

# Decreased baroreflex gain more strongly predicts microalbuminuria and increased pulsatile stress than decreased RR E/I ratio in patients with type 1 diabetes

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## BACKGROUND AND AIMS

Long-lasting type 1 diabetes (T1DM) may be associated with cardiac autonomic neuropathy (CAN), increased pulse pressure (PP) or pulsatile stress, and microalbuminuria ( $\mu A$ ), all cardiovascular risk factors. We compared the relationships of two markers of CAN, RR E/I (Expiratory/Inspiratory) ratio and baroreflex gain (BRG), with  $\mu A$  and pulsatile stress during an active orthostatic test in patients with T1DM

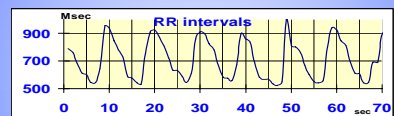


## MATERIALS AND METHODS



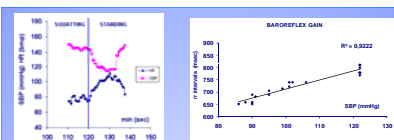
Deep breathing test  
6 breaths/min

Measurement of RR E/I ratio.



167 patients with T1DM  
(mean age 40 years,  
diabetes duration 20  
years, body mass index  
23.6 kg/m<sup>2</sup>, HbA1c 8.64%)

Continuous heart rate (HR)  
and arterial blood pressure (BP)  
monitoring (Finapres®)  
during a postural squat-  
stand test



The mirror changes in heart rate and systolic BP during the squat-stand transition allows the calculation of BRG (msec.mm Hg<sup>-1</sup>), by plotting the pulse intervals (RR) against systolic BP values, as classically assessed during a pharmacological test using the infusion of a vasodilator and a vasopressor agent.

Pulsatile stress was defined as the product of PP and heart rate, both during the whole test and during the squatting position only.



## RESULTS

N (F/M)

Age(yrs)

Diabetes duration (yrs)

$\mu A$  (mg/l)

HbA1c (%)

RR E/I ratio

BRG (msec.mm Hg<sup>-1</sup>)

Total pulsatile stress  
index (mmHg.min<sup>-1</sup>)

Squatting position  
pulsatile stress index  
(mmHg.min<sup>-1</sup>)

T1DM cohort separated in two  
subgroups according to the median  
value of RR E/I ratio

>1.25 <1.25 p

90 (42/48) 83 (45/38)

36 ± 11 45 ± 9 0.0001

16 ± 10 24 ± 11 0.0001

23 ± 79 48 ± 115 NS

8.6 ± 1.7 8.7 ± 1.3 NS

1.41 ± 0.14 1.14 ± 0.07 0.0001

3.19 ± 2.63 2.24 ± 1.84 0.0081

4666 ± 1299 5090 ± 1501 0.0545

4533 ± 1389 5352 ± 1697 0.0009

T1DM cohort separated in two  
subgroups according to the median  
value of BRG (mmHg.min<sup>-1</sup>)

> 2.20 <2.20 P

82 (41/41) 85 (45/40)

38 ± 12 42 ± 11 0.02

18 ± 12 21 ± 10 NS

10 ± 16 59 ± 133 0.019

8.6 ± 1.8 8.6 ± 1.2 NS

1.31 ± 0.16 1.25 ± 0.19 NS

4.51 ± 2.31 1.17 ± 0.61 0.0001

4521 ± 1310 5190 ± 1428 0.0019

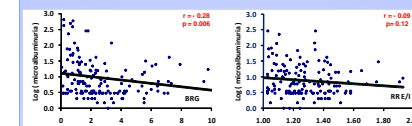
4396 ± 1472 5408 ± 1543 0.0001

- Compared to T1DM patients with high BRG, patients with low BRG tended to be slightly older and to have a slightly longer duration of diabetes, lower RR E/I ratio but had similar recent HbA1c levels

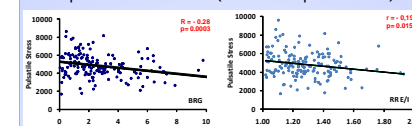
- T1DM patients with low BRG had an increased pulsatile stress index (specially in squatting position). Similarly,  $\mu A$  was higher in T1DM patients with low BRG, being expressed by the mean level or by its logarithm to adjust for a non Gaussian distribution.

- All together, 26.9 % of T1DM patients with low BRG had abnormal  $\mu A$  ( $\geq 30$  mg/l) versus only 5.3 % of patients with high BRG ( $p < 0.001$ ).

- There was an inverse correlation between BRG and log  $\mu A$  ( $r = -0.28$ ;  $p = 0.0006$ ), but not between RR E/I ratio and log  $\mu A$  ( $r = 0.09$ ;  $p = 0.12$ ).



- The correlation between BRG and pulsatile stress ( $r = -0.28$ ;  $p = 0.0003$ ) was stronger than that between RR E/I ratio and pulsatile stress ( $r = -0.19$ ;  $p = 0.0153$ ).



## CONCLUSIONS:

The calculation of BRG during a squat-stand test in subjects with T1DM allows to better detect patients with increased pulsatile stress and even more strongly patients with  $\mu A$  than the classical RR E/I ratio CAN index. Decreased BRG may be used to detect T1DM patients at high risk of cardiorenal complications.