## Effect of yogic intervention on the autonomic nervous system in the patients with coronary artery disease: a randomized controlled trial

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تأثير تدخلات اليوغا على الجهاز العصبي المستقل لدى مرضى الشريان التاجي: دراسة مُشَهَّدة عشوائياً أ. بال، ن سريفاستافا، ف.س.ناراين. جي جي أغروال. م. راني

الخلاصة: ضمت الدراسة 258 مريضاً من قسم الأمراض القلبية في جامعة شهاتراباتي ساهوجي مهراج الطبية، في لكنهو، ممن اختارهم الباحثون، وقد شُخِّص لديهم جيعاً مرض الشريان التاجي. وقسمهم الباحثون إلى مجموعتين؛ مجموعة اليوغا ومجموعة غير المارسين لليوغا، وتتألف كل مجموعة من 129 مريضاً، وقد استكمل 208 منهم متطلبات بروتوكول الدراسة، وكانت تدخلات اليوغا تستغرق 35-40 دقيقة/ يومياً، وتكرر خمسة أيام كل أسبوع على مدى 18 شهراً في قسم الفيزيولوجيا. وأجرى الباحثون اختبارات وظائف الجهاز العصبي المستقل لكلا المجموعتين في الوقت صفر وبعد 18 شهراً، ولاحظوا وجود انخفاض يُعْتدُ به إحصائياً في مَنْسَب كتلة الجسم، ومحيط الخصر، والضغط الانقباضي والضغط الانبساطي وسرعة القلب (0.05 < P) مما يدل على تأثير إيجابي يُعْتدُ به إحصائياً عند ممارسة اليوغا كإجراء إضافي لدى مرضى الشريان التاجي.

ABSTRACT In this study 258 patients from the Department of Cardiology in Chhatrapati Shahuji Maharaj Medical University, Lucknow were selected to participate. All had been diagnosed with coronary artery disease. They were randomly divided into 2 groups, the yoga group and the non-yoga group, 129 in each group; 208 completed the study protocol. The yogic intervention consisted of 35–40 min/day, 5 days a week over a period of 18 months in the Department of Physiology. Autonomic function testing was done in both the groups at zero time and after 18 months. We observed a statistically significant reduction in body mass index, waist circumference, systolic and diastolic blood pressure, and heart rate (P < 0.05), i.e. a significant positive effect was observed when yoga therapy was used as an adjunct in patients with coronary artery disease.

Effet d'une intervention basée sur le yoga sur le système nerveux autonome de patients atteints d'une maladie coronarienne : étude clinique randomisée et contrôlée

RÉSUMÉ Dans la présente étude, 258 patients consultant le service de cardiologie du centre hospitalier universitaire Chhatrapati Shahuji Maharaj de Lucknow (Inde) ont été sélectionnés pour cette intervention. Tous avaient reçu le diagnostic de maladie coronarienne. Les patients ont été répartis aléatoirement en deux groupes, le premier bénéficiant d'une intervention basée sur le yoga contrairement au second. Chaque groupe comptait 129 patients. Au total, 208 patients ont achevé le protocole de l'étude. L'intervention yogique durait 35 à 40 minutes par jour, 5 jours par semaine, sur une période de 18 mois dans le service de physiologie. Des tests de la fonction autonome ont été menés dans les deux groupes au début de l'étude et à 18 mois. Nous avons observé une réduction statistiquement significative de l'indice de masse corporelle, du tour de taille, de la pression systolique et diastolique, ainsi que de la fréquence cardiaque (P < 0,05). Un effet positif important a donc été observé lorsqu'une thérapie yogique a été appliquée en tant que traitement adjuvant chez des patients atteints d'une maladie coronarienne.

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Received: 01/12/10; accepted: 01/11/11

#### Introduction

Indians have considerably higher prevalence of premature coronary artery disease (CAD) and standardized mortality rates for CAD compared with Europeans, Chinese and Malays [1–4]. Projections on mortality rates due to CAD in India indicate almost 100% increase from 1985 to 2015 [5]. The Indian population is more prone to CAD at a younger age: Indians are affected 5–10 years earlier than people in other communities [6,7]. This condition is the leading cause of morbidity and mortality in most industrialized societies [8].

