Smoke less than 5 cig                         |                               |              |   |
| 0  | Smoke less than 1 pag                         |                               |              |   |
| 0  | Smoke more than 1 pa                          | аск/дау                       |              |   |
| 0  | Use to smoke but have quit                    |                               |              |   |
| •  |   |                               |              |   |
| 0  | Do you have any eatin                         | ng disorders?                 |              |   |
| Ex | ercise History:                               |                               |              | · |
|    | w often do you exercis                        | e?                            |              |   |
| 0  | 0 to 1 times/week                             |                               |              |   |

- 0 2 times/week
- 0 3 to 5 times/week
- 0 More than 5 times/week

| 0   | 46 to 60 minutes more than sixty min | utes                   |                      |
|-----|--------------------------------------|------------------------|----------------------|
| Pla | ce an X in the types of              | of exercise you partic | cipate in regularly. |
|     | F                                    | requency/week          | Duration (min)       |
| 0   | Cycling                              |                        |                      |
| 0   | Walking                              |                        |                      |
| 0   | Jogging                              |                        |                      |
| 0   | Swimming                             |                        |                      |
| 0   | Rowing                               |                        |                      |
| 0   | Aerobics                             |                        |                      |
| Ot  | her (explain)                        |                        |                      |
| W   | hat do you consider y                | our body type to be?   | ,                    |
| 0   | lean                                 |                        |                      |
| 0   | average weight slightly over weigh   | t                      |                      |
| Ŏ   | over weight                          |                        |                      |

How long are your exercise periods?

() 0 to 15 minutes

() 16 to 29 minutes

30 to 45 minutes

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#### APPENDIX B

## EFFECTS OF CAFFEINE ON THERMOREGULATION OF ACTIVE MEN AT REST AND EXERCISE.

# STATEMENT OF INFORMED CONSENT FOR OBTAINING MEASUREMENTS OF: MAXIMUM OXYGEN UPTAKE, BODY COMPOSITION, RECTAL, AND SKIN TEMPERATURES.

#### **Purpose of the Study**

The purpose of this study is to observe caffeine's effect on sweat loss, skin (surface) temperature, rectal temperature, mean body temperature, skin heat conductance, and heart rate during rest and exercise.

#### **Invitation to Participate**

You are being invited to participate in a thesis project. The purpose of this research is to determine if caffeine effects temperature regulation during rest and exercise. Your participation is voluntary, and you can choose not to participate at any time during the project. This experiment will require the completion of three trials within two weeks. The first trial (approximately one hour) will assess your maximum oxygen uptake, body composition (fat percentage), and height. The second and third trials (approximately two and a half hours) will be the same except one trial will have caffeine (in a dose equal to 10 milligrams of caffeine per kilogram of body weight) added to the drink that will be taken before the resting and bicycling period. All the experiments will be conducted at the San Jose State University campus.

#### **Basis for Selection**

You have been selected because you are: a healthy male, between the ages of 18 and 40 years old, relatively lean, and exercise aerobically at least three times per week for 30 minutes or more. If you decide to participate, your response to a health history questionnaire will be reviewed by an exercise physiologist and, if satisfactory, you will be asked to participated in the tests described below.

## **Explanation of Procedures**

## Body Composition (Fat Percentage) - (approximately 30 minutes)

Underwater weighing requires three measurements: residual lung volume (air left in the lungs after a maximal exhalation), body weight while under water, and normal body weight. Height will also be taken at this time. To measure residual lung volume you will be breathing room air through a mouthpiece. At the end of a normal expiration a valve will be turned so you will be breathing a mixture of helium and room air from a spirometer. Oxygen will be added to the spirometer when needed. After breathing this gas mixture for several minutes a maximal inhaled and exhale breath will be recorded. Several measurements may need to be taken.

To measure underwater weight you will be sitting on a sling so you are positioned neck deep in 820 to 850F water. After being completely submerged under the water you will need to exhale as much air as possible and remain under water for five to ten seconds as the measurement is taken. This underwater weight will be measured six to ten times with rest periods allow between measures.

