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P4.04: CARDIOVASCULAR PARAMETERS OTHER THAN MEAN ARTERIAL PRESSURE ARE PREDICTIVE OF DYNAMIC CHANGES IN AORTIC STIFFNESS IN THE RAT

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Table: Peak cap stress in kPa (mean±st. dev.)	Median Value	Peak cap stress in low group (kPa)	Peak cap stress in high group (kPa)	p-value
Min cap thickness	146 μ m	170 \pm 110	85 \pm 40	<0.05
Max necrotic core th.	537 μ m	172 \pm 113	83 \pm 30	<0.05
Necrotic core angle	58°	148 \pm 103	109 \pm 81	>0.05
Min intima thickness	196 μ m	142 \pm 130	116 \pm 38	>0.05
Min media thickness	78 μ m	138 \pm 100	119 \pm 90	>0.05
Min adventitia th.	47 μ m	124 \pm 82	134 \pm 108	>0.05

P4.03

THE RESERVOIR-WAVE PARADIGM INTRODUCES ARTEFACT INTO WAVE INTENSITY ANALYSIS: A COMPUTER MODELLING AND IN VIVO STUDY

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Background: Wave intensity (WI) has traditionally been calculated from measured pressure and velocity waveforms. In the recently proposed reservoir-wave approach, wave intensity is calculated using excess pressure (WI_{RW}), i.e. the difference between measured and reservoir pressures. Although it has been suggested that WI_{RW} is more accurate than WI [1], this has not yet been validated in a system with known wave reflection properties.

Methods. This study assessed WI and WI_{RW} in two one-dimensional models, first, a simple network containing three bifurcations (two producing positive reflection and one producing negative reflection) and second, a full model of the systemic arterial tree. In both, a forward component of pressure was prescribed at the (non-reflecting) inlet, thus forward waves were known *a priori*. 3-element windkessels formed the model outlets. Modelling results were compared with measurements in the ascending aorta of five adult sheep.

Results. WI accurately predicted the three reflection sites in the simple network, whereas WI_{RW} did not detect the first or third (positive) sites and overestimated the second (negative) site. In both models, an artefactual mid-systolic forward expansion wave appeared in WI_{RW} but not WI. In the systemic arterial model, WI_{RW} predicted predominant negative reflection during systole, whereas WI correctly predicted positive reflection; in vivo results were qualitatively similar.

Conclusion: WI_{RW} introduces artefactual expansion waves and attenuates or eliminates compression waves. This may limit the utility of WI_{RW} in the assessment of both forward-running waves and wave reflection in arterial networks.

[1] Tyberg J. et al, Med Biol Eng Comput, 47(2):221-232, 2009

P4.04

CARDIOVASCULAR PARAMETERS OTHER THAN MEAN ARTERIAL PRESSURE ARE PREDICTIVE OF DYNAMIC CHANGES IN AORTIC STIFFNESS IN THE RAT

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It has been observed that phenylephrine induced mean arterial pressure (MAP) changes in rats cause different values of aortic stiffness, measured by pulse wave velocity (PWV), at the same MAP during increasing, compared to decreasing pressure. The reason for this hysteresis in the MAP-PWV relationship is not understood. This study evaluates the influence of a variety of cardiovascular factors on PWV. Five, 12 week old Wistar-Kyoto rats were anaesthetised and two pressure sensors introduced into the thoracic and abdominal aorta. Pressure was increased with intravascular phenylephrine infusion (30 μ g/kg/min), and allowed to return to baseline following cessation of infusion. This was repeated following intravascular hexamethonium bolus (20mg/kg), blocking autonomic

activity to quantify the role of the sympathetic system. PWV, thoracic MAP, pulse pressure, form factor, maximum slope of pressure (dP/dt, surrogate measure of cardiac contractility), and R to R interval from the electrocardiogram were calculated for each individual pulse. These variables along with hexamethonium presence, and dynamic MAP direction were entered into a stepwise linear regression model (dependent variable: PWV). MAP, R to R interval, sympathetic activity, pulse pressure, dynamic MAP direction and dP/dt were significant predictors (Table). The results indicate that cardiovascular parameters other than MAP, especially pulse pressure, and notably sympathetic activity and R to R interval, are predictive of PWV. MAP direction remained in the model, indicating that other factors are at play, or that the aorta displays a short term viscoelastic memory. Further studies are required to isolate each of these factors and their effect on PWV.

	standardised coefficient	p	r ² change
MAP	0.242	<0.001	0.476
R to R interval	0.275	<0.001	0.105
sympathetic activity	-0.193	<0.001	0.034
pulse pressure	0.453	<0.001	0.005
MAP direction	0.129	<0.001	0.014
maximum dP/dt	-0.168	<0.001	0.011

P4.05

CHARACTERISTICS OF AORTIC STIFFNESS IN DIABETIC RATS TREATED WITH VITAMIN D

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Background: Diabetes mellitus is associated with macrovascular complications characterized by increased aortic stiffness. Mechanisms of diabetes-induced vascular impairment include persistent activation of the inflammatory system that may be improved by the immunomodulatory effects of orally administered vitamin D. This study aimed to examine the effects of diabetes on the elastic properties of the aorta and the protective effects of vitamin D treatment.

Methods: Diabetes was induced by injection of streptozotocin in male Wistar rats (age 4 months), followed by oral administration of cholecalciferol (500 IU/kg) for 10 weeks. Aortic pulse wave velocity (PWV) and other hemodynamic parameters were recorded over a mean arterial pressure (MAP) range of 50 to 200 mmHg using a dual pressure sensor catheter. Serum 25-hydroxyvitamin D (25[OH]D) level was measured using a radioimmuno assay.

Results: In diabetic rats, PWV was significantly elevated across MAP range between 120 and 200 mmHg. PWV across lower MAP range did not reveal any significant differences between all groups. Administration of vitamin D did not improve aortic stiffness, despite high levels of serum 25(OH)D in the treatment group, compared to the placebo group (513 \pm 132 nmol/L vs 108 \pm 38 nmol/L, respectively). Other hemodynamic parameters (heart rate, systolic and diastolic blood pressure, and pulse pressure) were not different between all groups at any given level of MAP.

Conclusion: PWV profile determined under isobaric conditions demonstrated increased aortic stiffness in diabetic rats that was not improved by vitamin D supplementation.