Association Between Endothelial Function (Assessed on Reactive Hyperemia Peripheral Arterial Tonometry) and Obstructive Sleep Apnea, Visceral Fat Accumulation, and Serum Adiponectin

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Background: Visceral obesity, low adiponectin, and severe obstructive sleep apnea (OSA) are associated with cardiovascular diseases, but the interactions among these factors on endothelial dysfunction are not well known.

Methods and Results: Endothelial function in 133 patients after polysomnography was evaluated as reactive hyperemia index (RHI) on reactive hyperemia peripheral arterial tonometry. Visceral obesity was defined as visceral fat area ≥100 cm² on computed tomography. RHI was significantly correlated with apnea hypopnea index (AHI), visceral fat area, and serum adiponectin (r=−0.24, P=0.0055, r=−0.19, P=0.031, and r=0.20, P=0.019, respectively). RHI in patients with visceral obesity was significantly decreased in the presence of severe OSA (AHI ≥30; P=0.042). On multivariate regression analysis, only severe OSA remained as an independent predictive factor of RHI (P=0.024, R²=5.4%). RHI in patients with severe OSA (n=44) was significantly improved after 3 months of continuous positive airway pressure (CPAP) treatment (1.78±0.40 before CPAP vs. 2.00±0.53 after CPAP, P=0.013), similarly to those with AHI <30 (P=0.45).

Conclusions: Severe OSA, but not visceral fat area or serum adiponectin, was independently associated with endothelial function according to RHI. In addition, impaired endothelial function was reversible following 3 months of CPAP treatment.

Key Words: Adiponectin; Continuous positive airway pressure; Endothelial function; Obstructive sleep apnea; Visceral obesity
function related to coexisting OSA and obesity. To our knowledge, there is no study using RH-PAT to investigate the additive effect of OSA on endothelial dysfunction due to visceral obesity.

We hypothesized that severe OSA exacerbated endothelial dysfunction in patients with visceral obesity. Thus, using RH-PAT, we investigated the association and interrelationships of endothelial function with visceral obesity, serum adiponectin, and OSA in addition to the effects of continuous positive airway pressure (CPAP) treatment.

**Methods**

**Subjects**

Study patients were consecutively recruited from the Sleep Unit of the Kyoto University Hospital between September 2009 and February 2013. All had been referred to this sleep unit due to suspicion of OSA with symptoms such as habitual snoring or daytime sleepiness. None had been previously diagnosed with or treated for OSA. Patients with pulmonary diseases, chronic infection, or history of cancer or collagen disease were excluded. This study was approved by the Kyoto University Graduate School and Faculty of Medicine Ethics Committee (approval number E-2091), and written informed consent was obtained from all patients.

**Polysomnography, CPAP and Follow-up**

The diagnosis of OSA was confirmed on polysomnography (PSG; SomnoStar pro, Cardinal Health, Dublin, OH, USA or Alice 4, Philips Respironics, Murrysville, PA, USA), which was started at 22:00 hours and ended at 06:00 hours the following morning. Surface electrodes were attached using standard techniques to obtain an electro-oculogram, electromyogram of the chin, and 12-lead electroencephalograph. Sleep stages were defined according to the criteria of Rechtschaffen and Kales. Ventilation was monitored on inductive plethysmography (Respitrace QDC, Viasys Healthcare, Palm Springs, CA, USA). Airflow was monitored with a nasal pressure transducer and supplemented by an oronasal thermal sensor. Arterial oxygen saturation (SpO$_2$) was monitored continuously with a pulse oximeter. Apnea was defined as the complete cessation of airflow and hypopnea as a clear decrease in airflow ≥50% lasting for ≥10 s accompanied by a decrease in SpO$_2$ of at least 3% and/or associated with arousal. Apnea-hypopnea index (AHI) was calculated as the number of episodes of apnea and hypopnea per hour over the total sleep time. Arousals were scored using the American Sleep Disorders Association’s 3-s definition, and the arousal index was calculated according to the number of arousals per h of sleep. Nocturnal oxygen desaturation was assessed as the minimum SpO$_2$ during sleep and SpO$_2$ <90% time per total sleep time. Patients with predominantly central sleep apnea were excluded. OSA severity was defined as follows: none-mild OSA (AHI <15), moderate OSA (15≤AHI<30) and severe OSA (AHI ≥30). Those who agreed to CPAP underwent a second PSG with CPAP titration. We implemented CPAP with the autoadjusting positive airway pressure function for all patients. During the second sleep study, optimal minimum and maximum positive airway pressure were adjusted by the attending sleep laboratory physicians to abolish all respiratory events.