## Maximal Oxygen Uptake - (approximately 30 minutes)

Maximal oxygen uptake will be determined using a cycle ergometer at a pedal cadence of 60 repetitions per minutes (rpm). After a five minute warm-up at a power output of 180 kilopond meters per minute (kpm/min), the resistance will

be increased by 180 kpm/min every minute until you are unable to maintain the 60 rpm speed. Throughout this maximal test, blood pressure will be measured every minute, where respiratory parameters and heart rate will be monitored continuously.

#### <u>Caffeine/Non-Caffeine Trials</u> – (approximately two and a half hours each)

As mentioned previously these trials will be the same except one trial will have caffeine added (in a dose equal to 10 milligrams of caffeine per kilogram of body weight) to a diet drink mixture and the other trial will not have caffeine added. Before both of these trials you will be required to not eat or drink any caffeine for four days before the test. During these trials skin temperature will be measured using skin probes that will be held on with elastic bands; rectal temperature will be measured using a rectal probe; ventilation will be measured which will require you to breath into a mouth piece; and heart rate will be monitored which will require electrode application.

After drinking the liquid you will sit on a chair for 30 minutes while temperatures, respiration, and heart rate are recorded every ten minutes. These measurements will continue into the exercise phase, where you will pedal the ergometer at a rate and resistance equal to 65% of you maximal work capacity. A fan will blow air on you as you complete 70 minutes of cycling.

#### **Risks and Discomforts**

#### **Underwater Weighing**

Although the water quality in the tank is maintained daily there is still a possible risk of infection from the water. This risk is small since the water is chemically treated and filtered daily. As with most pool underwater experiences there is a chance of chlorine irritation, swallowing water, and choking.

#### **Residual Lung Volume**

There may be some discomfort from the nose clip and from breathing through the mouthpiece. Some people experience dizziness and lightheadedness when preforming breathing measurements.

#### Maximal Oxygen Uptake

There will be some discomfort that is common with high levels of exercise in this test: sweating, increased heart rate, increased breathing rate, elevated blood pressure, and fatigue. You may experience dryness in the mouth, throat, and chest. Toward the end of this maximal test you may incur abnormal blood pressure, fainting, dizziness, muscle cramps, muscle fatigue, and abnormalities in heart beat. You may feel lightheaded and slightly nauseous for a short while following the test. If abnormalities are detected in pulmonary function, electrocardiographic recordings, or if other abnormal physiological signs occur the test will be stopped and you will be asked not to continue the rest of the experiment.

#### Caffeine Effects/Caffeine Withdrawal

Adverse reactions to caffeine ingestion can include: restlessness, irritability, nervousness, insomnia, headache, nausea, vomiting, elevated blood glucose, tingling of the face, flushing, an increase or decrease in heart rate, and muscle twitching. At the dosage applied in this study caffeine may occasionally cause cardiac irregularities. Habitual caffeine consumers may experience headaches after being deprived of caffeine.

#### Sixty-Five Percent Bicycling Work Load

During this test you will experience the same types discomfort as you would during heavy exercise: sweating, increased heart rate, increased breathing rate, elevated blood pressure, and fatigue. You may experience lightheadedness and dryness in the mouth, throat, and chest.

## Benefits from Participating in this Study

You will benefit from this study by receiving information about your body fat percentage and maximal oxygen consumption. If you are a caffeine user you may be interested in how caffeine effects your body temperature.

## Assurance of Confidentiality

The information that will be obtained will be treated as privileged and confidential and will not be released or revealed without your expressed written consent. The data obtained, however, may be used for a statistical or scientific purpose with your right of privacy retained. Your right to confidentiality will be protected.

#### Withdrawal from the Study

Your participation in this research is voluntary. You may withdraw from this study at any time (including during a test) without prejudice. You may also decline to answer any question or item on the health history questionnaire. Maximal oxygen consumption testing will be supervised and conducted a Certified Exercise Test Technologist. Certified CPR personnel will also be present during all testing.

If you have any questions about this research feel free to ask them. If questions come up later or in case of an emergency please contact me, Nancy Dunagan, at (408) 248-7578 or Dr. Craig Cisar (408) 924-3018. In case there is a complaint you can contact Dr. Craig Cisar (408) 924-3018. If there are questions about research subject's rights, or in the event of research-related injury you may contact Dr. Serena Stanford, Associate Academic Vice President of Graduate Studies and Research at (408) 924-2480.

