### Supplementary Study Summary Table 1. Effects of meditation on psychological, psychosocial, and physiological response to stress

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
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</thead>
</table>
| Momeni, J, J Am Soc Hypertension, 2016 (1) | • Single-blind randomized controlled trial to that assessed blood pressure, perceived stress, and anger in 60 cardiac patients.  
  • Experimental group (N=30) received 8 weeks MBSR training while those in control group received no psychological training (waitlisted control group N=30). | • Systolic blood pressure, perceived stress, and anger were significantly improved (p<.01) in MBSR group as compared to control group. | • Wait-listed control group  
  • Small sample size |
| Epel ES, Transl Psychiatry, 2016 (2) | • 97 Healthy women aged 30–60 entered into 1 week spa retreat: 30 regular meditators and 64 non meditators (31 vacation only and 33 vacation + meditation) with post intervention 1 and 10 month follow up | • Regular meditators had lower telomerase at baseline, and a significant increase in peripheral blood cell telomerase activity post treatment not observed in the other two groups. | • Small sample size  
  • Quasi controlled design (confounding of vacation with meditation) |
| Koncz, R, J Occup Environ Med, 2016 (3) | • University employees were randomized to a 6-week mindfulness-based stress release program (SRP) (N=50) or a waitlist control group (N=29)  
  • SRP program is a structured program consisting of “practices using body, breath, and cognitive strategies and reflective activities to enhance professional and personal life” and is considered to be less intensive than MBSR  
  • Perceived stress, workplace well-being, and engagement were measured at baseline and at completion of the program | • Participants in SRP group had significant improvements in level of distress [-3.0 (95% CI -5.5 to -0.6  
  p=0.02), university workplace wellbeing (2.5, 95% CI 0.5 to 4.5,  
  p=0.02), and vigor (0.39, 95% CI 0.65 to 3.07; p <0.01) at follow-up compared with baseline. No improvements were observed in control group. | • Wait-listed control group  
  • Attendance at sessions and practice time not assessed. |
| Himashree, G., et al. Altern Ther Health Med 2016 (4) | • 200 soldiers fully acclimatized to high altitude were randomized to routine physical training activities vs. comprehensive yoga package (physical asanas, pranayama, and meditation) | • The yoga group had lower body fat %, respiratory rate, DBPs, and anxiety scores. They had higher EtCO2, forced vital capacity, forced expiratory volume in the first second (FEV1), and VO2Max. Also, the yoga group showed a significant reduction in serum cholesterol, LDL, and triglycerides. | • Greatest benefit in those markedly hypertensive (SBP>160 mmHg)  
  • No long-term f/u of durability of benefit |
<table>
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<tr>
<th>Study</th>
<th>Design</th>
<th>Participants</th>
<th>Outcomes</th>
<th>Notes</th>
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</thead>
<tbody>
<tr>
<td>Bharshankar JR, et al. Indian J Physiol Pharmacol. 2015 (5)</td>
<td>Case control study of 50 Raja-yoga meditators practicing meditation for 5 years and 50 age matched non-meditators.</td>
<td>Mean resting HR, SBP and DBP were less in meditators. Galvanic Skin Response in meditators was significantly more (p &lt; 0.001). Mean increase BP response to Hand Grip Test and Cold Pressor Test was significantly less in meditators than non-meditators (p &lt; 0.001).</td>
<td>• Small sample size&lt;br&gt;• Results, suggest a shifting of the autonomic balance to parasympathetic side in Raja-yoga meditators, which suggests meditation combat the ill effects of stress.</td>
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<tr>
<td>Black, DS, JAMA Intern Med, 2015 (6)</td>
<td>Randomized clinical trial&lt;br&gt;Participants (older adults with sleep disturbances) randomized to a standardized mindful awareness practice (MAP) intervention (N=24) or sleep hygiene education (SHE) intervention (N=25).&lt;br&gt;Each group met for 2 hours per week for 6 weeks with assigned homework</td>
<td>Participants in the MAP group had better improvements in sleep than the SHE group.&lt;br&gt;MAP group participants reported significant improvements relative to the SHE group in health outcomes, depressive symptoms, and levels of fatigue.&lt;br&gt;NF-kappa B concentrations significantly declined over time for both groups</td>
<td>• Small sample size&lt;br&gt;• Case control design</td>
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<tr>
<td>Bower, JE, Cancer, 2015 (7)</td>
<td>Randomized trial that examined brief mindfulness-based intervention for younger female breast cancer survivors.&lt;br&gt;Women diagnosed with early stage breast cancer at or before age 50 who had completed cancer treatment were randomly assigned to a 6-week Mindful Awareness Practices (MAPS) intervention group (n = 39) or to a wait-list control group (n = 32).&lt;br&gt;Participants completed questionnaires before and after the intervention to assess stress and depressive symptoms (primary outcomes) as well as physical symptoms, cancer-related distress, and positive outcomes. Blood samples were collected to examine genomic and circulating markers of inflammation. Participants also completed questionnaires at a 3-month follow-up assessment.</td>
<td>Perceived stress reduced (p=.004) post intervention,&lt;br&gt;Decreased pro-inflammatory gene expression (P = .009) and inflammatory signaling (P = .001) at post intervention.&lt;br&gt;Intervention effects on psychological and behavioral measures not maintained at 3 month follow up.</td>
<td>• Small sample size&lt;br&gt;• Use of wait-list control group</td>
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<tr>
<td>Authors</td>
<td>Journal</td>
<td>Study Details</td>
<td>Findings</td>
<td>Comments</td>
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<td>Schutte, NS, Psychoneuroendocrinology, 2016 (8)</td>
<td>Meta-analysis of 4 clinical trials (total N=191) examining telomerase activity in association with meditation.</td>
<td>A meta-analytic effect size of $d = 0.46$ indicated that mindfulness meditation leads to increased telomerase activity in peripheral blood mononuclear cells. These results</td>
<td>Small number of studies included in analysis.</td>
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<tr>
<td>Younge, JO, PLoS One, 2015 (9)</td>
<td>Randomized controlled single-blind trial that examined the physiological and psychological outcomes of a 12 weeks online mindfulness training program (N=215) as compared to usual care (N=109) in patients with cardiac disease. Primary outcome was exercise capacity as measured by a 6 min walk test. Other outcomes included heart rate, blood pressure, respiratory rate, NT-proBNP, subjective health status, perceived stress, psychological well-being, social support, and a composite end-point (all-cause mortality, heart failure, symptomatic arrhythmia, cardiac surgery, and percutaneous cardiac intervention).</td>
<td>Compared to the control group, participants in the online MBSR program demonstrated improved exercise capacity (effect size; 13.2, 95% CI: -0.02-26.4, $p=0.050$). Participants in the MBSR group also had a lower heart rate (effect size, beats per minute;-2.8, 95% CI: -5.4;0.2, $p=0.033$)</td>
<td>No control group</td>
<td></td>
</tr>
<tr>
<td>Azam, MA, Int J Psychophysiology, 2015 (10)</td>
<td>Stratified-randomized trial. Following a laboratory cognitive stressor, participants (maladaptive perfectionists – N=21, and controls – N=39, were randomly assigned to a 10 min audio instructed mindfulness meditation condition or a 10 min rest condition with audio description of mindfulness meditation.</td>
<td>Significant elevated heart rate variability (HRV) during meditation for controls but not for maladaptive perfectionists. Findings suggest that mindfulness meditation promotes relaxation following cognitive stress but the maladaptive perfectionist personality hinders relaxation possibly due to decreased cardiac vagal tone</td>
<td>Small sample size. 10 minutes of audiotaped guided meditation may not be sufficient</td>
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<tr>
<td>de Fátima Rosas Marchiori M, et al. Geriatric Gerontol Int., 2015 (11)</td>
<td>59 volunteers, aged ≥60 years with SBP 130-159 mmHg and DBP 85-99 mmHg, were randomly divided into meditation twice a day for 20 min for 3 months vs. control wait-list control</td>
<td>SBP, CRP and IL-6 levels did not differ between groups. QOL improved in psychological aspects (e.g. loneliness) and overall in the meditation group vs. control.</td>
<td>Small sample size. No change in physiologic parameters</td>
<td></td>
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</table>
Carlson, LE, Cancer, 2015 (12)  
- Randomized controlled trial comparing Mindfulness Based Cancer Recovery program to a supportive-expressive group. A one day stress management seminar was used for the control group.  
- 88 distressed cancer survivors with a diagnosis of stage 1 or stage 11 cancer who completed treatment at least three months prior participated.  
- No differences were found in regards to telomere length between the mindfulness group and the supportive expressive group – but a trend was observed in the combined intervention group as compared to the control group ($F(1,84) = 3.82, p=.054$, $\eta^2=.043$).  
- No associations were found between changes in telomere length and changes in mood or stress scores over time.  
- Small sample size  
- Control group consisted only of a one-day stress management seminar.

