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P9.09: DIFFERENCE IN AGE-RELATED PATTERNS OF ARTERIAL STIFFNESS AND WAVE REFLECTIONS AMONG PATIENTS WITH KIDNEY DISEASE: RESULTS OF THE UK RESEARCH ALLIANCE INTO KIDNEY DISEASE AND ARTERIAL STIFFNESS (UREKA) COLLABORATION

L.A. Tomlinson, B. Caplin, S.J. Carr, C. Delles, C.J. Ferro, J. Goddard, S.G. Holt, P.A. Kalra, P.S. Lacy, C.M. McEniery, C.W. McIntyre, E.P. McQuarrie, C. Rajkumar, T. Savage, M.W. Taal, C.R. Tompson, J.N. Townend, D.J. Webb, D.C. Wheeler, I.B. Wilkinson

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(129.3±74.0) and without prevalent CVD(120.2±68.6). Participants with a high waist (4th quartile, gender specific) had higher IL18 levels than those with a low waist(1th quartile)(125.9±77.9 versus 117.2±61.0). The increase in IL18 was accompanied by an increase in subclinical atherosclerosis, as reflected by a lower ABI (1.09 versus 1.11), a thicker IMT(0.86 versus 0.83 mm) and increased arterial stiffness, as reflected by an increased PWV(10.3 versus 9.6 m/s) and an increase in all derived central pressure parameters. **Conclusion:** In our population-based cohort obesity, as reflected by an increased waist circumference, was accompanied by increased IL-18 levels and an increase in non-invasively determined subclinical atherosclerosis. Our data support the hypothesis that the increased CVD risk in obesity might be caused by increased inflammation, although prospective studies are needed to conclude on causality of this relation.

P9.06

SERUM URIC ACID LEVELS AND ARTERIAL STIFFNESS AND CARDIAC AND CAROTID ARTERY STRUCTURE IN A GENERAL POPULATION IN NORTHERN ITALY

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Background: The relationship between serum uric acid (UA) levels and CV disease has been described since the late 19th century. The role of UA as an independent risk factor for CV events and its association with TOD is however less clear. Aim of the study was to assess the relationship between UA and TOD in a general population (Vobarno Study).

Methods: 385 subjects (age 56±9yrs, 44%males, 64% hypertensives, 32% treated) underwent laboratory examinations and clinic and 24 hours BP measurement. Left ventricular and carotid artery structure were assessed by ultrasound and carotid-femoral PWV was measured using Complior.

Results: Subjects with increased UA (>6 mg/dl in ♀ and >7 mg/dl in ♂) were older, had greater BMI, higher BP, glucose, cholesterol and triglycerides levels and lower HDL cholesterol and e-GFR. Subjects with increased UA had also increased PWV (11.1±4.1 vs 13.3±3.7m/sec, p<0.0001), and a slight increase in left ventricular mass index (LVMI) (38.7±10.6 vs 43.0±11gr/m^{2.7}, p<0.05) and IMT (Meanmax 1.1±0.28 vs 1.2±0.29 mm, p<0.05). After adjusting for confounders, including e-GFR, in a multivariable model, PWV was significantly greater in subjects with increased UA (11.1±2.41 vs 13.4±3.7 m/sec, p<0.001), while no significant difference in LVMI and IMT was observed. A significant correlation between UA levels and PWV(r=0.279, p<0.001), LVMI(r=0.157, p<0.001), meanmax IMT(r=0.159, p<0.001) was observed. After adjusting for confounders, serum UA levels were independently correlated to PWV, but not to LVMI and IMT.

Conclusions: Subjects with increased UA have increased arterial stiffness, but comparable left ventricular anatomy and carotid artery structure. The increase in arterial stiffness might contribute to the higher CV risk in these subjects.

P9.07

RELATION OF CENTRAL AND BRACHIAL BLOOD PRESSURE TO LEFT VENTRICULAR HYPERTROPHY. THE CZECH POST-MONICA STUDY

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Objective: Recently central aortic blood pressures were shown to be better predictors of target organ damage, cardiovascular events and mortality when compared with conventional brachial blood pressure. Whether central blood pressure is a better predictor of left ventricular hypertrophy (LVH) determined by electrocardiographic criteria is not known.

Methods: Radial applanation tonometry and ECG were performed in 563 individuals from the Czech post-MONICA study (a randomly selected 1% representative population sample, mean age 46±11 years, 44% of men). LVH was determined using electrocardiographic criteria. Brachial blood pressure was measured using mercury sphygmomanometer according to standardized protocol; central systolic blood pressure was derived from radial pulse wave using generalised transfer function.