It is believed that yogic practices were originated in the civilization of the Indian subcontinent. Yogic practices such as shatkarmas, asana, pranayama and meditation practices are very useful for physical, mental and spiritual development. Through these practices, individual consciousness is purified and the purified consciousness level can achieve salvation. Stress and anxiety levels can also be reduced through the practices of yoga. Relaxation asana like shavasana, makarasana, shashankasana are widely used for their stress releasing response.

The objective of this study was to determine the effect of yogic practices on sympathetic (blood pressure response to lying to standing test and handgrip test) and parasympathetic (30th:15th ratio, E:I ratio, valsalva ratio) function in patients with CAD.

#### Methods

The study was part of a larger study conducted under the Extra Mural Research Project of the Department of Department of Ayurveda, Yoga & Naturopathy, Unani, Siddha and Homoeopathy (AYUSH), at the Indian Ministry of Health and Family Welfare. Some parts of the methodology have been described in a previously published study which was part of the same project [9].

Sample size was calculated as 200 This was increased to 258 to cover drop-outs, etc. Patients from the Department of Cardiology, Chhatrapati Shahuji Maharaj Medical University, Lucknow, India who fulfilled the inclusion criteria were invited to participate in the yoga interventional study. We approached 464 patients; 258 eligible patients agreed to participate (Figure 1). The study was carried out from February 2007 to July 2010. The progress of the participants through the study is described in Figure 1. The institutional Research Ethics Committee approved the study. All participants gave written informed consent to take part in the study before anthropometric measurements were taken.

A senior cardiologist referred the patients after examining their physical health and medication status. Patients were selected on the basis of



Figure 1 Flow-chart of the participants (patients with coronary artery disease, Lucknow, India) in the 18-month yogic intervention study showing progression of both intervention and control groups

previous history and blood pressure level. Prescribed medications (metoprolol/atenolol, asprin, clopidogrel, atorvastatin/rosuvastatin, ramipril/ losartan/telmisartan) were continued in both groups.

Using a random number generator, each of the 258 patients selected was randomly assigned to one of 2 groups: (study group) medication + yoga, and (control group) medication only (n = 129 in each group). A professional not associated with this study generated the randomization scheme.

Yoga instructors (post graduate) with more than 2 years experience were selected to lead yogic practices. Yoga sessions were free of cost and all the necessary facilities were provided for the participants (airy room, yoga mats, *jal neti* pot, etc.).

Most of the participants were from Lucknow. Patients who dropped out of the study did not differ significantly in terms of age and sex; 208 completed the study protocol (Figure 1). Before the study all the participants were asked to maintain their routine activities and not initiate any new physical activities for the duration of the study.

Patients with proven CAD were recruited. Disease was diagnosed by history of electrocardiograms, echocardiography and treadmill testing. Their willingness to complete the entire span of the project (18 months) was assured. Patients who had other co-morbid conditions (e.g. malignant hypertension, diabetes mellitus, chronic obstructive pulmonary disease, asthma, diseases of the nervous system, endocrinal disorders, congenital heart disease) and patients with known complications of CAD, those on pacemakers, and those who had undergone bypass surgery were excluded from the study. Participants who did not participate in a minimum number of yoga intervention classes (< 60%) were not included in the final assessment (Figure 1).

#### Autonomic functions tests

A battery of 6 standard tests was used to assess the integrity of autonomic function status of the participants [10]. These tests were helpful in assessment of the reactivity of divisions of the autonomic nervous system, i.e. tests for parasympathetic and sympathetic function. The participants were made comfortable and the entire procedure was explained before the recording was done. They were asked to lie quietly on a couch and were connected to an electrocardiogram (ECG) monitor. Blood pressure was measured using a sphygmomanometer with a standard cuff around the left arm. After this preparation the participants were asked to remain supine and rest for about 10 minutes before the tests began. First the resting cardio-respiratory parameters (heart rate, and blood pressure) were recorded in a lying position.

# Test for parasympathetic reactivity

# Heart rate response to standing from the supine posture (30:15 ratio) [11]

For this test the heart rate was recorded in the lying posture by ECG; after recording a stable baseline resting heart rate, the patient was asked to get up unaided and the heart rate was continuously recorded. From the post-standing record, the ratio of the longest to the shortest R–R interval (approximately corresponding to the 30th and 15th beats, by convention: maximum R–R interval around 30th beat, minimum R–R interval around 15th beat) was recorded.