Cash, E. Ann Behav Med, 2015 (13)  
- Randomized clinical trial of MBSR in women with fibromyalgia  
- Examined pain, perceived stress, sleep quality, fatigue, symptom severity, and salivary cortisol at baseline, post-program, and 2 month follow up  
- 51 women in treatment group, 40 in wait-list control group  
- MBSR significantly reduced perceived stress, symptom severity, and sleep disturbances with changes sustained at follow up.  
- MBSR did not change level of pain, physical functioning, or cortisol profile.  
- Frequency of home MBSR practice significantly associated with greater symptom relief.  
- Small sample size  
- High attrition rate (attendance fell over 33% from first session to fourth session)

Creswell, JD, Psychoneuroendocrinology, 2014 (14)  
- Examined the extent to which a brief mindfulness training intervention buffered self-reported psychological and cortisol responses to TSST in 66 young adults.  
- Participants were randomly assigned to either a 3 day (25 min per day) mindfulness meditation training or an analytic cognitive training control program.  
- Controlled for treatment expectancies.  
- Perceived stress was reduced in brief mindfulness training group however demonstrated increased cortisol reactivity to TSST as compared to the control group.  
- No changes were observed in systolic or diastolic blood pressure between the two groups.  
- Did not include a validated state mindfulness measure  
- Since blood pressures were not taken continuously during/after the TSST, BP reactivity may not have been fully captured.
Arch, JJ, Psychoneuroendocrinology, 2014 (15)  
- Examined the extent to which a brief self-compassion training program (consisting of 10 min. recordings listened to daily for 3 days) moderated biophysiological responses to the Trier Social Stress Test in women (N=105).  
- Compared intervention to attention and no training control conditions  
- Collect salivary cortisol, salivary alpha amylase, and heart rate variability in response to Trier Social Stress Test  
- Brief self-compassion training attenuated sympathetic, cardiac parasympathetic, and subjective anxiety to Trier Social Stress test as compared to attention and no training control conditions  
- No differences were noted in cortisol response to the TSST between the self-compassion group and the control groups.  
- Small sample size

Kaliman, P. Psychoneuroendocrinology, 2014 (16)  
- Examined impact of a day of intensive mindfulness meditation in experienced individuals (N=19) on expression of circadian, chromatin modulatory, and inflammatory genes in peripheral blood mononuclear cells compared to a control group (N=21) of individuals with no meditation experience who engaged in leisure activities in the same environment as intervention group  
- Blood was collected before and after the intervention for analysis of gene expression. In addition, individuals underwent the Trier Social Stress Test (TSST).  
- Core clock gene expression at baseline was similar between groups and their rhythmicity was not influenced by meditation.  
- Epigenetic regulatory enzymes and inflammatory genes were similar at baseline for the two groups.  
- Reduced expression of histone deacetylase genes (HDAC 2, 3, and 9), alterations in global modification of histones (H4ac;H3Lme3), and decreased expression of pro-inflammatory genes (RIPK2 and COX2 were found in meditators as compared to controls.  
- Faster recovery of cortisol levels after the TSST was associated with lower gene expression levels of RIPK2 and HDAC2.  
- Small sample size

Lengacher CA, Biol Res Nursing 2014 (17)  
- 162 breast cancer survivors were randomized or wait-listed.  
- 6 week Mindfulness-based stress reduction (MBSR) on telomere length (TL) and telomerase activity (TA) at 6 and 12 weeks.  
- MBSR led to increased telomerase activity but no increase in telomere length  
- Small-modest sample size  
- Mindfulness-based stress reduction influenced telomerase activity in women with breast cancer
<table>
<thead>
<tr>
<th>Authors</th>
<th>Description</th>
<th>Findings</th>
<th>Limitations</th>
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<tbody>
<tr>
<td>Lipschitz, DL,</td>
<td>Randomized controlled trial of 57 cancer survivors with sleep disturbances. Participants were randomized into either a sleep hygiene education control group (n=18) or a Mind-Body Bridging program (N=19) or a Mindfulness Meditation program (N=20).</td>
<td>Mean sAA upon awakening levels declined in the Mind-Body Bridge group as compared to the Sleep Hygiene Education group. Self-reported sleep improved in all three interventions with largest improvements demonstrated in Mind-Body Bridge group. Cortisol levels were not altered by any of the interventions.</td>
<td>Small sample size, Saliva samples were collected over the course of a single day rather than 2 or 3 consecutive days, Intervention was only three weeks long.</td>
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<td>Psychoneuroendocrinology, 2013 (18)</td>
<td>The Mind-Body Bridge program is a program that teaches individuals how to become aware of dysfunctional mind-body states. Each intervention consisted of one session per week for three consecutive weeks. Saliva cortisol and serum alpha amylase (sAA) measured at baseline and one week after last session.</td>
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<td>Malarkey, WB,</td>
<td>Controlled randomized of university faculty and staff at risk for cardiovascular disease (N=186) comparing a low dose Mindfulness Based Intervention group to an active control group receiving lifestyle education program. Low dose Mindfulness Based Intervention (MBI-ld) consisted of one hour sessions for 8 consecutive weeks. Participants were expected to practice 20 minutes per day.</td>
<td>The low dose Mindfulness Based Intervention significantly improved mindfulness post intervention and this change was sustained 1 year later as compared to the education group. No significant changes were found between groups in regards to cortisol, IL-5, or self-reported measures of stress, depression, or sleep quality.</td>
<td>Did not compare low dose MBSR to traditional MBSR program</td>
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<td>Brain Behav Immun, 2013 (19)</td>
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</table>
### Bhasin MK, PLoS ONE 2013 (20)
- Prospective study of 26 healthy subjects who had no prior relaxation response training (RR; diaphragmatic breathing, mantra repetition, and mindfulness meditation)- eliciting experience (Novices, N1) who underwent 8 weeks of RR-eliciting training (Short-term Practitioners, N2).
- Parallel cross-sectional study of another 26 healthy subjects with significant prior RR- practice (4–20 years; Long-Term Practitioners, M) and compared with novices either before or after their 8-week RR training.
- Both short-term and long-term practitioners evoked significant temporal gene expression changes with greater significance in the long-term practitioners as compared to novices. RR practice enhanced gene expression related to energy metabolism, mitochondrial function, insulin secretion and telomere maintenance, and reduced expression of genes linked to inflammatory response and stress-related pathways.
- Small sample size
- Quasi-experimental design
- Relaxation response induces temporal transcriptome changes in energy metabolism, insulin secretion and inflammatory pathways. Some genes were modified only in long-term practitioners, whereas others were modified in both short- and long-term practitioners with a greater intensity in the latter.

### Rosenkranz, MA, Brain Behav Immun, 2013 (21)
- Randomized controlled trial (N=49) comparing 8 weeks MBSR program to Health Enhancement Program (HEP)
- Psychological stress and endocrine response were measured before and after the Trier Social Stress Test.
- Inflammation was measured by using capsaicin topical cream to induce inflammation before and after intervention.
- Cortisol responses to TSST were similar between MBSR and HEP groups.
- Reduction in psychological distress and symptoms in response to TSST were similar between groups.
- Those randomized to MBSR group had significantly smaller post-stress inflammatory response as compared to HEP.
- Generalizability to populations with chronic illness
- Unable to determine interaction between stress and inflammation because, due to potential participant burden, stress condition was not employed in the absence of inflammation and an inflammation condition was not tested in the absence of stress.