Results: Of 563 subjects 39 (7%) had ECG signs of LVH. In the univariate analysis patients with LVH were older (50.4±11.3 vs. 46.6±11.3, p=0.04), had higher central systolic (129.7±31.8 vs. 116.7±16.8, p<0.0001), diastolic (83.3±10.3 vs. 79.2±9.7, p=0.04), pulse (46.5±13.1 vs. 37.5±12, p<0.0001) and mean pressure (103.5±14.8 vs. 96±11.8, p<0.01), higher brachial systolic (136.6±19.4 vs. 122.8±14.8, p<0.0001), pulse (67.2±19.3 vs. 52.6±16.8, p<0.0001) and mean (91.8±10.3 vs. 87.2±8.9, p=0.02) pressure and aortic pulse wave velocity (8.3±2.1 vs. 7.5±1.8, p=0.02) than subjects without LVH. However, in the binary logistic regression only central systolic pressure (OR 2.2, 95% CI 1.4-3.4, p=0.001) and male sex (OR 4.8, 95% CI 1.3-17.6, p=0.02) were predictors of LVH.

Conclusion: Noninvasively determined central systolic blood pressure is more strongly related to LVH determined by electrocardiographic criteria than brachial systolic and pulse pressure. This is a further support of closer association of central blood pressure with target organ damage. Prospective studies with central blood pressure as a treatment target will be needed in the future.

P9.08

THE ASSOCIATION BETWEEN METABOLIC SYNDROME AND AORTIC STIFFNESS IN GENERAL POPULATION

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Background: Despite being a cluster of conventional risk factors, metabolic syndrome (MetSy) has been recognized as independent predictor of cardiovascular diseases. We aimed to establish the association between metabolic syndrome and aortic pulse wave velocity (aPWV) as a measure of arterial stiffness in Czech general population.

Methods: 576 subjects (mean age 48.03y (SD 14.8)), 41.5% males), a Pilsen sub-sample of postMONICA survey, were included into cross-sectional study. aPWV was estimated using Sphygmocor device, subjects with MetSy were identified using common NCEP-ATPIII definition.

Results: Subjects with MetSy showed significantly higher aPWV (9.02 vs. 7.42 m/sec, p<0.001), also if diabetic (8.75 vs. 7.18, p<0.001) or diabetic and hypertensive patients (7.96 vs. 6.84, p<0.001) were excluded from the analysis (p value adjusted for age).

The significance of association between MetSy and aPWV remained significant after adjustment for age, gender, current smoking, mean arterial pressure, serum glucose and other risk factors as potential confounders (b=-0.088, p=0.023).

Conclusion: In our sample of general population, we found that MetSy represents an additive risk factor of increased aortic stiffness independent of age, blood pressure, glucose status and other conventional factors.

P9.09

DIFFERENCE IN AGE-RELATED PATTERNS OF ARTERIAL STIFFNESS AND WAVE REFLECTIONS AMONG PATIENTS WITH KIDNEY DISEASE: RESULTS OF THE UK RESEARCH ALLIANCE INTO KIDNEY DISEASE AND ARTERIAL STIFFNESS (UREKA) COLLABORATION

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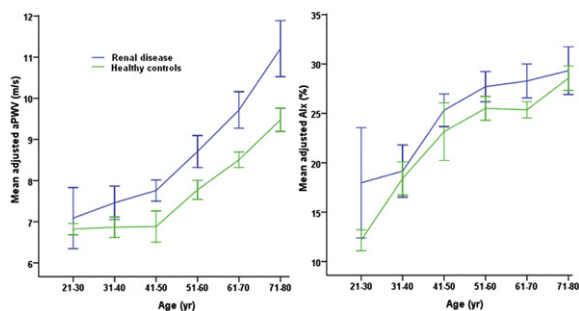
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Background: Patients with CKD may have higher aortic PWV (aPWV) and augmentation index (Alx) but studies are limited by size and lack of control population and differences may be due to joint risk factors. We examined whether aPWV and Alx are increased in CKD patients with no vascular comorbidities compared to controls and how change with age varies between populations.

	Kidney disease	Healthy controls
Age (y)	55±14	56±18
Male (%) [*]	66	48
eGFR [*]	43±21	96±30
Syst./Diast	133/80	134/80
BP	±20/11	±19/11
Adj PWV [*]	9.0±2.5	7.8±1.8
Adj Alx [*]	26.0±9.2	19.3±8.5

Methods: Cardiovascular risk factor data, aPWV and Alx were obtained from CKD patients at 8 UK renal centres and participants in ACCT, a study of the general population. Those with diabetes and/or vascular disease were excluded. The relationship of age to aPWV and Alx was compared between patients with CKD (stages 1-5, not on dialysis, non-vascular diagnosis, n=524) and controls (eGFR >60ml/min, n=1535). Controls were stratified by age and gender to ensure comparability.



Results: Adjusted aPWV and Alx were higher in CKD patients ($P<0.001$). There was a significant interaction ($P<0.001$) between age and the presence of CKD on aPWV, but this was not seen for Alx ($P=0.19$).

Conclusions: Kidney disease in the absence of co-morbidities is associated with increased arterial stiffness compared to controls. In CKD patients aPWV increases more rapidly with age than controls but there was no difference in the pattern of change of Alx with age between groups.