#### AGREEMENT TO PARTICIPATE IN RESEARCH

#### SAN JOSE STATE UNIVERSITY

RESPONSIBLE INVESTIGATOR: Nancy A. Dunagan

TITLE OF PROTOCOL: Effects of Caffeine on Thermoregulation of Active Men at Rest and Exercise.

#### Consent

By signing this form, I agree that:

- (a) I have read the information provided above and have decided to participate in this study;
- (b) I understand the discomforts and risks involved;
- (c) I understand that I can withdraw from this study at any time without prejudice to my relations with SJSU.
- (d) I understand that my identity will be kept confidential.
- (e) I understand that I may ask questions about this study at any time.

| SIGNATURE OF SUBJECT      |  |
|---------------------------|--|
| DATE                      |  |
| PRINTED NAME              |  |
| SIGNATURE OF WITNESS      |  |
| SIGNATURE OF INVESTIGATOR |  |

## APPENDIX C

## RESIDUAL LUNG VOLUME AND UNDERWATER WEIGHING DATA SHEET

| Name  |                            |                   |
|---|----------------------------|-------------------|
| Vital Capacity (L)  | _ <del>_</del>             |                   |
| Starting lung volume (VC + 0.5L)  |                            |                   |
| Initial N2 reading after purging spin<br>trial 1 trial 2                                | rometer (%)<br>trial 3     | average           |
| **(ADD STARTING LUNG VOL  | <br>UME)**                 |                   |
| N2 reading at end expiration of roo   | om air (%)                 |                   |
| N2 reading at equilibrium during fa   | ast hard breathing (%)     |                   |
| N2 reading at end expiration after  | rebreathing technique (%)  |                   |
| Temperature of expired air (°C)Body Weight (KG)   |                            |                   |
| Height cm   |                            |                   |
| Under Water body weight (KG)  |                            |                   |
| Average of three highest underwa  | ter weight scores within 0 | 0.05kg of each of |
| Harness weight (KG)  Water temperature (F)  Water density (KG/L)  Min % fat 1 Max % fat |                            |                   |

#### APPENDIX D

#### **List of Caffeine Sources**

The following are caffeine containing foods and drinks that you should not have for four days prior to the test. If you accidentally do have some caffeine please contact Nancy Dunagan (408-248-7578) to re-schedule the test date. Please, read the content labels of the food and drink you ingest to double check and make sure caffeine does not go into your body. Your system needs to be clean from caffeine in order to obtain accurate results in this experiment.

#### **Drinks**

Coffee of any type (not even decaffeinated)

Tea of any type Cola of any type

Tab

Dr. Pepper

Mr. Pibb

Aspen

Mountain Dew

Mellow Yellow

Cocoa of any type

Chocolate of any type

#### Food

Candies and Cookies containing chocolate and cocoa Check packages on frozen dairy products for caffeine

Check gelatin and puddings for caffeine

Drugs - Prescription

APC's (Aspirin, Phenacetin, Caffeine)

Cafergot

Darvon

Fiorinal

Migral

#### Over the counter drugs

Vanquish, Excedrin
Extra Strength Excedrin
No-Doz
Vivarin

Ban-Drowz
Tirend
Kirkaffeine
Quick Pep

Appendirne Caffedrine Dexatrim Dristan

Anacin, Aspirin compound, Bromoselter

Cope, Emperin compound, Midol

These lists were derived from Chung (1986), Kastrup (1989), and from Jacobson and Kulling (1989).

## APPENDIX E ADVERTISEMENT TO SOLICIT SUBJECTS

# EFFECTS OF CAFFEINE ON THERMOREGULATION OF ACTIVE MEN AT REST AND EXERCISE.

Are you interested in finding out your fat percentage and maximum oxygen consumption? You could have these measurements made free of charge and at the same time contribute to scientific research. To volunteer for this study you need to be:

| measurements made free of charge and at the same time contribute to scientific research. To volunteer for this study you need to be:   |
|--|
| MALE   |
| BETWEEN 18 AND 40  |
| AVERAGE TO LEAN BODY TYPE  |
| EXERCISE AEROBICALLY AT LEAST 3 TIMES PER WEEK (CYCLE, JOG, ETC.) FOR AT LEAST 30 MINUTES  |
| EACH EXERCISE SESSION  |
|  |
| TIME REQUIREMENT:  |
| 1–2 hour session and   |
| 2–2.5 hour sessions  |
|  |
| If you are interested in helping with data collecting, or volunteering to be a subject call Nancy Dunagan at (408) 248-7578 to find out more about the study. There is no obligation. Participation in this research is on a voluntary basis |