### Qu S, PLoS ONE 2013 (22)
- Ten health adults did two courses of 4 consecutive days of a comprehensive yoga program, at the same time of the day (6.30 am – 8.30 am) or yoga and related practices or nature walk with relaxing music.
- Gene expression changes were noted as early as 2 hours. 97 unique genes were affected by yoga and related practices vs. 24 by the control regimen. 36% of the control group genes were also influenced by the yoga regimen, suggesting overlap in effect on biological processes.
- Small sample size
- Intervention is poorly described
- Rapid gene expression changes in peripheral blood lymphocytes upon practice of a comprehensive yoga program.
<table>
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<tr>
<th>Author</th>
<th>Journal</th>
<th>Study Design</th>
<th>Findings</th>
<th>Notes</th>
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</table>
| Nyklicek, L             | Health Psychol, 2013 (23) | • 88 community-dwelling volunteers reporting elevated levels of perceived stress were randomly assigned to a MBSR program or waitlist control group.  
• Participants underwent a social stressor consisting of mental math and making a speech before and after the intervention.  
• Measurements before, during and after the social stressor included heart rate variability, blood pressure, and salivary cortisol. | • Controlling for age, body mass index, and beta blockers, participants in the MBSR group demonstrated larger decreases in systolic blood pressure, diastolic blood pressure.  
• No effect was obtained for other physiological measures. | • Absence of critical committee during TSST could have reduced cortisol response to stressor.  
• Recovery period after social stressor was only 10 minutes which may have limited the ability to capture changes during recovery. |
| Carlson, J              | J Clin Oncol, 2013 (12)  | • Randomized controlled trial to compare mindfulness–based cancer recovery (MBCR) program to supportive-expressive group therapy (SET) in distressed survivors of stage 1 to III breast cancer (N=271)  
• 1-day stress management class was used as a control condition  
• Measures (mood, diurnal salivary cortisol, stress, quality of life, and social support) were collected at baseline and after the intervention by evaluators blinded to the study condition. | • Cortisol slopes were maintained in MBCR (p=.011) and SET group (p=.002) participants in comparison to those in the control group whose cortisol slopes became flatter.  
• Stress symptoms were improved in MBCR group compared to SET (p=.009) and control (p=.024) groups.  
• Those participating in the MBCR group demonstrated greater improvements in quality of life compared to the control group (p=.005) and social support compared to those in the SET group (p=.012). | • Only breast cancer patients in study –findings may not be generalizable.  
• High attrition (34.5% in MBCR group). |
| Jacobs, TL              | Health Psychol, 2013 (24)  | • Observational study examining self-reported mindfulness and evening cortisol at the beginning and after a 3 month Shamatha meditation retreat (N=57)  
• The group met 2x/day for 1-hr sessions to engage in guided meditations and dialogue but primarily practiced solitary meditation for much of the day (M = 6.3 hr/day, SD = 1.34). | • Mindfulness increased from pre-retreat (M = 5.16, SD = .77) to post-retreat(M= 5.76, SD=.72), F(1, 56) = 36.20, p=.001  
• Cortisol levels did not change  
• Mindfulness was inversely related to cortisol levels both pre and post retreat. | • To allow for acclimatization, cortisol measures were taken 2 weeks after arrival to the retreat site, which meant that participants had already been meditating for up to 9 days before the initial cortisol measure was obtained. |
Levine et al. Meditation and Cardiovascular Risk Reduction: A Scientific Statement From the American Heart Association  
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<tr>
<th>Reference</th>
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<th>Findings</th>
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</table>
| Jensen, J Exp Psychol Gen, 2012 (25) | Blinded design - 48 young, healthy meditation novices were randomly assigned to MBSR, non-MBSR, or inactive control group.  
- At posttest, those in the inactive control group were randomly split into incentive and non-incentive controls  
- Attention, self-report of mindfulness, perceived stress, and salivary cortisol were measured at pre and post intervention. | Attentional effects of MBSR, non-MBSR, and the financial incentive were comparable or significantly larger in the incentive group.  
- Selective attention improved significantly more in the MBSR group than non-MBSR and inactive control group. F(6, 84)=2.30, p=.052.  
- Conscious perception and visual working memory capacity were only improved in the MBSR group F(1, 22)= 7.31, p=.05)  
- MBSR participants had significant reduction of perceived stress (p=.04) and improvement in salivary cortisol levels (p<.05). |
| Creswell JD, Brain Behav Immun 2012 (14) | 40 healthy older adults (mean age 65 years) in a 8 week randomized controlled trial, Mindfulness-Based Stress Reduction (MBSR) program vs. wait-list control | MBSR downregulated NF-κB gene expression profile & a trend to reduce C Reactive Protein  
- Small sample size  
- MBSR training reduced loneliness and proinflammatory gene expression in older adults |
| Matousek, RH, Complement Ther Clin Pract 2011 (26) | 33 women who had completed treatment for breast cancer participated in the study  
- MBSR group met weekly for 2.5 hours for 8 consecutive weeks  
- Cortisol Awakening Response (CAR) was assessed at three days prior to the MBSR program and three days after.  
- Depressive symptoms, perceived stress, and medical symptoms were measured pre and post intervention. | Cortisol levels demonstrated a prolonged increase after awakening at the post MBSR assessment. This was accompanied by significant improvements in self-reported stress, depressive symptoms, and medical symptoms.  
- Small sample size  
- No control or comparison group |
### Jacobs TL, Psychoneuroendocrinology 2011; 36: 664–681 (27) (27)

- 3-month meditation retreat (concentrative meditation techniques and complementary practices used to cultivate benevolent states of mind: 30 active and 30 matched waitlist) on telomerase activity and two measures of stress: Perceived Control and Neuroticism

- Telomerase activity was significantly greater in retreat participants than in controls at the end of the retreat
- Retreat group: increases in Perceived Control, decreases in Neuroticism, and increases in both Mindfulness and Purpose in Life were greater in the

### Klatt, MD, Health Educ Behav, 2009 (28)

- University employees and staff randomized to a low dose MBSR program (N=24 or wait list control group (N=24).
- Low dose MBSR program consisted of 1 hour weekly sessions for 6 consecutive weeks with 20 minutes of daily practice.
- Perceived stress, sleep quality and mindfulness assessed at baseline and at end of 6 weeks intervention.
- Salivary cortisol was collected three times a day for 2 consecutive days every week for the duration of the intervention.

- Participants in the low dose MBSR group had significant reduction of perceived stress (p=.0025) and increase in mindfulness (p=.0149).
- No changes in average daily salivary cortisol levels over time for participants in both groups and no differences from the pretest to the posttest were found.

### Pace, TW, Psychoneuroendocrinology, 2009 (29)

- Examined the effect of compassion meditation on innate immune, neuroendocrine, and behavioral responses to psychosocial stress and examined the degree to which meditation practice influenced stress reactivity in 61 healthy adults randomized to 6 weeks of training in compassion meditation (N=33) or in a health discussion control group (N=28).
- Response to TSST was measured by repeated measures of interleukin -6 (IL-6), cortisol and total distress scores on the Profile of Mood States (POMS).

- No main effect of group assignment on TSST responses was found for IL-6, cortisol, or POMS scores.
- Increase meditation practice was correlated with decreased TSST-induced IL-6 (p=.0008) and POMS distress scores (p=.014).

- Small sample size
- Did not perform TSST prior to intervention – may be possible that individuals who had reduced inflammatory response to social stress may have been more willing or able to engage in meditation practice.
- Those randomized to the meditation group may have had higher expectations of outcomes than those randomized to the control group.
<table>
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<tr>
<th>Source</th>
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<tbody>
<tr>
<td>Witek-Janusek, Brain Behav Immun, 2008 (30)</td>
<td>Non randomized controlled design to evaluate the effect of MBSR on immune function, quality of life, and coping in women recently diagnosed with breast cancer. Participants self-selected into the MBSR group (N=44) or control group (usual care) (N=31). Data was collected from a cancer free group of women (N=30) for comparison of immune measures. Over time, women in MBSR group re-established NKCA and cytokine production levels while women in the usual care group demonstrated continued reductions in NKCA and IFN-gamma. IL4, IL-6, and IL-10 increased. Women in MBSR group had reduced cortisol levels and improved QOL and coping effectiveness compared to those in the usual care group. Small sample size. Non-randomization of participants.</td>
</tr>
<tr>
<td>Carlson, LE, Brain Behav Immun, 2007 (31)</td>
<td>49 women with breast cancer and 10 men with prostate cancer were enrolled in an 8 weeks MBSR program. Health behaviors, quality of life, mood, stress, salivary cortisol levels, immune cell counts, intracellular cytokine production, blood pressure and heart rate were assessed at baseline, post-intervention, and 6 and 12 months post intervention. Symptoms and stress were significantly improved after the intervention and improvements were maintained at 12 months post intervention. Cortisol and proinflammatory cytokines decreased over follow up period. Blood pressure significantly decreased from baseline to post intervention. No control or comparison group. Multiple statistical comparisons.</td>
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<tr>
<td>Tang, YY, Proc Natl Acad Sci USA, 2007 (32)</td>
<td>Randomly assigned undergraduate Chinese students to 5 days of meditation practice with integrative body–mind training (20 minutes per day) (N=40) or 5 days of relaxation training (20 minutes per day) (N=40). Those in meditation practice group demonstrated improved attention (p&lt;.01), lower anxiety (p&lt;.01), depression (p&lt;.05), anger (p&lt;.05), and fatigue (p&lt;.01). Cortisol response to 3 min of mental arithmetic was significantly lower in the meditation group than the relaxation group (p&lt;.01) after 20 minutes of practice. Small sample size.</td>
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<td>Author(s)</td>
<td>Journal</td>
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| Jain, S. | *Annals of Behavioral Medicine*, 2007 (33) | Randomized controlled trial examining the effects of a 1-month mindfulness meditation to a somatic relaxation training to a control group in 83 students. | • Both meditation and relaxation groups demonstrated improvements in mood and decreases in distress when controlling for social desirability as compared to the control group (p>.05).  
• Meditation group demonstrated significant pre-post decreases in distractive and ruminative thoughts compared to control group (p<.04). | Small sample size |
| Robert MacComb, JJ | *JJ Altern Complement Med*, 2004 (34) | Women with documented histories of cardiovascular disease were randomly assigned to a MBSR group (N=9) or control group (N=9).  
Pre-post hormonal measures and physical functioning were collected  
Submaximal exercise responses were measured after the 8 weeks intervention. | • No significant main effects or interaction for resting levels of stress hormones or physical functioning, or submaximal exercise responses  
• Significant differences in breathing patterns (p<.01). | Small sample size |
| Speca, M. | *Psychosomatic Medicine*, 2000 (35) | 90 outpatient cancer patients were randomized to a weekly meditation group lasting 1.5 hours or 7 weeks with home meditation practice or a wait-list control group.  
Participants completed the Profile of Mood States and the Symptoms of Stress Inventory before and after the intervention | • Participants in meditation group reported significantly lower total mood disturbance and fewer overall symptoms of stress (p<.05). | Wait list control group |
Supplementary Study Summary Table 2. Effects of meditation on blood pressure