At the 3-month follow-up, we urged the patients to undergo a third sleep study to confirm whether an adjustment of the CPAP setting was necessary. To investigate the effect of CPAP treatment, at the third sleep study endothelial function data and blood samples were collected in the same way as at the first sleep study.

**RH-PAT**

We evaluated endothelial dysfunction with RH-PAT in the morning following diagnostic and follow-up PSG. Previously, RH-PAT has been used to measure endothelial function in OSA patients. With this method, specially designed finger probes (EndoPAT 2000, Itamar Medical, Caesarea, Israel) were placed on the index finger of each hand. Pulsatile arterial volume changes induced pressure alterations in the finger cuff, which were sensed by pressure transducers and recorded by the computer. The RH-PAT protocol consisted of a 5-min baseline measurement, after which a blood pressure cuff on the test arm was inflated to 60 mmHg above the baseline systolic blood pressure or at least 200 mmHg for 5 min. Occlusion of pulsatile arterial flow was confirmed by the reduction of the PAT tracing to zero. After 5 min, the cuff was deflated, and the PAT signal after cuff release compared with the baseline PAT signal was calculated via computer algorithm that automatically normalized for the baseline signal and was indexed to the control arm. As the index of endothelial function, the reactive hyperemia index (RHI) was calculated as the ratio of the average amplitude of PAT signal over 1 min starting 1.5 min after cuff deflation (control arm, A; occluded arm, C) divided by the average amplitude of the PAT signal of the 2.5-min time period before cuff inflation (baseline; control arm, B; occluded arm, D). RHI=(C/D)/(A/B)×baseline correction. Higher RHI represents a better preserved endothelial response to hyperemia.

**Blood Sampling and Serum Adiponectin**

Blood samples were drawn at 07:00 hours after the subjects had fasted beginning at 20:00 hours the previous night. Blood samples were centrifuged immediately at 1,370×g at 4°C for 10 min. The separated samples were stored at −80°C until the assay. Serum adiponectin was measured using a latex agglutination kit (Otsuka Pharmaceutical, Tokyo, Japan). The minimum detection limit for adiponectin with this kit was 0.5 μg/ml. The intra- and inter-assay coefficients of variation for adiponectin were 0.8–1.09% and 1.6–3.5%, respectively.

**Visceral and Subcutaneous Fat Area**

Visceral and subcutaneous fat area was assessed using an Aquilion 64 computed tomography (CT) system (Toshiba Medical Systems, Tochigi, Japan) running on 135kVp, 440 mA, 0.5-s scan time and 10.0-mm slice thickness. We used a single CT scan obtained at the level of the umbilicus, and the visceral and subcutaneous fat areas were quantified using a specialized image analysis program (AZE Virtual Place 99, AZE of America, Irvine, CA, USA).

**Definition of Comorbidities**

According to the criteria for Japanese subjects, we defined comorbidities as follows. BMI ≥25 kg/m$^2$ was defined as obesity. Visceral obesity was determined to be positive at visceral fat area ≥100 cm$^2$ in both men and women. WC was measured at the level of the navel with the patient standing, and abdominal obesity was determined to be positive at WC ≥85 cm for men and ≥90 cm for women. A diagnosis of MetS required the subject to have abdominal obesity and 2 or 3 of the following: (1) dyslipidemia (triglycerides ≥150 mg/dl and/or high-density lipoprotein cholesterol <40 mg/dl or specific treatment for these lipid abnormalities); (2) hypertension (systolic blood pressure ≥130 mmHg and/or diastolic blood pressure ≥85 mmHg or treatment of previously diagnosed hypertension); and (3) hyperglycemia (fasting plasma glucose ≥110 mg/dl or specific
133 patients underwent PSG, venous blood testing, abdominal CT and RH-PAT

23 patients were diagnosed with non-mild OSA
54 patients were diagnosed with moderate OSA

56 patients were diagnosed with severe OSA and introduced to CPAP therapy

12 patients could not have follow-up PSG
4 patients refused follow-up PSG
7 patients discontinued CPAP therapy
1 patient changed hospital

44 patients with severe OSA had follow-up PSG, venous blood testing and RH-PAT after 3 months

Results

Subject Characteristics
For 133 patients, PSG, venous blood testing, abdominal CT and RH-PAT were performed. Fifty-six patients were diagnosed as having severe OSA and were indicated to undergo CPAP therapy. Seven patients, however, subsequently withdrew from CPAP therapy, 1 patient changed hospital, and 4 patients refused follow-up PSG. As a result, 44 patients with severe OSA underwent follow-up PSG, venous blood testing, and RH-PAT at 3 months (Figure 1). There were no significant differences in characteristics (such as age, sex, blood pressure, BMI, Brinkman index, AHI) between patients who did and did not undergo follow-up PSG (Table S1). The baseline characteristics and PSG data of the total 133 patients are given in Tables 1, 2. (Subject characteristics according to sex are given in Table S2.)