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## APPENDIX F DATA COLLECTION SHEETS

#### CAFFEINE/NON CAFFEINE TRIAL

1 OR 2

| Name   |
|--|
| ENVIRONMENTAL TEMPERATURE.   |
| Wet bulb temp (sling psychrometer) two lowest agreeing temp°C                                      |
| Dry bulb temp°C  |
| Relative humidity  |
| Barometric pressure  |
| STPD (standard temp and press dry)   |
| STPD sat   |
| BTPS (Body tem and press sat)  |
| Pre-exercise weight  |
| Oral Temperature 1 pre-exercise  |
| - Blindfold and put a nose clip on subject. Drink mixture should be 37°C of 237 ml (80z) lemonade. |
| - Drink temperature  |
| Wind speed   |
| Post-exercise weight   |
| Oral Temperature 2 post exercise   |

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| Name | Date |
|------|------|
|      |      |

Test #1 or #2

Resting Time starts upon completion of drinking the beverage. Give volunteer one minute to drink liquid.

| Time     | RecT#1        | Chest#3 | Back#4 | FArm#6 | Calf#8 | UArm#9 | Thigh#10 |
|----------|---------------|---------|--------|--------|--------|--------|----------|
| 1 m      |               |         |        |        |        |        |          |
| 5 m      |               |         |        |        |        |        |          |
| 10m      |               |         |        |        |        |        |          |
| 20m      |               |         |        |        |        | ·      |          |
| 29m      |               |         |        |        |        |        |          |
| Exercise | Begins at 30m |         |        |        |        | 11.2   |          |
| 35m      |               |         |        |        |        |        |          |
| 40m      |               |         |        |        |        |        |          |
| 50m      |               |         |        |        |        |        |          |
| 60       |               |         |        |        |        |        |          |
| 70       |               |         | ·      |        |        |        |          |
| 80       |               |         |        |        |        |        |          |
| 90       |               |         |        |        |        |        |          |
| 100      |               |         |        |        |        |        |          |

cool down

## GAS ANALYSIS AND HEART RATE DURING CAF/NON CAF TRIALS

| Name | Date |
|------|------|
|------|------|

#### TRIAL 1 OR 2

| TIME                | $v_{E}$ | VO <sub>2</sub> | VCO <sub>2</sub> | HEART RATE |
|---------------------|---------|-----------------|------------------|------------|
| TAKE DRINK          |         |                 |                  |            |
| POST DRINK<br>1 MIN |         |                 |                  |            |
| 2 MIN               |         |                 |                  |            |
| 3 MIN               |         |                 |                  |            |
| 8 MIN               |         |                 |                  |            |
| 9 MIN               |         |                 |                  |            |
| 10 MIN              |         |                 |                  |            |
| 18 MIN              |         |                 |                  |            |
| 19 MIN              |         |                 |                  |            |
| 20 MIN              |         |                 |                  |            |
| 28 MIN              |         |                 |                  |            |
| 29 MIN              |         |                 |                  |            |
| 30 MIN              |         |                 |                  |            |

#### GAS ANALYSIS AND HEART RATE DURING CAF/NON CAF TRIALS

| Name | Date |
|------|------|
|      |      |

#### TRIAL 1 OR 2

| TIME                 | $V_{\rm E}$ | VO <sub>2</sub> | VCO <sub>2</sub> | HEART RATE |
|----------------------|-------------|-----------------|------------------|------------|
| START EXER<br>38 MIN |             |                 |                  |            |
| 39 MIN               |             |                 |                  |            |
| 40 MIN               |             |                 |                  |            |
| 48 MIN               |             |                 |                  |            |
| 49 MIN               |             |                 |                  |            |
| 50 MIN               |             |                 |                  |            |
| 58 MIN               |             |                 |                  |            |
| 59 MIN               |             |                 |                  |            |
| 60 MIN               |             |                 |                  |            |
| 68 MIN               |             |                 |                  |            |
| 69 MIN               |             |                 |                  |            |
| 70 MIN               |             |                 |                  |            |
| 78 MIN               |             |                 |                  |            |
| 79 MIN               |             |                 |                  |            |
| 80 MIN               |             |                 |                  |            |
| 88 MIN               |             |                 |                  |            |
| 89 MIN               |             |                 | :                |            |
| 90 MIN               |             |                 |                  |            |
| 98 MIN               |             |                 |                  |            |
| 99 MIN               |             |                 |                  |            |
| 100 MIN              |             |                 |                  |            |