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
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</table>
| Bai Z, J Human Hypertension, 2015 | MA of 12 RTC’s of TM versus control for effect on blood pressure  
Total 996 patients | TM improved BP 4.26/2.33 mmHg when compared to control P<.05 | The completion rate was <75% in 6 of 11 studies. Dropouts might have enhanced tendency to favor TM  
Only one study reported on all primary and secondary outcomes with intention-to-treat analysis,  
The efficacy of TM on BP tended to decrease with the study durations |
| de Fátima Rosas Marchiori M, Geriatr Gerontol Int., 2015 | RCT of twice-daily meditation for 20 min for 3 months vs. wait-list control  
59 volunteers, aged ≥60 years with SBP 130-159 mmHg and DBP 85-99 mmHg | At one month SBP was lower in meditation group but at 3 months BP did not differ | Small sample size  
No change in physiologic parameters at end of study |
| Blom, Am J Hypertension, 2014 | RTC of 8 weeks of mindfulness meditation on 24 hour BP control  
101 subjects (38% male) | Decrease in 24 hour BP of 0.4 mmHg in both treatment and control (wait list group)  
No significant between group reductions in blood pressure |  |
| Hughes JW, Psychosomatic medicine. 2013 | RCT of mindfulness-based stress reduction (MBSR) vs progressive muscle relaxation (PMR) over 8 weeks  
56 pre-hypertensive adults (50.3 years of age, on no BP meds). Clinic BP was the primary outcome. | In an intention to treat, clinic SBP fell 4.8 mmHg with MBSR vs 0.7 mmHg with PMR (P=0.016) | Small sample size  
| Schneider RH, Circ Cardiovasc Qual Outcomes. 2012 | RCT 201 adults with coronary artery disease treated with a TM program or health education | Systolic blood pressure fell 4.9 mmHg in TM vs. control (P=0.01) | Blood pressure reduction was a secondary outcome |
Levine et al. Meditation and Cardiovascular Risk Reduction: A Scientific Statement From the American Heart Association © 2017 The Authors. Published on behalf of the American Heart Association, Inc., by Wiley. This is an open access article under the terms of the Creative Commons Attribution-NonCommercial-NoDerivs License, which permits use and distribution in any medium, provided the original work is properly cited, the use is non-commercial and no modifications or adaptations are made.

<table>
<thead>
<tr>
<th>Palta P, J Urban Health, 2012</th>
<th>8 week RTC on mindfulness meditation versus social support on BP control</th>
<th>12 intervention and 8 control subjects</th>
<th>there was a 11/4 mmHg decrease in systolic/diastolic blood pressure in those randomized to 8 weeks of treatment</th>
<th>there was an analysis adjusted significant 22/17 mmHg difference in blood pressure between the two groups at follow-up</th>
<th>Small number of patients studied</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gregoski MJ, et al. J Adolesc Health. 2011</td>
<td>Randomized trial - breathing awareness meditation (BAM), Botvin Life Skills Training (LST), and health education control (HEC)</td>
<td>Study population consisted of 166 normotensive African American adolescents</td>
<td>BAM had greatest reduction in SBP and SBP, DBP and HR over the 24-hour period, overnight and during school hours. (Bonferroni adjusted, p &lt;0.05)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Nidich SI, Am J Hypertens. 2009</td>
<td>Randomized trial in 298 university students treated with a transcendental meditation program or wait-list control.</td>
<td>3 month intervention</td>
<td>Overall no difference in SBP &amp; DBP between groups.</td>
<td>In hypertension risk subgroup (n=112), SBP fell 5 mmHg with TM compared to increased 1.3 mmHg for control (P= 0.014)</td>
<td>Hypertension risk subgroup was a secondary analysis</td>
</tr>
<tr>
<td>Anderson, JV, Am J Cardiol, 2008</td>
<td>MA of RTC of TM that randomly assigned individuals to different target BP levels</td>
<td></td>
<td>Transcendental Meditation, compared to control, was associated with the following changes: -4.7 mm Hg (95% confidence interval (CI), -7.4 to -1.9 mm Hg) and -3.2 mm Hg (95% CI, -5.4 to -1.3 mm Hg)</td>
<td></td>
<td>Study designs and BP methods of blood pressure measurement, as well as dropout rates, limit the extrapolation of results</td>
</tr>
<tr>
<td>Manikonda JP, J Hum Hypertens. 2008</td>
<td>8 week pilot study of either contemplative meditation combined with breathing techniques (CMBT) or no intervention in this</td>
<td>observer-blind design</td>
<td>SBP after 8 weeks of meditation fell 15 mm Hg (vs 3 mm Hg in controls (P&lt;0.0001)</td>
<td></td>
<td>Small sample size • Short duration</td>
</tr>
</tbody>
</table>

- Systematic review and MA of stress reduction therapies
- Seventeen trials of 960 participants with elevated BP

| Reductions in blood pressure with Transcendental Meditation were 5.0/2.8 mmHg (systolic/diastolic); p=0.002 (systolic) and p=0.02 (diastolic)
| No significant reductions in blood pressure with biofeedback, relaxation-assisted biofeedback, progressive muscle relaxation, and stress management training.
| Review did not study other forms of meditation

-5.0/-2.8 mm Hg (P = 0.002/0.02)
Supplementary Study Summary Table 3. Effects of meditation on smoking and tobacco use

