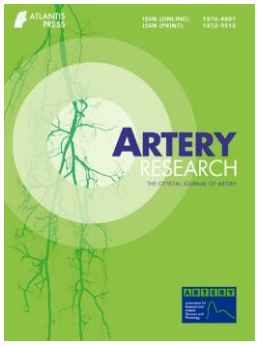

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4.4: ASSOCIATIONS BETWEEN OBJECTIVELY MEASURED PHYSICAL ACTIVITY ENERGY EXPENDITURE AND CENTRAL HAEMODYNAMICS. THE ADDITION-PRO STUDY

A.S.D. Laursen, A.-L.S. Hansen, N. Wiinberg, S. Brage, A. Sandbæk, T. Lauritsen, M.E. Jørgensen, B. Kiens, N.B. Johansen

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Background. No longitudinal study addressed whether systolic blood pressure level (SBPL) or variability (SBPV) predict arterial properties or vice versa.
Methods and Results. In families randomly recruited from a Flemish population, we determined SBPL and SBPV from 5 consecutive blood pressure readings. The indexes of SBPv were variability independent of mean (VIM), the difference between maximum and minimum SBPL (MMD), and average real variability (ARV). We measured carotid intima-media thickness (cIMT) and distensibility (cD) by ultrasound and carotid-femoral pulse wave velocity (cfPWV) by tonometry (SphygmoCor, version 8.2). Effect sizes were computed for 1 SD increments in the predictors, while accounting for covariables and family clusters. Among 1171 participants (51.0% women; mean age, 39.8 years), followed up for 2.48 years (median), higher SBPL predicted ($P \leq 0.036$) higher cIMT (+14 μm), lower cD (−1.57 10-3/kPa) and faster cfPWV (+0.298 m/s) at follow-up, whereas none of the SBPV indexes predicted the arterial traits at follow-up ($P \geq 0.11$). In a subset of 749 participants, followed up for another 3.07 years, lower cD predicted ($P \leq 0.026$) higher SBPL (+1.69 mm Hg), VIM (+0.304 units), MMD (+1.05 mm Hg) and ARV (+0.389). Higher cfPWV predicted a 1.06 mm Hg increase SBPL ($P = 0.027$).
Conclusions. Temporality and effect size suggest that SBPL but not SBPV cause arterial stiffening and cIMT thickening. Carotid stiffening, independent of SBPL, predicts SBPV, possibly because baroreflexes originating from a stiffer carotid artery wall are impaired. Finally, stiffening of the aorta contributes to the age-related SBPL possibly because faster returning reflected waves augments SBPL.

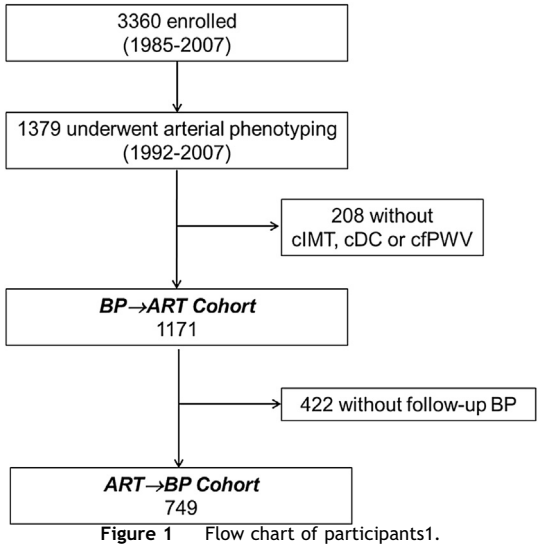

Figure 1 Flow chart of participants1.