#### Slow deep breathing test

After 3–5 min in a sitting position, the participants were asked to take deep, slow breaths at a rate of 6 breaths per minute [12] in the supine position for 1 min [11]. An automatic event marker was used to mark the inspiratory (I) and expiratory (E) phases, indicating delivery of signals. The E:I ratio was the ratio of averaged maximum and

minimum R–R intervals during expiration and inspiration respectively during the manoeuvre. We checked whether the respective phase of respiration coincided with the signal marking, which was confirmation of the precise execution of the test. If not, the test was repeated after adequate interval or at the end.

E:I ratio = average maximum R–R interval during expiration/average minimum R–R interval during inspiration.

#### Valsalva manoeuvre [13]

The valsalva manoeuvre comprises abrupt, transient and voluntary elevation of intra-thoracic and intraabdominal pressure provoked by straining [14], expiring forcefully through a mouthpiece attached to a manometer to generate a pressure of 40 mmHg and maintaining this pressure for 15 seconds after which the participants released the effort and the mouthpiece was removed. Due care was taken to prevent deep breathing before and after the release of strain. The participants were asked to sit quietly and breathe normally after the test.

Valsalva ratio = longest R–R interval after manoeuvre (phase IV)/shortest R–R interval during manoeuvre (phase II).

### Tests for sympathetic reactivity Blood pressure response to lying to standing test

The participants were asked to stand up quickly in 3 seconds and remain standing quietly for 2 minutes putting equal weight on each leg and without any support [15]. Blood pressure was recorded at 0 min (baseline; in lying position) and 0.5, 1.0 and 2.0 min while standing up. The differences in systolic and diastolic pressure from baseline were calculated. The participants were asked to sit quietly after the test.

#### Hand grip test [10]

A very light, small, hand-held handgrip dynamometer, which shows isometric strength, was used for this

130.1 (10.3)

83.7 (8.0)

73.0 (9.1

test. After instruction in the use of the dynamometer, baseline blood pressure was recorded. The participants were then asked to grip the dynamometer with their dominant hand maximally. This was repeated 3 times at short intervals. The maximum of these 3 readings was taken as maximum voluntary contraction. For the test, the subjects were asked to maintain the pressure on the dynamometer for 5 min at 30% of maximum voluntary contraction. Blood pressure was recorded on the non-exercising arm at 1 min and 2 min after the onset of isometric contraction (before release of grip). The changes in systolic and diastolic blood pressure were measured.

#### **Yogic intervention**

Under the guidance and supervision of yoga experts, participants in the yoga group performed yoga practices as directed. These practices are described in detail in a previously published study which was part of the same project [9].

The total duration of these practices was 35–40 min/day, 5 days a week over the 18 months of the study in the Department of Physiology at Chhatrapati Sahuji Maharaj Medical University.

Using the independent *t*-test of differences [16,17] to analyse the differences between baseline values and post-intervention (after 18 months) values for each variable in both groups (yoga and non-yoga). P < 0.05 was considered statistically significant,

#### Results

The baseline characteristics for both groups are presented in Table 1. Patients of both sexes were included and the overall age range was 35–82 years. In the control group mean age was 56.4 [standard deviation (SD) 10.9] years and in the yoga group mean age was 59.1 (SD 9.9) years.

After 18 months 208 participants remained in the study, 105 in the

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Characteristic	Control group <sup>a</sup> ( <i>n</i> = 129)	Yogic group <sup>b</sup> ( <i>n</i> = 129)			
	Mean (SD)	Mean (SD)			
Age (years)	56.4 (10.9)	59.1 (9.9)			
Height (cm)	161.6 (8.4)	161.5 (9.0)			
Weight (kg)	65.6 (12.3)	64.9 (9.3)			
WC (cm)	94.2 (10.5)	93.0 (8.3)			
HC (cm)	95.6 (11.3)	94.5 (9.4)			

### Table 1 Baseline characteristics of patients with coronary artery disease who participated in the yoga intervention study, Lucknow, India, 2007–2009

°104 males, 25 females.

Heart rate (beats/min)

DBP = diastolic blood pressure.

SBP (mmHg)

DBP (mmHg)

<sup>b</sup>103 males, 26 females. SD = standard deviation; WC = waist circumference, HC = hip circumference, SBP = systolic blood pressure;

129.8 (14.1)

82.3 (8.6)

71.6 (11.4)

intervention group and 103 in the control group: 50 had been lost to followup, 26 in the control group and 24 in the intervention group (Figure 1).