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
</tr>
</thead>
</table>
| Oikonomou, Journal of health Psychology, 2016 (36) | • Meta-analysis of 4 randomized controlled trials  
• Studies between 2011-2014, included 474 patients  
• Included studies examined smoking abstinence in short (4–6 weeks) and long term (17–24 weeks) comparing mindfulness training for smokers to a control group | • 25.2% of participants in the mindfulness group remained abstinent in the long term (17-24 weeks) compared to 13.6% of those who received usual care therapy (RR: 1.88; 95% confidence interval, 1.04-3.40)  
• No significant differences were found in the short term (4-6 weeks) (RR: 1.52; 95% CI, 0.95–2.45)  
• Small number of studies included in the meta-analysis (4), of which 3 are conducted by the same author | |
| Ruscio, Nicotine & Tobacco Research, 2016 (37) | • Randomized controlled trial of a brief mindfulness practice (Brief-MP) intervention on self-reported smoking behavior delivered to smokers on a Personal Digital Assistant (PDA) in the field  
• Participants carried a PDA for 2 weeks and were instructed to initiate 20 minutes of meditation (or control) training on the PDA daily (n=24 (MP) vs. n=20 (control)) | • Brief-MP (vs. Control) reduced overall negative effect, reduced craving immediately post-meditation and reduced cigarettes smoked per day over time | • Small sample size  
• 27% of the participants did not complete the study (12 of 44)  
• The study advertisements mentioned meditation, which could have appealed to individuals who were more educated, more interested in meditation, or more motivated to quit smoking, and reduces the generalizability of the findings |
| Davis JM, BMC Complement Altern Med, 2015 (38) | • Prospective observational study  
Participants (n=26) were asked to watch eight classes of web-based video instruction describing mindfulness skills and how to use these skills to overcome various core challenges in tobacco dependence  
Participants received eight weekly phone calls from a smoking cessation coach who provided general support and answered questions about the videos | • 7-day point prevalence smoking abstinence at 4 and 6-months post-quit of 23.1% and 15.4% respectively  
Participants showed a significant pre- to post-intervention increase in mindfulness as measured by the Five-Factor Mindfulness Questionnaire  
Participants also demonstrated a significant pre- to post-intervention decrease in the Anxiety Sub-scale of the Depression Anxiety and Stress Scale | • Small sample size  
• Lack of control group  
• Possible selection bias as participants were required to have internet access |
| --- | --- | --- | --- |
| Davis JM, J Subst Abuse Treat, 2014 (39) | • Randomized controlled trial comparing mindfulness training to a matched control based on the American Lung Association’s Freedom From Smoking program  
135 low socioeconomic status smokers were randomized to Mindfulness Training for Smokers (MTS) or Freedom from smoking-enhanced (FFS-E)  
Participants in the MT and FFS-E groups received 24 hours of instruction in each group | • Intent-to-treat analysis of 7-day point prevalence abstinence between two groups was almost identical at 4 weeks (MTS = 35.3%; FFS-E = 34.3%; p = 1.00, OR = 1.04, CI = 0.51-2.19)  
At 24 weeks, MTS compared to FFS-E showed higher numerical abstinence rates (MTS = 25.0%; FFS-E = 17.9%; p = 0.35, OR = 1.53, CI = 0.67-3.51), but failed to reach statistical significance  
Mindfulness training was associated with decreased urges, increased mindfulness, and decreased stress and experiential avoidance | • Participants were not blinded to their respective treatments  
• Intervention attrition was 32.4% in the MTS group and 26.9% in the FFS-E group |
<table>
<thead>
<tr>
<th>Study Reference</th>
<th>Study Design and Findings</th>
</tr>
</thead>
<tbody>
<tr>
<td>Davis JM, Subst Use Misuse, 2014 (40)</td>
<td>Randomized trial comparing mindfulness training for smokers (MTS) to a usual care therapy (Controls), which included the availability of a tobacco quit line and nicotine patches. 198 low socioeconomic status smokers were randomized to MTS or Quit Line. Primary outcome measure of the study (7-day point-prevalence abstinence at 4 and 24-weeks post-quit) did not reach statistical significance in an intent-to-treat analysis. Did reach statistical significance comparing treatment initiators at 4-weeks (MTS = 45.8%, Controls = 25.4%) and at 24-weeks (MTS = 38.7%, Controls = 20.6%, OR = 2.33, p = .05). The study showed high pre-intervention attrition and high 24-week assessment visit attrition. Participants not blinded to their respective interventions. Compares MTS to a less intensive usual-care therapy and as such lacks a time/intensity matched control.</td>
</tr>
<tr>
<td>Tang YY, Proc Natl Acad Sci USA, 2013 (41)</td>
<td>Healthy college students recruited through campus advertisements for learning meditation/relaxation to reduce stress and improve cognitive performance. Randomized to Integrated body-mind (IBMT) technique for meditation (15 smokers and 18 non-smokers) vs. Relaxation (RT) technique (12 smokers and 15 non-smokers). The participants received 30-min of IBMT or RT group practice every night for 10 consecutive sessions, for a total of 5 h of training. Among smokers, meditation training produced a significant reduction in smoking of 60%; no reduction was found in the relaxation control. Resting-state brain scans showed increased activity for the meditation group in the anterior cingulate and prefrontal cortex, brain areas related to self-control. Small study. Recruitment via advertisements, so possible selection bias. Participants with the goal of quitting smoking were not included.</td>
</tr>
</tbody>
</table>
Brewer, Drug Alcohol Depend, 2011 (42)  
- 88 smokers were randomly assigned to receive Mindfulness Training (MT) or the American Lung Association’s Freedom From Smoking (FFS) treatment  
- Both treatments were delivered twice weekly over four weeks (eight sessions total) in a group format  
- 88% of individuals who received MT and 84% of individuals who received FFS completed treatment  
- Compared to those randomized to the FFS intervention, individuals who received MT showed a greater rate of reduction in cigarette use during treatment and maintained these gains during follow-up  
- They also exhibited a trend toward greater point prevalence abstinence rate at the end of treatment (36% vs. 15%, p = .063), which was significant at the 17-week follow-up (31% vs. 6%, p = .012).  
- First randomized clinical trial to evaluate the efficacy of Mindfulness Training as a stand-alone treatment for smoking cessation compared to an active, empirically-supported control condition  
- Exclusion of individuals using psychoactive medications  
- Single site study

Davis JM, BMC Complement Altern Med, 2007 (43)  
- Pilot study designed to test the feasibility of using Mindfulness Based Stress Reduction (MBSR) as a smoking intervention  
- At the 6-week post-quit visit, 10 of 18 subjects (56%) achieved biologically confirmed 7-day point-prevalent smoking abstinence  
- Small sample size  
- Short follow-up  
- Lack of control group  
- 28% (5 of 18 subjects) attrition rate  
- No concurrent use of pharmacotherapy for smoking cessation
Supplementary Study Summary Table 4. Effects of meditation on insulin resistance and metabolic syndrome.

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
</tr>
</thead>
</table>
| Paul Labrador, Archives of Internal Medicine, 2006 (44) | • Randomized control trial with 103 subjects diagnosed with coronary artery disease randomized to health education versus transcendental meditation for 16 weeks. | • At study end, compared to changes in the control group, in the transcendental meditation group there were significant reductions in systolic blood pressure and insulin resistance, and a trend towards improved heart rate variability.  
• There were no significant changes in diastolic blood pressure, lipoprotein levels, C-reactive protein, BMI, or brachial artery reactivity. | • Limitations of this trial included its numerous end points, relatively small size and short duration. |
| Khatri, Diabetes Research and Clinical Practice, 2007 (45) | • Randomized control trial of 101 subjects diagnosed with metabolic syndrome.  
• Usual care versus usual care plus yoga for 12 weeks | • Significant improvement in waist circumference, blood pressure, blood glucose, HbA1c, triglycerides, and HDL-C in Yoga Group. | • Limited description of study population, methods and intervention time and frequency of meditation/ Yoga. |
| Vaccarino, Psychosomatic Medicine, 2013 (46) | • Randomized control in 68 black Americans with metabolic syndrome.  
• Comparing the effect of consciously resting meditation (CRM), a sound (mantra)-based meditation, with a control intervention of health education (HE) on endothelial function in the setting of metabolic syndrome as primary end point and metabolic risk factors, psychosocial and behavioral variables were secondary endpoints | • CRM, did not improve endothelial function significantly more than a control intervention of HE (p=0.51).  
• Improved metabolic syndrome parameters like Diastolic BP, Weight, lipid profile and metabolic risk factors score. | • Study is limited by small number and high attrition rate. |
<table>
<thead>
<tr>
<th>Author(s)</th>
<th>Title and Journal</th>
<th>Study Design</th>
<th>Participants</th>
<th>Intervention</th>
<th>Outcomes</th>
<th>Limitations</th>
</tr>
</thead>
<tbody>
<tr>
<td>Bijlani RL</td>
<td>The Journal of Alternative and Complimentary Medicine, 2005 (47)</td>
<td>Single group with 98 subjects with hypertension, coronary artery disease, diabetes mellitus, and multiple comorbidities.</td>
<td>Improved lipid profile at the end of the study. The changes were more marked in subjects with hyperglycemia or hypercholesterolemia</td>
<td>Subjects were too heterogeneous and study is limited by meagre intervention period of 9 days</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Sivasankaran</td>
<td>Clinical Cardiology, 2006 (48)</td>
<td>Prospective Cohort study with 2 cohort of subjects with and without established CAD.</td>
<td>Significant reductions in blood pressure, heart rate, and BMI in the total cohort with yoga.</td>
<td>Study limited with small group of subjects and 20% of patients failed to complete some portion of the study protocol</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Younge</td>
<td>Psychosomatic Medicine, 2015 (49)</td>
<td>Cross sectional study with 2579 subjects free of Cardiovascular disease from Rotterdam, Netherlands were interviewed for mind body practices (prayer, meditation, yoga, tai chi, qi-gong, breathing exercises, or any other form of mind-body related relaxation techniques).</td>
<td>Fifteen percent of the participants engaged in a form of mind-body practice of which only (n = 97) were meditating.</td>
<td>Observational cross section study where conclusions cannot be drawn for causality.</td>
<td></td>
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</tbody>
</table>
**Supplementary Table 5. Studies of meditation on subclinical atherosclerosis:**