4.3
CAROTID HEMODYNAMICS DURING SYMPATHETIC NERVOUS SYSTEM STIMULATION VIA HANDGRIP AND COLD PRESSURE TESTING IN YOUNG HEALTHY SUBJECTS

F. J. Londono [1], T. S. Klopmeijer [2], D. Georgakopoulos [3], E. G. Lovett [3], P. Segers [1]
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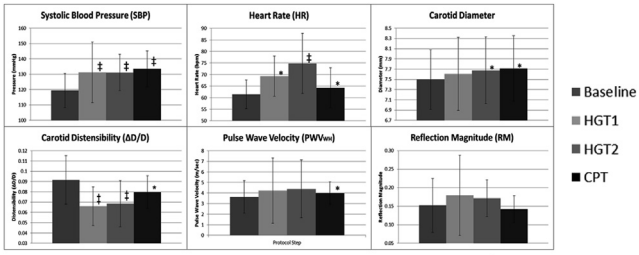
Introduction. Assessing arterial properties and hemodynamics in response to a physiological perturbation might provide additional information on an individual's "vascular health". The aim of this study was to assess the feasibility of measuring changes in carotid stiffness and hemodynamics in response to sympathetic nervous system stressors (Hand Grip Test (HGT) or Cold Pressor Test (CPT)).
Methods. A non-invasive protocol, consisting of two HGT (30% (HGT1) and 40% (HGT2) of maximal voluntary contraction) and CPT, was performed in 12 young healthy volunteers (6 males/6 females). Measurements included continuous finger blood pressure recordings (NexFin; non-dominant hand) and ultrasound measurement of common carotid diameter distension and flow velocity at discrete moments in time during the protocol (GE Vivid 7 system). Carotid distensibility (ÄD/D), local wave speed using the waterhammer equation (PWV_{WH}) and reflection magnitude (RM; ratio of backward to forward diameter wave) were derived from the data.
Results. Consistent with the overall increase in blood pressure, carotid diameter increased while ÄD/D decreased. PWV_{WH} and RM showed an increase during both HGT and a decrease during CPT (see Figure).
Conclusion. It is feasible to monitor the carotid hemodynamic response to a sympathetic nervous system stimulus. In this young, healthy population, the net result of the increased diameter and decreased distensibility on local PWV_{WH} was similar for all tests (increase) and on the magnitude of reflections was different for HGT (increase) and CPT (decrease).



The difference from baseline is indicated as significant * ($p<0.05$) or highly significant ‡ ($p<0.001$).

4.4
ASSOCIATIONS BETWEEN OBJECTIVELY MEASURED PHYSICAL ACTIVITY ENERGY EXPENDITURE AND CENTRAL HAEMODYNAMICS. THE ADDITION-PRO STUDY

A. S. D. Laursen [1,2], A.-L. S. Hansen [1,3], N. Wiinberg [4], S. Brage [5], A. Sandbæk [6], T. Lauritsen [6], M. E. Jørgensen [1], B. Kiens [2], N. B. Johansen [1,7]
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Objective. Physical activity (PA) has been associated with reduced cardiovascular disease (CVD). However, improvements in conventional risk factors do not explain the full benefit of PA on CVD risk. Therefore, we examined the association between PA and central haemodynamics to provide new insight into the link between PA and CVD.
Methods. We performed cross-sectional analyses of data from a health examination of 1,607 Danish adults at low to high diabetes risk. PA energy expenditure (PAEE) was measured by combined accelerometry and heart rate monitoring (ActiHeart®) expressed as kJ/kg/day. Aortic stiffness was assessed by applanation tonometry (SphygmoCor®), as aortic pulse wave velocity (aPWV), and central blood pressure was estimated from wave forms recorded at the radial artery. Associations between PAEE and central haemodynamics were examined by linear regression successively adjusted for sex, age, waist circumference, and smoking. Additionally, aPWV was adjusted for mean blood pressure. aPWV was logarithmically transformed. Individuals with previously myocardial or cerebral infarction were excluded ($n = 183$).
Results. Median age was 66 years (IQR: 61;71), 52% was men, median PAEE was 28 kJ/kg/day (IQR: 20;39), and median aPWV 8.0 m/s (IQR: 6.9;9.4). A higher PAEE of 10 kJ/kg/day was associated with a 1.0 % lower aPWV (CI: -0.17;-0.03). Associations with systolic blood pressure and pulse pressure were not statistically significant (-0.5 mmHg (CI: -0.11;0.01) and -0.02 mmHg (CI: -0.06;0.03)).