APPENDIX G-1
Repeated Measures Two Way Analysis of Covariance Summary Table for Oxygen Uptake

| SOURCE               | SS             | DF  | MS    | F      | SIGNIFICANCE |
|----------------------|----------------|-----|-------|--------|--------------|
| Within Cells         | 0.34           | 9   | 0.04  | 2.35   | 0.160        |
| Caffeine             | 0.09           | 100 | 0.09  |        | 0.100        |
| Within Cells<br>Time | 4.62<br>266.48 | 10  | 26.65 | 577.03 | 0.000        |
| Within Cells         | 0.51           | 100 | 0.01  | 2.00   | 0.032        |
| Interaction          | 0.11           | 10  | 0.01  | 2.09   | 0.032        |

## APPENDIX G-2 Repeated Measures Two Way Analysis of Covariance Summary Table for Heart Rate

| Within Cells | 1006.99   | 9   | 111.89   |        |       |
|--------------|-----------|-----|----------|--------|-------|
| Caffeine     | 67.37     | 1   | 67.37    | 0.605  | 0.458 |
| Within Cells | 10627.67  | 100 | 106.28   |        |       |
| Time         | 384786.79 | 10  | 38478.68 | 362.06 | 0.000 |
| Within Cells | 1866.77   | 100 | 18.67    |        |       |
| Interaction  | 394.62    | 10  | 39.46    | 2.11   | 0.030 |

#### **APPENDIX G-3**

## Repeated Measures Two Way Analysis of Covariance Summary Table for Mean Rectal Temperature

| Within Cells | 0.74  | 9  | 0.08 |        |       |
|--------------|-------|----|------|--------|-------|
| Caffeine     | 0.02  | 1  | 0.02 | 0.19   | 0.677 |
| Within Cells | 6.27  | 90 | 0.07 |        |       |
| Time         | 78.25 | 9  | 8.69 | 124.85 | 0.000 |
| Within Cells | 1.06  | 90 | 0.01 |        |       |
| Interaction  | 0.20  | 9  | 0.02 | 1.86   | 0.068 |

#### **APPENDIX G-4**

## Repeated Measures Two Way Analysis of Covariance Summary Table for Mean Skin Temperature

| Repeated 111 |       |    |      | <del></del> |       |
|--------------|-------|----|------|-------------|-------|
| Within Cells | 7.63  | 9  | 0.85 |             | 0.545 |
| Caffeine     | 0.33  | 1  | 0.33 | 0.39        | 0.547 |
| Within Cells | 42.41 | 90 | 0.47 | _           |       |
| Time         | 7.88  | 9  | 0.88 | 1.86        | 0.069 |
| Within Cells | 5.43  | 90 | 0.06 |             |       |
| Interaction  | 0.26  | 9  | 0.03 | 0.48        | 0.886 |

### **APPENDIX G-5**

## Repeated Measures Two Way Analysis of Covariance Summary Table for Skin Heat Conductance

| •       |   |   |  |   |
|---------|---|---|--|---|
| 707.31  | 9   | 78.26   |  |   |
| 128.30  | 1   | 128.30  | 1.64   | 0.232   |
| 2624.31 | 90  | 29.16   |  |   |
|         | 9   | 5077.92   | 174.05   | 0.000   |
|         | 90  | 9   | 6.64   |   |
| 105.72  | 9   | 11.75   | 1.77   | 0.085   |
|         | 707.31<br>128.30<br>2624.31<br>45701.29<br>597.65 | 707.31 9<br>128.30 1<br>2624.31 90<br>45701.29 9<br>597.65 90 | 707.31     9     78.26       128.30     1     128.30       2624.31     90     29.16       45701.29     9     5077.92       597.65     90     9 | 707.31 9 78.26<br>128.30 1 128.30 1.64<br>2624.31 90 29.16<br>45701.29 9 5077.92 174.05<br>597.65 90 9 6.64 |