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
</tr>
</thead>
</table>
| Zhang Y et al. Res Sports Med 2013 (50) | • Design: Within group change (no comparator intervention)  
• Population: 20 female hypertensive patients (mean age 57 ±3.5 years)  
• Intervention: Program of traditional Chinese mental and physical exercises performed for 60 minutes twice a week x 24 weeks. One of the mental exercises, “Tu-Na-Yang-Sheng” included breathing and meditation  
• Outcome: ABI (as well as SBP, DBP, and PP)  
• Follow-up: 24 weeks | • Right ABI increased from 1.06 ± 0.08 to 1.12 ±0.08, p=0.041  
• Left ABI increased from 1.06 ± 0.09 to 1.11 ± 0.09, p=0.100  
• Also improvements in SBP, DBP, PP | • A program of physical and mental Chinese exercises (which includes a meditative component) may improve ABI (only statistically significant in right ABI noted but sample is small).  
• Multi-modality, so unable to discern the effects of the meditative components vs. the other stretching and postural components.  
• Small sample  
• Hypertensive women only  
• No comparator group (each participant was their own control)  
• By nature, intervention was noted blinded.  
• Mean ABI was normal to begin with – did not look at a group with PAD. |
Gupta SK, et al. Indian Heart Journal 2011 (51)

- **Design:** Pre-post (within group change)
- **Population:** 123 individuals with angiographically-documented moderate to severe CAD.
- **Intervention:** Rajyoga meditation for stress management, healthy diet (low fat, high fiber vegetarian diet), moderate aerobic exercise. Patients first spent 7 days in-house training, with retraining at 6 month
- **Outcome:** Change in CAD severity by angiography
- **Follow-up:** 2 years for angiographic change in coronary stenosis; 6.48 years for cardiac events

- **Decline in absolute % diameter of coronary stenosis and cardiac events were correlated with percent adherence to intervention.**
  - In patients with highest adherence, percent diameter stenosis regressed by 18.2 ±12.0 absolute percentage points (29% relative improvement, p<0.0001).
  - Least adherence had a progression of 10.6 ± 13.2 absolute percentage points (23% relative worsening, p<0.0001)
  - 91% patients showed a trend towards regression and 51.4% lesions regressed by more than 10 absolute percentage points.
  - Cardiac events were 11 in group with most adherence, and 38 in least adherence. (risk ratio of least vs most; 4.32; 95% CI: 1.69-11.71; p<0.002).

- **A lifestyle invention that included meditation was associated with regression in CAD relative to adherence**
  - By nature, intervention not blinded. Also intervention required a 7 day in-house “sojourn”.
  - Only 76% of patients completed 2 year follow-up angiography
  - No comparison group; patients served as their own control and compared by adherence scores
  - Intervention could not be blinded
  - Multimodality intervention including meditation, diet, exercise – unable to discern the effects of meditation alone
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| Fields JZ et al. Am J Cardiol 2002 (52) | • Design: RCT  
• Population: 57 healthy older adults older than 65 years (mean age 74 years), of which 46 completed post-test ultrasound  
• Intervention (3 arms):  
1. Maharishi Vedic Medicine (MVM) which is Transcendental Meditation practiced 20 min twice daily. This group also got herbal supplements high in antioxidants, were instructed on a Vedic medicine diet (low in fat, high in fruits & vegetables), incorporated Vedic exercises (yoga poses and walking), and attended monthly f/u meetings.  
2. “Modern Medicine” arm that included a conventional diet, exercise, and multivitamin approach.  
3. Usual care (no added therapy)  
• Outcome: cIMT by B-mode U/S  
• Follow-up: 1 yr | • Significant within change in MVM arm (-0.15 +/- 0.21, p=0.004). No significant within change in other treatment groups.  
• Among high risk subjects with multiple CAD risk factors, cIMT decreased more in MVM group (-0.32 ± 0.23 mm) than in the usual care (+0.022 ± 0.085; p=0.009) or modern medicine (-0.082 ± 0.095, p=0.10) groups.  
• cIMT decreased in a larger fraction of MVM subjects (16 out of 20) than in modern (5 out of 9) or usual care groups (7 of 14).  
• Trend for more cIMT decrease among those with better adherence (r= -0.34, p=0.08) | • MVM (which is TM) reduced carotid atherosclerosis among older adults  
• Reductions in cIMT were greater than the Castillo-Richmond study suggesting benefit for multimodality approach with MVM.  
• By nature, intervention not blinded.  
• Small sample size  
• Multi-modality approach. The MVM arm also included an herbal supplement, counseling on healthy diet and incorporated exercise which limits the ability to discern the effect of MVM alone.  
• Older adults only were included. |
| Castillo-Richmond et al. Stroke 2000 (53) | Design: RCT  
Population: 138 hypertensive African Americans enrolled but only 60 completed post-test carotid ultrasound.  
Intervention: Transcendental Meditation vs a Health Education program. TM is a mental technique practiced twice a day for 20 minutes. Initial teaching instructions conducted in both groups within 1 week, follow-up meetings 1 week later, than every 2 weeks for 2 months, and once a month for 3 months.  
Outcome: carotid intimal medial thickness (cIMT) by B-mode U/S.  
Follow-up: 6-9 months (mean 6.8±1.3 months) | TM group showed a significant decrease of -0.098 mm (95% CI -0.198 to 0.003 mm) in cIMT compared with an increase of 0.054 mm (95% CI -0.05 to 0.158 mm) in the control group (P=0.038 for between group difference).  
Correlation between attendance rates of meetings and change in cIMT scores was significant for TM group but not health education group  
TM group also had statistically significant within-group changes in SBP, DBP, pulse and pulse pressure while health education group improved only SBP and DBP.  
TM reduced carotid atherosclerosis among African American hypertensive adults  
High rate of attrition questions the generalizability of these findings. 170 participants were randomized but only 60 had completed post-test interpretable cIMT scans, although attrition was equal in both groups.  
By nature, intervention not blinded.  
Low number of men (~30%)  
Only African Americans |
Levine et al. Meditation and Cardiovascular Risk Reduction: A Scientific Statement From the American Heart Association
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<table>
<thead>
<tr>
<th>Study</th>
<th>Design: RCT</th>
<th>Population: 28 patients with CAD</th>
<th>Intervention: randomized to a lifestyle intervention vs control. Lifestyle intervention included a very low fat vegetarian diet, moderate aerobic exercise, stopping smoking, stress management program (that included stretching, breathing techniques, meditation, progressive relaxation, and imagery) which they were asked to practice at least 1 hour per day, and group therapy</th>
<th>Outcome: Coronary atherosclerosis by quantitative coronary angiography</th>
<th>Follow-up: 1 yr and 5 years</th>
</tr>
</thead>
<tbody>
<tr>
<td>Ornish et al. Lancet 1990 (1 yr f/u) (54) and JAMA 1998 (5 yr f/u) (55)</td>
<td>At 1 yr, 82% overall experienced some regression of their CAD. Average % diameter stenosis regressed from 40% to 38% in intervention group, yet progressed from 43% to 46% in controls. More regression of atherosclerosis occurred at patients in the intervention group at 5 years follow-up than was seen at the 1 yr follow-up. There was a 4.5% and 7.9% relative improvement in coronary stenosis in the intervention arm at 1 and 5 years, respectively vs. 5.4% and 28% relative worsening at 1 and 5 years in control.</td>
<td>A multi-modality lifestyle intervention (that includes meditation stress management as one component) confers coronary atherosclerosis regression</td>
<td>Small sample size. And only 20 out of 28 (71%) had 5 yr follow-up data</td>
<td>Intervention not blinded</td>
<td>Multi-modality of the lifestyle intervention limits the ability to discern the effects of mediation vs. the other components such as the extremely low fat vegetarian diet.</td>
</tr>
</tbody>
</table>
Supplementary Study Summary Table 6. Studies of meditation on endothelial function.