Conclusion. A higher level of PAEE is associated with lower levels of aortic stiffness indicating that the beneficial effects of PA on CVD are partially mediated by aortic stiffness.

4.5
FREQUENCY RESPONSE OF BLOOD PRESSURE CUFFS BASED ON STEP RESPONSE AND FORCED SINUSOIDAL HARMONIC EXCITATION

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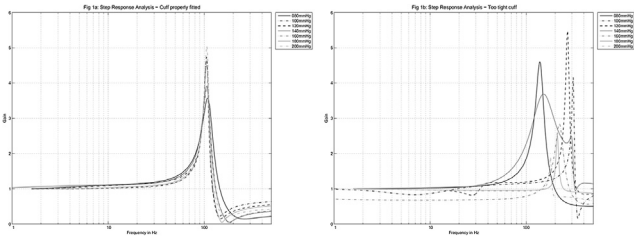
Background. Pulse wave analysis (PWA) using cuff based methods emerged within the last years. Compared to traditional sensors used in PWA like catheters and piezo gauges very little is known on the frequency response of cuff based systems according to standard measurement and systems theory. Therefore the aim of this work is the investigation of the capability of blood pressure cuffs to register the dynamical behavior of the arterial pulse appropriately.
Methods. To evaluate the dynamic behavior of cuff based sensor chains we performed both tests on step response (similar to "pop test") as well as forced sinusoidal harmonic excitation by the means of a fully automated and standardized custom testing bench. The stepwise variation of cuff pressures and/or bladder volumes and cuff sizes was intended to account for various anatomical situations faced in clinical practice.
Results. The resonant frequencies of the evaluated systems are in the range of 110 Hz (Fig. 1a). Similar results have been obtained for both step response method and harmonic excitation. This behavior did not vary over a pressure range from 80 to 200 mmHg. Nevertheless we observed that pressure-volume relation had significant influence on the cuffs frequency response. In particular too tightly fitted cuffs lead to chaotic results (Fig. 1b). Too loosely fitted cuffs causes a loss in resolution.
Conclusions. Based on our actual data blood pressure cuffs provide acceptable capabilities to cope with the dynamics of the arterial pulse for PWA use. This is only valid for properly fitted and inflated cuffs.



Oral Session 5
Invited Lecture and Free Communication Oral Presentations
In association with the Pulse of Asia

5.1
VULNERABLE CAROTID PLAQUES ARE ASSOCIATED WITH THE DEVELOPMENT OF EARLY RESTENOSIS AFTER CAROTID ENDARTERECTOMY

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Samara State Medical University, Samara, Russian Federation

Background. Carotid endarterectomy (CEA) is widely recognised as an effective surgical method in both symptomatic and asymptomatic patients with significant internal carotid artery stenosis. Although the association between vulnerable carotid plaques and stroke is well established, little is known about their role in development of myointimal hyperplasia after carotid surgery. In the current study we aimed to establish correlation between plaque morphology (as observed by ultrasonic scan) and the degree of myointimal hyperplasia after CEA.
Methods. A total of 567 patients with a median age of 65.0 $\pm$ 5,6 years who underwent CEA were examined using duplex ultrasound scanning prior to surgery and 12 months after. The morphology of plaques in terms of their echogenicity was graded as echolucent, predominantly echolucent, predominantly echogenic, echogenic, or calcified. The plaque surface was categorized as smooth, irregular, or ulcerated. Chi-square test and multivariate logistic regression were used for statistical analysis.
Results. Internal carotid artery restenosis due to intimal hyperplasia $\geq$50% were detected in 67 patients (11,82%). The incidence of carotid restenosis was significantly higher in patients with ulcerated carotid plaques ($P < 0.05$). There was no difference between rates of restenosis in patients who had plaques with smooth or irregular surface. Predominantly echolucent carotid plaque appeared to be an independent predictor of carotid restenosis in 12 months after CEA ($P < 0.05$).
Conclusion. The results of our study suggest that vulnerable carotid plaques can lead to myointimal hyperplasia after CEA and can be considered as an independent predictor of early restenosis after carotid surgery.