## APPENDIX H-1

### **Results from Selected Caffeine Studies on Animals**

| Amount of caffeine (caf.)                                | Time after ingestion              | Subjects                      | Exercise   | Results  | Reference  |
|--|-----------------------------------|-------------------------------|--|--|--|
| 10–20 mg/kg  | 120 min                           | Rat                           | none   | Increased rectal temperature in 8°, 22°, and 30°C environments.  | Lin, Chandran,<br>and Liu (1980)                   |
| 12.5 mg/kg,<br>25 mg/kg,<br>50 mg/kg, and<br>100 mg/kg   | up to 3.5 hrs                     | Rat                           | none   | 12.5 and 25 mg/kg produced increased body temperatures; 50 and 100 mg/kg produced decreased body temperatures.   | Schlosberg<br>(1983)                               |
| 12.5 mg/kg,<br>25 mg/kg,<br>50 mg/kg, and<br>100 mg/kg   | daily for<br>7 days               | Rat                           |  | By the 4th day 50 and 100 mg/kg doses produced increased body temperatures that continued through the 7th day. Both 12.5 and 25 mg/kg doses increased body temperature all 7 days. |  |
| 10, 20,<br>40 mg/kg                                      | 30 min                            | Rat                           | none   | 10mg/kg, no effect. 20 and 40 mg/kg, higher interscapular temperatures.  | Wellman, and<br>Marmon (1985)                      |
| 0.8 mg/kg<br>nicotine +<br>10 mg/kg<br>caffeine          | 30 min                            | Rat                           | none   | Increased interscapular temperature.   | Wellman,<br>Marmon, Reich,<br>and Ruddle<br>(1986) |
| 5 mg/kg,<br>25 mg/kg,<br>50 mg/kg                        | just prior to exercise            | Rat                           | treadmill<br>running 45 or<br>90 min                           | No effect on plasma FFA or blood glycerol.   | Winder (1986)                                      |
| 5 mg caf.,<br>25 mg caf., or<br>0.9% sodium-<br>chloride | immediately<br>before<br>exercise | Rats,<br>endurance<br>trained | treadmill<br>running<br>45 min,<br>90 min, or to<br>exhaustion | Caf. did not enhance endurance time or<br>change liver glycogenolysis, liver cAMP,<br>muscle glycogen, plasma FFA, blood<br>glucose, or blood lactate.                             | Arogyasami,<br>Yang, and<br>Winder (1989)          |

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### **APPENDIX H-2**

#### **Results from Selected Caffeine Studies on Humans**

| Amount of caffeine (caf.)     | Time after ingestion         | Subjects                       | Exercise  | Results   | Reference  |
|-------------------------------|------------------------------|--------------------------------|---|---|--|
| ~330 mg                       | 60 min<br>before<br>exercise | male<br>cyclists               | cycle at 80% VO <sub>2</sub> max until exhaustion                   | Caffeine increased exercise time and decreased rate of perceived exertion.  | Costill, Dalsky,<br>and Fink<br>(1978)                   |
| 5mg/kg                        | 1 hour and<br>3 hours        | male                           | 2 hour<br>run in<br>hot/humid<br>conditions                         | No significant differences in rectal temp., plasma volume, or electrolyte levels, but tendencies toward higher rectal temp. and lower plasma vol. for caffeine group.                                   | Gordon et al.<br>(1982)                                  |
| 4 mg/kg                       | up to<br>2 hrs               | trained/<br>untrained<br>males | none  | Trained subject had greater changes, but all increased resting metabolic rate, plasma epinephrine, and free fatty acid utilization, and decreased respiratory exchange ratio and plasma norepinephrine. | LeBlanc, Jobin,<br>Cote, Samson,<br>and LaBrie<br>(1985) |
| 100-200mg                     | 1 and 2 hrs                  | male                           | none  | Increase in metabolic stimulation and systolic and diastolic blood pressures.   | Ammaturo, and<br>Monti (1986)                            |
| ephedrine + 30 mg<br>caffeine |                              | lean/post-<br>obese<br>males   |   | energy expenditure in post-obese man  | Dullo, and<br>Miller (1986)                              |
| 10 mg/kg and<br>15 mg/kg      | 60 min<br>before<br>exercise | female                         | treadmill running at 70-75% of VO <sub>2</sub> max until exhaustion | 15 mg dose increased maximal running time, lowered R, decreased FFA, and decreased rate of perceived exertion and respiratory exchange ratio (R <sub>E</sub> ).   | McNaughton<br>(1986)                                     |
| 5 mg caf.                     | 60 min<br>before<br>exercise | male                           | ergometer,<br>graded load   | Caf. did not effect heart rate, minute ventilation R <sub>E</sub> , glucose, or VO <sub>2</sub> at rest or during exercise.   | Bond et al. (1987)                                       |
| 10 mg/kg and<br>15 mg/kg      | 60 min<br>before<br>exercise | female                         | ergometer,<br>increase<br>load<br>30kpm/min                         | Both doses of caf. increased time to exhaustion, and had lower blood lactate levels.  | McNaughton,<br>and Davies<br>(1987)                      |