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
</tr>
</thead>
</table>
| Vaccarino V et al. Psychosomatic Medicine 2013 (46) | • Design: RCT  
• Population: 68 year old black Americans with metabolic risk factors  
• Intervention: Randomized to consciously resting meditation (CRM) vs. high education  
• Outcome: Brachial reactivity (FMD)  
• Follow-up: 6 and 12 months | • CRM did improve FMD at 12 months but not statistically significantly more than the health education group despite more favorable trends in metabolic risk factors. Mean change was 2.1% (95% CI 0.5%-3.7%, p=0.009) in CRM group and 1.4% (95% CI = -0.2% to 2.9%, p=0.094) for health education, p-interaction 0.51.  
• Non-endothelium dependent dilation and arterial elasticity also did not change in either group. | • Meditation did not improve endothelial function more than controls  
• Outcome was endothelial function, not atherosclerosis  
• As nature of study, intervention not blinded.  
• Small sample size  
• Only African Americans |
| Paul-Labrador M et al. Arch Intern Med 2006 (44)  | • Design: RCT  
• Population: 103 subjects with stable CAD  
• Intervention: Transcendental Meditation (TM) vs health education  
• Outcome: Brachial Reactivity assessed by flow mediated dilation (FMD)  
• Follow-up: 16 weeks | • No significant effect on brachial reactivity with TM, despite beneficial changes in SBP, insulin resistance, and heart rate variability.  
• There was a non-significant improvement in FMD (-0.11%) in the TM group and a non-significant decline in the health education group (+0.81); (p=0.24 for difference between groups). | • This study failed to show a benefit of TM on endothelial function.  
• Outcome was endothelial function, not atherosclerosis  
• As nature of study, intervention not blinded.  
• High level of statin use and near optimal LDL-C levels in population may have limited TM to confer any additional benefit  
• Relatively small size and short duration  
• All had CAD |

- Design: Within group change, no comparator intervention
- 33 subjects (mean age 55±11 years). 30% had CAD.
- Intervention: Yoga plus meditation combined classes given 90 mins a day three times a day for 6 weeks (each 90 min session was 15 min meditation, 15 min yogic breathing, 20 min of deep relaxation (shavasana), 40 min postural exercises (asanas).
- Outcome: Brachial reactivity (endothelial-dependent vasodilation) and also endothelial-independent vasoreactivity with nitroglycerin
- Follow-up: 6 weeks

- Overall no significant improvement brachial reactivity with yoga and meditation compared with baseline (16.7% relative improvement from a baseline reactivity of 7.2-8.4%; p=0.3).
- In the group with CAD, there was a trend for relative improvement in brachial reactivity with intervention, (69% relative improvement from a baseline of 6.38-10.78%; p = 0.09). No significant change in subgroup without CAD.
- No significant change in endothelial-independent vasodilation overall or in either subgroup.

- Yoga plus meditation may improve endothelial function in individuals with CAD.
- Outcome was endothelial function, not atherosclerosis
- As nature of study, intervention not blinded.
- No comparison group
- Intervention was yoga and meditation combined. Thus unable to discern the effect of meditation alone.
- Small study, short duration
- 20% failed to complete some portion of study protocol
### Supplementary Study Summary Table 7. Effects of meditation on inducible myocardial ischemia

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
</tr>
</thead>
</table>
| Cunningham et al. AJC 2000 (56) | • Longitudinal pre/post TM intervention study in 9 postmenopausal women with cardiac syndrome X  
  • Subjects underwent exercise treadmill testing (Bruce Protocol) pre and post 3 months of TM                                                                                                        | • Compared to baseline, TM improved time to 1 mm ST segment depression \( p = 0.03 \), maximum ST segment depression \( p=0.03 \), frequency of angina episodes \( p=0.04 \), and quality of life \( p=0.003 \).  
  • No significant differences in heart rate, blood pressure, or duration of exercise before and after TM.                                                                                      | • Non-randomized  
  • Patients served as their own control group  
  • TM compliance was high                                                                                                          |
| Zamarra et al. AJC 1996 (57) | • Single blinded study in 21 subjects that compared 7.6 months of TM in 12 CAD patients vs. 9 CAD patients who were waitlisted for TM and served as controls  
  • Subjects underwent symptom-limited exercise tolerance testing (upright cycle protocol) before and after intervention                                                                 | • TM led to greater exercise tolerance - 14.7% increase in exercise duration \( p=0.013 \), 11.7% increase in maximal work load \( p=0.004 \), and 18.1% delay of onset of ST depression \( p=0.029 \). | • Only 16 completed the study due to various reasons (10 in TM and 6 waiting controls)  
  • TM compliance was high                                                                                                           |
Ornish et al. JAMA 1983 (58)

- Randomized study comparing the effects of short-term (24 days) stress management training plus dietary changes in 23 subjects with IHD vs 23 controls.
- Subjects underwent exercise nuclear ventriculography imaging with wall motion and ejection fraction (EF)
- Stress management training and dietary changes resulted in a 44% increase in exercise duration (p<0.001) and 55% increase in total work performed (p<0.001).
- Improved left ventricular wall motion during peak exercise and a net change in EF from rest to maximum exercise of +6.4%.
- Lifestyle intervention lowered total cholesterol (p<0.001) and triglyceride levels (p<0.01); deceased HDL levels (p<0.0001), but the ratio of total cholesterol/HDL showed no difference between the two groups.
- Since both dietary changes (mainly vegan-based diet and 1400 kcal/day) and stress management training were included, the relative contribution of stress reduction cannot be assessed.
- Changes in EF and wall motion assessed but inducible myocardial ischemia or changes in coronary blood flow not measured.
Supplementary Study Summary Table 8. Meditation and primary prevention of cardiovascular disease.

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
</tr>
</thead>
</table>
| Barnes VA, Journal of Social Behavior and Personality, 2005 (59) | • RCT of African Americans ≥55 years old with mild hypertension (n=109) assigned to (a) transcendental meditation (TM); (b) progressive muscle relaxation (PMR) or (c) a health education control (EC) program  
  • Treatment conducted twice daily for 20 minutes, over a 3-month period, after which patients were encouraged to continue with their treatment program on their own long-term.  
  • Follow-up of mortality events conducted an average of eight years after randomization in the original phase of this trial | • Relative risk for all-cause mortality in the TM group compared with PMR was 0.43 (95% CI 0–1.16, p < .08), and for the TM group compared with EC was 0.51 (95% CI 0–1.33, p < .12).  
  • Relative risk of cardiovascular deaths for TM compared to PMR was 0.33 (95% CI 0–2.27, p < .16), and for TM compared to EC was 0.25 (95% CI 0–1.60, p < .08).  
  | Study limited by small sample size  
  Mortality not the original primary endpoint  
  Follow-up of mortality events conducted an average of eight years after randomization in the original phase of this trial  
  Compliance to the intervention was not monitored after the three-month follow-up was completed. |
| Schneider RH, Am J Cardiol, 2005 (60) | • Data pooled from 2 RCT (n=202) (see Alexander and Barnes, above) that compared TM, other behavioral interventions, and usual therapy for hypertension  
  • Programs practiced for 20 minutes twice daily for 3 months  
  • All-cause mortality primary endpoint; cardiovascular and cancer mortality secondary endpoints.  | • Mean follow-up was 7.6 ± 3.5 years.  
  • Compared with controls, TM showed a 23% decrease in all-cause mortality (relative risk 0.77, p = 0.039).  
  Secondary analyses showed a 30% decrease in cardiovascular mortality (relative risk 0.70, p = 0.045) and a 49% decrease in the rate of cancer mortality (relative risk 0.49, p = 0.16) in the TM group compared with controls  
  | Retrospective study limited by modest sample size  
  Only mortality data from national databases were collected  
  Follow-up of mortality events conducted three to eight years after randomization  
  Compliance to the intervention was not monitored after the three-month follow-up was completed. |
Alexander CN, J Pers Soc Psychol 1989 (61)

- RCT of institutionalized elderly (n=73) assigned to (a) no treatment (n=11); (b) transcendental meditation (TM) (n=20); (c) mindfulness training (MF) in active distinction making (n=21), or (d) mental relaxation (MR) with low mindfulness (n=21).
- Programs practiced for 20 minutes twice daily for 3 months
- Assessed cognitive function and health (i.e., blood pressure, mental health, general health and longevity) endpoints

- Mindfulness techniques associated with improvements in blood pressure, cognitive functioning and mental health.
- Mindfulness techniques improved 36 month survival rates; mortality 100% for TM; 87.5% for MF; 77.3% for no treatment; and 65% for relaxation (p<0.00025).