5.2
CENTRAL PULSE PRESSURE: A POSSIBLE ROBUST MARKER OF THE CARDIAC HEMODYNAMIC LOAD

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Tokyo Medical University, Tokyo, Japan

Background. No prospective study has examined whether markers of the central hemodynamics, e.g., central pulse pressure, might be a more robust marker of change in the cardiac hemodynamic load, estimated by the change during the study period of the serum levels of N-terminal fragment B-type natriuretic peptide (NT-proBNP), as compared to markers of the arterial stiffness or brachial blood pressure variables in subjects with preserved cardiac function.
Methods. The brachial-ankle pulse wave velocity, radial augmentation index (rAI), second peak of the radial pressure waveform, systolic and pulse pressure of the second peak of the radial pressure waveform (SP2 and PP2) and serum NT-proBNP level were measured at the start (first examination) and at the end (second examination) of this 3-year study in middle-aged healthy Japanese men ($n = 1851$).
Results. A stepwise multivariate linear regression analysis demonstrated that only PP2, among the parameters related to arterial stiffness and central hemodynamics and also brachial blood pressure variables, was significantly associated with the serum NT-pro BNP levels in the subjects. Furthermore, only the changes of the PP2 during the study period, among the parameters related to arterial stiffness and central hemodynamics, were significantly correlated with those of the serum NT-pro BNP levels during the study period (beta $= 0.131$, $p < 0.001$).
Conclusion. Central pulse pressure, as reflected by PP2, may be a robust marker of the cardiac hemodynamic load and reflect changes in the cardiac hemodynamic load even in persons with preserved cardiac function.
Keywords. Natriuretic peptides, Central blood pressure, Cardiac hemodynamic load

5.3
INORGANIC NITRITE, CONDUIT ARTERIES & CENTRAL BLOOD PRESSURE

S. Omar [1,2], H. Fok [1,2], A. Nair [1,2], J. Hunt [1,2], B. Jiang [1,2], P. Chowienczyk [1,2], A. J. Webb [1,2]
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[2]*Guy's & St. Thomas' NHS Foundation Trust, London, United Kingdom*

Background. Organic nitrates (e.g. nitroglycerin) are highly selective dilators of muscular conduit arteries. By contrast, the endogenous inorganic nitrite anion (NO_2^-) is thought to be a hypoxia-dependent dilator of small resistance arterioles, via its reduction to vasodilating nitric oxide (NO) by deoxyhaemoglobin.
Objective. To establish selectivity of nitrite for resistance versus conduit arteries.
Methods and Results A series of forearm blood flow (FABF) studies were performed in healthy volunteers. Intra-brachial sodium nitrite (8.7 μmol/min) markedly increased radial artery diameter (assessed using ultrasound) by 37.6 $\pm$ 9.7% ($P < 0.001$), with HbO_2 ~99%. Furthermore, nitrite (0.087-87 μmol/min) displayed similar selectivity as nitroglycerin (0.003-1 μg/min) for conduit arteries, compared to resistance arterioles (FABF). Intravenous administration of sodium nitrite (8.7 μmol/min) dilated the contralateral radial artery by 10.7 $\pm$ 1.8% ($P < 0.01$) and lowered central systolic blood pressure (BP) by ~12mmHg from 98.3 $\pm$ 12.3 to 86.7 $\pm$ 15.1 mmHg ($P = 0.02$) without any change in peripheral BP; nitrite also reduced augmentation index and pulse wave velocity. In contrast to nitrite's effects on FABF, induction of hypoxia (breathing 12% O_2) paradoxically inhibited nitrite-induced dilatation of the radial artery to a similar extent as hyperoxia/breathing 100% O_2 (both $P < 0.001$ compared to normoxia).