P9.10 IN THE ELDERLY, ENDOTHELIAL DYSFUNCTION AND LOW-GRADE INFLAMMATION DO NOT PLAY A PROMINENT ROLE IN LOCAL ARTERIAL STIFFENING – THE HOORN STUDY -

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Purpose: To investigate, in the elderly, the association between plasma biomarkers of endothelial dysfunction (ED) and low-grade inflammation (LGI) on the one hand and greater local arterial stiffness of the carotid, femoral and brachial arteries on the other.

Methods: Plasma biomarkers for ED (vWf, sVCAM-1, sE-selectin, sICAM-1) and LGI (CRP, SAA, IL6, TNF- α , sICAM-1) were determined and combined into mean z-scores, in a cohort stratified by glucose tolerance status (GTS) (n=745; DM2=275, IGM=183, NGM=287; age 68.7±7.0 years). Ultrasonography was used to measure arterial properties and local arterial stiffness estimates were calculated: distensibility (DC) and compliance coefficients (CC), in all arteries, and the carotid Young's elastic modulus (YEM). Linear regression analyses were used to investigate the above associations.

Results: The study population was characterized by a high prevalence of prior cardiovascular disease (CVD) (48%), hypertension (70%) and use of

lipid-lowering (17%) and anti-hypertensive (39%) medication. After adjustment for sex, age, mean arterial pressure and GTS, ED was not associated with carotid, femoral or brachial stiffness (e.g. for the carotid artery (β (95%CI): DC: 0.19(-0.24;0.62), CC: -0.01(-0.03;0.02) and YEM: 0.03 (-0.03;0.08)). LGI was not associated with carotid, femoral or brachial stiffness, except for YEM (0.07(0.02;0.12); other data not shown).

Discussion: In an elderly population at high CVD risk, ED and LGI were not associated with local arterial stiffness, except for LGI and YEM. This suggests that ED and LGI, as estimated by these markers, do not play a prominent role in arterial stiffening, as estimated by these local arterial stiffness estimates.

P9.11 ARTERIAL STIFFNESS IS ASSOCIATED WITH DECREASED KIDNEY FUNCTION IN A PRIMARY CARE POPULATION: RESULTS FROM THE HIPPOCRATES-STUDY

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Introduction: Chronic kidney disease (CKD) is associated with cardiovascular morbidity and mortality. A common observation in both CKD and cardiovascular disease is increased arterial stiffness. Although many studies have focussed on patients with advanced CKD or established cardiovascular disease, relatively few studies have investigated whether increased arterial stiffness is predictive of renal damage in a less selected primary-care population.

Objective: In this study, we aimed to investigate whether increased arterial stiffness is associated with impaired renal function in a primary care population without overt CKD.

Design and Method: We performed a cross sectional analysis of data from the HIPPOCRATES-study, a study investigating hypertension and cardiovascular complications in a primary care population. Carotid-femoral Pulse-Wave Velocity (cfPWV), blood-pressure measurements and laboratory data were available. The estimated Glomerular Filtration Rate (eGFR) was calculated using the Cockcroft-Gault formula, adjusted for the Body Surface Area.

Results: We studied 587 patients (283 males). The mean age of the population was 61.1 ± 10.6 years. The mean GFR was 69.8 ± 15.6 ml/min. In a linear regression model unadjusted for age the mean CF-PWV was inversely associated with GFR. (β -0.16; $p<0.0001$). However, this relationship did not persist after correction for age. Body-mass index was an independent determinant of eGFR in both models.

Conclusion: In a primary care population carotid-femoral PWV is significantly associated with decreased kidney function, however this effect is mainly determined by age and body-mass index.

Therapeutic Aspects

P10.01

AORTIC STIFFNESS IN POLYMYALGIA RHEUMATICA: EFFECTS OF STEROID TREATMENT

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Increased arterial stiffness and cardiovascular risk have been observed in inflammatory diseases. Polymyalgia rheumatica (PMR) is a disease which affects primarily the elderly and exhibits evidence of a systemic inflammatory response, but little is known about aortic involvement in PMR. We investigated whether aortic stiffness is increased in PMR and whether it improves after steroid treatment.

Thirty-nine PMR patients (age 72±8 years, men 45%, blood pressure 134/75±16/9 mmHg) and 39 age-, sex- and blood pressure-matched control subjects underwent aortic pulse wave velocity (PWV) determination (applanation tonometry, Sphygmocor). Aortic augmentation as a measure of the impact of the reflection wave on central hemodynamics was also measured, and corrected for heart rate. Twenty-nine of the PMR patients were reexamined after 4-week treatment with prednisone (starting dose, 12.5-50 mg/day).

Aortic PWV was significantly higher in PMR patients than in control subjects (12.4±4 vs 10.2±2 m/s, $p<0.01$). Treatment was followed by a reduction in heart rate (from 78±12 to 70±10 bpm, $p<0.001$), and no significant change in BP (from 134/75±16/8 to 134/75±15/9 mmHg, both $p=n.s.$). As shown in the Figure, aortic PWV decreased significantly after steroid treatment (from 11.8±4 to 10.5±3 m/s, $p=0.015$), and the difference was independent from blood pressure and heart rate changes. Treatment was also associated with a significant reduction in aortic augmentation. Augmentation index