- Study limited by small sample size
- Highly biased study (i.e., data were unavailable for 3 “no treatment subjects,” so 14 nonrandomized subjects were included in the mortality analysis of these subjects).
- Follow-up of mortality events conducted three years after randomization
- Compliance to the intervention was not monitored after the three-month follow-up
- Longevity assessed from nursing home records
**Supplementary Study Summary Table 9. Studies of meditation in patients with established cardiovascular disease.**

<table>
<thead>
<tr>
<th>Reference</th>
<th>Study type, design, type of meditation, and population</th>
<th>Primary Findings</th>
<th>Comments</th>
</tr>
</thead>
</table>
| DuBroff R, Alternative Therapies, 2015 (62) | • Longitudinal pilot study of 22 patients with documented CAD treated with ayurvedic therapy (dietary recommendations and restrictions, meditation, breathing exercises, yoga and herbs)  
  • So’Ham meditation practiced initially 10 minutes twice daily, with instructions for patients to increase durations weekly | • At 90 days, there was significant improvement in arterial pulse wave velocity and significant reductions in BMI, total cholesterol, LDL cholesterol and triglycerides | • Study limited by small sample size (19 patients completed the study) and no control group  
  • Exact contribution of meditation to study findings cannot be discerned |
| Younge JO, Eur J Preventive Cardiology, 2015 (63) | • Systematic review and meta-analysis of randomized controlled trials of mind-body practices for patients with cardiac disease  
  • 11 studies included various types of meditation, mindfulness based stress reduction, stress management, and relaxation  
  • Studies variably included patients with coronary artery disease or heart failure | • Pooled analyses revealed effect sizes of 0.45 (95%CI 0.20–0.72) for physical quality of life, 0.68 (95%CI 0.10–1.26) for mental quality of life, 0.61 (95%CI 0.23–0.99) for depression, 0.52 (95%CI 0.26–0.78) for anxiety, 0.48 (95%CI 0.27–0.69) for systolic blood pressure and 0.36 (95%CI 0.15–0.57) for diastolic blood pressure. | • Study authors concluded “promising but heterogeneous results were seen on overall effect sizes of mental and physical quality of life, anxiety, depression, and blood pressure”  
  • Study authors rated overall quality of the studies as low and that no firm conclusions could be drawn |
<table>
<thead>
<tr>
<th>Study</th>
<th>Summary</th>
<th>Findings</th>
<th>Limitations</th>
</tr>
</thead>
<tbody>
<tr>
<td>Parswani MJ, <em>International Journal of Yoga</em>, 2013 (64)</td>
<td>Randomized study of 30 male patients with a diagnosis of coronary artery disease treated with either mindfulness based stress reduction (mindfulness meditation) consisting of 8 weekly instructional session and 30 minutes meditation daily at home or “treatment as usual”</td>
<td>At the end of the intervention, for the mindfulness based stress reduction group, there were significant within group and between group decreases in anxiety, depression, perceived stress, and systolic blood pressure At 3 month follow-up, there was a further significant reduction in blood pressure</td>
<td>Study limited by small sample size (15 patients per group) and relatively short-term follow-up Intervention consisted not only of meditation but also exercise and diet, though only significant changes seen in the group additionally treated with meditation</td>
</tr>
<tr>
<td>Nehra, <em>Dysphrenia</em> (now called <em>Open Journal of Psychiatry &amp; Allied Sciences</em>), 2013 (65)</td>
<td>Randomized study of 50 patients with coronary artery disease (MI or angina) randomized to 8 week program of mindfulness based stress reduction (mindful meditation) or usual care control group Mindfulness based stress reduction program included 2.5 hour weekly meetings, six-hour daylong retreat, and home practice at least 45 minutes daily, and included sitting meditation, hatha yoga, and the body scan</td>
<td>At 10-17 week post-assessment follow-up, there were, compared to changes in the control group, significant decreases in perceived stress, cognitive health complaints, and somatic health complaints in the mindfulness based stress reduction program group</td>
<td>Study limited by modest sample size (25 patients per group)</td>
</tr>
<tr>
<td>Delui MH, <em>The Open Cardiovascular Medicine Journal</em>, 2013 (66)</td>
<td>Randomized trial of 45 patients with cardiovascular disease and depression referred for cardiac rehabilitation randomized to relaxation, mindful meditation, or control Mindful meditation instruction included ten 20-25 minute sessions and home practice</td>
<td>At the end of the study (duration unclear) repeat testing showed significant reductions in depression, systolic blood pressure and heart rate in the meditation group compared to the control group</td>
<td>Study limited by small sample size (15 patients per group) Diastolic blood pressure and anxiety score not significantly reduced by any intervention</td>
</tr>
</tbody>
</table>
Schneider RH, Circ Cardiovasc Qual Outcomes, 2012 (67)  
- Randomized study of 201 black men and women with angiographic evidence of at least 1 coronary artery with >50% stenosis treated with either transcendental meditation (20 minutes twice daily) or health education  
- After a mean of 5.4 years, primary composite endpoint (all-cause mortality, nonfatal MI, nonfatal stroke) significantly lower in the transcendental meditation group (adjusted HR=0.52; 95% CI=0.29-0.92; p=0.025)  
- Secondary composite endpoint (CV mortality, nonfatal MI, nonfatal stroke, coronary revascularization, hospitalization for IHD or heart failure) non-significantly reduced (adjusted HR-0.76; 95% CI=0.51-1.13; p=0.17)  
- Study conducted in two phases after a hiatus in funding with 58 of the 201 subjects not participating in phase 2  
- Study completed in 2007 and published in 2012  
- Significant net difference of -4.9 mmHg in SBP in TM group (95% CI=-8.3 to -1.5 mm Hg; p=0.01)

Gupta SK, Indian Heart J, 2011 (51)  
- Longitudinal study of 123 patients with angiographically documented stable CAD (67% with history of MI) treated with comprehensive lifestyle modification, including Rajyoga meditation  
- Intervention included stress management through Rajyoga meditation, vegetarian diet, and moderate aerobic exercise  
- 2 year angiographic follow-up performed on 76% of participants. Average percent diameter stenosis decreased by 6.10 absolute percentage points (p<0.003)  
- Specific independent contribution of meditation to study findings cannot be determined

Paul-Labrador, Arch Intern Med, 2006 (44)  
- Randomized trial of 103 patients with documented and stable coronary artery disease randomized to 16 weeks of transcendental meditation or active control (health education)  
- Transcendental meditation intervention included personalized and group instruction and maintenance meetings  
- At study end, compared to changes in the control group, in the transcendental meditation group there were significant reductions in systolic blood pressure and insulin resistance, and a trend towards improved heart rate variability  
- There were no significant changes in diastolic blood pressure, lipoprotein levels, C-reactive protein, BMI, or brachial artery reactivity  
- Study findings somewhat limited by numerous study endpoints
Tacon AM, Fam Community Health, 2003 (68) and Robert-McComb JJ, Journal of Alternative and Complementary Medicine, 2004 (34)

- Randomized study of 18 patients with cardiovascular disease randomized to either mindfulness based stress reduction instruction (2 hours each week for 8 weeks) and home practice or a waiting list for such training
- Intervention included training in body scan, sitting meditation, and hatha yoga
- At study end, in the intervention group there were significant improvements in measured anxiety, emotional control, and coping styles
- There was no significant change in “health locus of control”
- Study limited by small sample size (9 patients per group) and relatively short term follow-up
- Participants included a mixed population of those with angina, hypertension, “cardiovascular disease”, and cardiac valve disorders
- A second publication of seemingly the same study population reported no significant changes in stress hormones or submaximal exercise responses

Sullivan, Am Heart J, 2009 (69)

- Prospective cohort study of 208 patients with heart failure (46% due to ischemic etiology) geographically assigned to a mindfulness-based intervention (8 weekly meetings plus practice of skills at least 30 minutes each day) or standard care
- Mindfulness-based intervention included mindfulness based stress reduction plus education on improving coping skills and an expressive support group discussion
- At 12 month follow-up, intervention resulted in significantly lower anxiety, depression, and heart failure symptoms and clinical scores
- No treatment effect on rehospitalization or death at 1 year
- Study limitation is use of a geographic control
- Specific contribution of mindful meditation itself to study findings cannot be determined
Levine et al. Meditation and Cardiovascular Risk Reduction: A Scientific Statement From the American Heart Association

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<table>
<thead>
<tr>
<th>Study</th>
<th>Description</th>
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<tbody>
<tr>
<td>Zamarra JW, Am J Cardiol, 1996 (57)</td>
<td>• 21 male patients with documented CAD (≥70 lesion on angiography and/or prior MI) with inducible myocardial ischemia on upright cycle ETT were assigned to either transcendental meditation or wait-list control group. &lt;br&gt;• Transcendental meditation intervention included 10 hours of basic instruction, follow up meetings, and home practice 20 minutes twice daily</td>
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<td>• At a mean 7.6 month follow-up, repeat ETT demonstrated that compared to the control group, the meditation group had significant increases in exercise duration, maximum workload, and time to ST depression onset</td>
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<td>• Study limited by small sample size (only 10 in intervention group and 6 in control group completed study)</td>
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Reference List