## Results from Selected Caffeine Studies on Humans (Concluded)

| Amount of caffeine   | Time after ingestion                       | Subjects             | Exercise  | Results   | Reference                                       |
|--|--|----------------------|---|---|---|
| sucrose (~81g) or caffeine (~384mg) or sucrose (~72g) + caffeine (~396mg) or placebo | 60 min<br>before and<br>during<br>exercise | trained<br>males     | treadmill running at ~80% of VO <sub>2</sub> max until exhaustion | Caf. and Su + Caf. increased time to exhaustion equally over control group.   | Sasaki,<br>Maeda, Usui,<br>and Ishiko<br>(1987) |
| 5 mg/kg  | 60 min and<br>150 min                      | untrained<br>females | 90 min walk<br>at 55% of<br>VO <sub>2</sub> max                   | Caffeine increased VO <sub>2</sub> and respiratory exchange ratio after 60 minutes rest; increased VO <sub>2</sub> and FFA, and decreased R <sub>E</sub> during 90 minutes of exercise. | Chad, and<br>Quigley<br>(1989)                  |
| 100 mg every 2 hrs<br>or 12hrs   | up to 24<br>hrs                            | lean/post-<br>obese  | none  | All subjects had increased resting metabolic rate during the 12 hours of caf. ingestion.  | Dullo, and<br>Miller (1989)                     |

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|  |                                |                      | and during submaximal exercise  |  |
|  |                                |                      | $29.2 \pm 6.2 \text{ yr}, \text{ $\dot{V}O_{2 \text{ max}}$= 3.73 $\pm$}$ |  |
| $0.46 \mid \min^{-1}$ , relative body fat = $12.3 \pm 3.7\%$ , mean $\pm$ SD) with, (CT) and without (NCT) the ingestion   |                                |                      |   |  |
| of 10 mg of caffeine per kg of body weight. Oxygen uptake ( $\dot{VO}_2$ ), heart rate (HR), and rectal (Tre) and  |                                |                      |   |  |
| mean skin ( $\overline{T}$ sk) temperatures were recorded for 100 minutes starting one minute after ingestion of   |                                |                      |   |  |
| caffeine or a placebo. Data were collected throughout 30 minutes of rest (sitting) and the following   |                                |                      |   |  |
| 70 minutes of sitting leg ergometer exercise using the same constant load (1,088 ± 153 kgm·min <sup>-1</sup> ) in  |                                |                      |   |  |
| both NCT and CT. The load resulted in a mean relative exercise intensity equal to approximately 68%  |                                |                      |   |  |
| of $VO_{2 \text{ max}}$ . Skin heat conductance (Hsk) and sweat rate were calculated. Two-way analysis of  |                                |                      |   |  |
| covariance revealed no significant (P>0.05) differences between NCT and CT in $\dot{VO}_2$ , HR, Tre, $\overline{Ts}$ or Hsk. A dependent t-test indicated no significant difference between NCT and CT in sweat rate. The |                                |                      |   |  |
| or Hsk. A dependent t-test inc   | CT and CT in sweat rate. Thus, |                      |   |  |
| a high level of caffeine ingestion has no detrimental effects on body temperatures and thermoregulato  |                                |                      |   |  |
| responses during moderately  | ons.                           |                      |   |  |

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