In the intervention group mean body mass index (BMI) decreased to 24.3 (SD 3.5) kg/m<sup>2</sup> from the baseline of 24.8 (SD 3.8) kg/m<sup>2</sup>, whereas in the control group mean BMI remained substantially the same (Table 2). Mean systolic blood pressure (SBP) fell to 123.1 (SD 9.4) mmHg from the baseline level of 130.1 (SD 10.3) mmHg and in the control group mean SBP fell slightly to 129.1 (SD 9.3) mmHg from the baseline value of 129.8 (SD 14.1) mmHg. The DBP level in the intervention group fell to 80.48 (SD 5.12) mmHg from the baseline level of 83.7 (SD 8.0) mmHg and in control group mean diastolic blood pressure (DBP) rose to 83.8 (SD 5.7) mmHg from the baseline level of 82.3 (SD 8.6) mmHg.

There were significant changes observed in BMI (P < 0.0001), waist:hip ratio (P < 0.005), SBP (P < 0.002), DBP (P < 0.0002) and heart rate (P < 0.0006) (Table 3).

#### Discussion

High blood pressure is a major cause of cardiovascular disease and primary

hypertension is a frequent pathological condition. Sympathetic hyperactivity may be involved in primary hypertension. Hypertensive patients have been found to have a significantly higher sympathetic response to central and peripheral stimulations and a significantly lower parasympathetic response when compared to normotensive controls [18]. In our study, BMI, waist:hip ratio, SBP, DBP and resting heart rate decreased significantly more in the intervention group than in the control group. In a study in Bombay, the shavasana group had a significant decrease in blood pressure [19]. Yogic practices have been shown to have significant benefits for cardiovascular health: through these practices heart rate, systolic blood pressure and diastolic blood pressure decrease significantly [20–22]. In another study it was observed that after fast pranayama basal heart rate was reduced [23]. A similar study conducted in Puducherry, India and it was found that after yogic practices heart rate was reduced significantly in hypertensive patients [24].

In our study resting heart rate decreased significantly in the yoga intervention group but not in the control group. Vempati and Telles reported that the rate of respiration and heart

Characteristic	Control group		Yogic group	
	Pre ( <i>n</i> = 129)	Post ( <i>n</i> = 103)	Pre ( <i>n</i> = 129)	Post ( <i>n</i> = 105)
	Mean (SD)	Mean (SD)	Mean (SD)	Mean (SD)
BMI (kg/m²)	25.1 (4.8)	25.1 (4.6)	24.8 (3.8)	24.3 (3.5)
Waist:hip ratio	1.0 (0.1)	1.0 (0.1)	1.0 (0.1)	1.0 (0.1)
SBP (mmHg)	129.8 (14.1)	129.1 (9.3)	130.1 (10.3)	123.1 (9.4)
DBP (mmHg)	82.3 (8.6)	83.8 (5.7)	83.7 (8.0)	80.5 (5.1)
Heart rate (beats/min)	71.6 (11.4)	73.3 (8.7)	73.0 (9.1	70.5 (7.5)
Fall in SBP (mmHg) <sup>a</sup>	10.3 (4.3)	9.3 (4.2)	10.5 (4.4)	10.1 (4.3)
Increase in DBP after sustained hand grip (mmHg)	10.6 (6.1)	10.5 (5.6)	12.6 (4.8)	12.1 (4.7)
Expiration-inspiration ratio	1.2 (0.3)	1.1 (0.2)	1.1 (0.2)	1.2 (0.1) )
30:15 beat ratio	1.1 (0.2)	1.1 (0.2)	1.1 (0.3)	1.1 (0.2)
Valsalva ratio	1.6 (0.8)	1.5 (0.6)	1.5 (0.7)	1.6 (0.9)
LF/HF ratio	1.6 (1.3)	1.6 (1.2)	1.7 (1.3)	1.6 (1.2)

Table 2 Characteristics of patients with coronary	tery disease in control and intervention groups before and after a yo	ga
intervention study		5

<sup>a</sup>Differences in systolic blood pressure from the baseline (lying position) and after 2 minutes while standing.

SD = standard deviation; BMI = body mass index; SBP = systolic blood pressure; DBP = diastolic blood pressure; LF = low frequency; HF = high frequency.

rate decreased significantly after yogabased relaxation [25]. There was no significant change in the heart rate in the control group in our study. The results indicate that sympathetic reactivity reduced to some extent in yoga group, but not statistically significantly. This is in contrast with the findings of the study of Mourya et al. [26].

Yogic practices can affect very short term to body mass index and blood pressure levels. Body mass index and waist hip ratio are the physiological parameters for measuring obesity. Both the variables decreased significantly when compared with control. Both the studies got similar results after short term yogic intervention [27,28]. There is also evidence that om chanting effects changes in various biochemical levels such as endorphins, cortisol, adrenocorticotrophic hormone, interleukin-1, and immunoglobulin A, all of which affect the immune system. In a study in St Petersburg it was found that in subjects who adopted the cobra posture (bhujangasana), the production of steroid hormones was controlled by neural mechanisms [29].

Table 3 Comparison of difference in study scores before and after yogic intervention among yogic and control groups					
Characteristic	Difference from baseline		<i>P</i> -value		
	Control group	Yogic group			
	Mean (SD)	Mean (SD)			
BMI (kg/m <sup>2</sup> )	-0.03 (1.07)	0.52 (0.89)	0.0001**		
Waist:hip ratio	-0.004 (0.09)	0.02 (0.03)	0.005**		
Heart rate (beats/min)	-1.78 (9.13)	2.58 (8.79)	0.0006**		
SBP (mmHg)	0.70 (10.81)	5.41 (11.40)	0.002**		
DBP (mmHg)	-1.51 (8.69)	3.27 (8.01)	0.0002**		
Fall in SBP (mmHg)	0.98 (4.76)	0.36 (5.28)	0.38		
Sustained hand grip DBP (mmHg)	0.09 (4.57)	0.43 (4.55)	0.59		
Valsalva ratio	0.07 (0.60)	0.13 (0.07)	0.44		
Expiration-inspiration ratio	0.01 (0.31)	0.06 (0.29)	0.88		
Standing ratio	0.03 (0.22)	0.05 (0.26)	0.49		
LF/HF ratio	0.01 (1.18)	0.18 (1.12)	0.27		

\*\*Statistically significant (P < 0.01).

SD = standard deviation; BMI = body mass index; SBP = systolic blood pressure; DBP = diastolic blood pressure; LF = low frequency; HF = high frequency.

#### Limitations of the study

The limitations of this study include the fact that our study group was taken from only the urban population (the study was conducted in a medical university situated in urban area so it was not possible for members of the rural population to attend a yoga class of such long duration in the mornings). Parameters tested in this study were indirectly related to CAD; there were no parameters directly related. Measuring cardiac output, ECG monitoring, etc., however, did provide accurate information related to CAD. The diet factor plays a vital role in yogic interventional studies. However, in this study we did not take into consideration the effect of diet. Measuring the effects of diet is very complicated in outpatients; it is only possible in inpatients. In this study we requested that patients maintain their normal daily routine.

Qualitative analysis of the participants was also useful for discussing the parameters which changed after yogic intervention. This study may provide a base for clinicians and policy-makers indicating that yogic intervention could be prescribed as an adjunct to medical treatment.

#### Acknowledgements

We are very grateful to the entire yoga team for its coordination. We are also grateful to all the staff of the Departments of Physiology, Pathology and Cardiology at Chhatrapati Shahuji Maharaj Medical University, Lucknow, for their cooperation during the entire study.

**Funding:** This study received the financial support of the Department of AYUSH, Ministry of Health and Family Welfare, Government of India.

Competing interests: None declared.

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#### Global atlas on cardiovascular disease prevention and control: policies, strategies and interventions

As the magnitude of cardiovascular diseases (CVDs) continues to accelerate globally, the pressing need for increased awareness and for stronger and more focused international and country responses is increasingly recognized. The abovementioned atlas on cardiovascular disease prevention and control is part of the response to this need.

It documents the magnitude of the problem, using global cardiovascular mortality and morbidity data. It demonstrates the inequities in access to protection, exposure to risk, and access to care as the cause of major inequalities between countries and populations in the occurrence and outcome of CVDs. The report has graphs showing mortality rates of CVDs by age, by country/region, and is divided into three main sections: Section A: Cardiovascular diseases due to atherosclerosis; Section B: Other cardiovascular diseases; and Section C: Prevention and control of CVDs – Policies, strategies and interventions.

This publication can be found at: http://www.who.int/cardiovascular\_diseases/publications/atlas\_cvd/en/index. html