Anthropometric parameters and blood pressure were measured immediately after PSG recording ended.

Statistical Analysis
Statistical analysis was done using JMP version 9.0.0 (SAS Institute, Cary, NC, USA). P<0.05 was considered to be statistically significant. Results are expressed as mean±SD. First, we classified the patients depending on the severity of OSA and compared their clinical backgrounds. The significance of intergroup differences was determined on analysis of variance. When a significant difference was found, we used Tukey’s post-hoc test to identify where the difference was significant. Chi-squared test was used to compare categorical variables. Second, we evaluated the relationships among RHI and clinical variables using Pearson’s or Spearman’s correlation coefficients. Unpaired t-test was used for comparing RHI between 2 groups. Based on the results of these analyses, multiple regression analysis was done to determine the contribution of clinical variables to endothelial function. The variables entered into the multivariate analysis were those with P<0.10 on univariate analysis. Third, we evaluated the impact on endothelial function of the coexistence of OSA and visceral obesity. In this subgroup analysis, the significance of intergroup differences was determined using Steel-Dwass test. Fourth, paired t-test was used to compare clinical variables before and after CPAP treatment.

Endothelial Function, Visceral Fat Accumulation, Serum Adiponectin and OSA
RHI was significantly correlated with AHI (r=−0.24, P=0.0055), visceral fat area (r=−0.19, P=0.031), and serum
significantly affect RHI (Table 3). On multivariate regression analysis, severe OSA remained as an independent predictive factor for RHI (P=0.024, R²=5.4%), but visceral fat area and serum adiponectin level did not (Table 3).

**Additive Changes in RHI in Severe OSA Due to Visceral Obesity**

On subgroup analysis, RHI in patients with visceral obesity significantly decreased in the presence of severe OSA (P=0.042). Patients with coexistent visceral obesity and severe OSA had significantly lower RHI than those with neither (P=0.016; Figure 3).

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**Table 1. Baseline Subject Characteristics**

<table>
<thead>
<tr>
<th></th>
<th>All patients</th>
<th>AHI &lt;15</th>
<th>15≤AHI&lt;30</th>
<th>30≤ AHI</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>No. subjects</td>
<td>133</td>
<td>23</td>
<td>54</td>
<td>56</td>
<td></td>
</tr>
<tr>
<td>Age (years)</td>
<td>56.8±12.7</td>
<td>55.5±16.8</td>
<td>57.4±11.2</td>
<td>56.7±12.2</td>
<td>0.82</td>
</tr>
<tr>
<td>Sex (M/F)</td>
<td>105 (79)/28 (21)</td>
<td>14 (61)/9 (39)</td>
<td>44 (81)/10 (19)</td>
<td>47 (84)/9 (16)</td>
<td>0.062</td>
</tr>
<tr>
<td>SBP (mmHg)</td>
<td>128±19</td>
<td>122±17</td>
<td>125±16</td>
<td>132±21</td>
<td>0.041*</td>
</tr>
<tr>
<td>DBP (mmHg)</td>
<td>80±12</td>
<td>76±10</td>
<td>78±11</td>
<td>84±13</td>
<td>0.0070*</td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>26.3±4.8</td>
<td>25.1±4.2</td>
<td>25.4±3.3</td>
<td>27.5±5.9</td>
<td>0.026*</td>
</tr>
<tr>
<td>WC (cm)</td>
<td>92±10.6</td>
<td>87.6±10.0</td>
<td>90.0±9.2</td>
<td>95.7±11.1</td>
<td>0.0014*</td>
</tr>
<tr>
<td>Abdominal obesity</td>
<td>97 (73)</td>
<td>13 (57)</td>
<td>37 (69)</td>
<td>47 (84)</td>
<td>0.029*</td>
</tr>
<tr>
<td>Waist hip ratio</td>
<td>0.97±0.049</td>
<td>0.95±0.056</td>
<td>0.96±0.048</td>
<td>0.99±0.040</td>
<td>0.0002*</td>
</tr>
<tr>
<td>Visceral fat area (cm²)</td>
<td>121±69</td>
<td>80.6±53</td>
<td>120±70</td>
<td>139±67</td>
<td>0.0024*</td>
</tr>
<tr>
<td>Visceral obesity</td>
<td>80 (60)</td>
<td>9 (39)</td>
<td>29 (54)</td>
<td>42 (75)</td>
<td>0.0057*</td>
</tr>
<tr>
<td>Subcutaneous fat area (cm²)</td>
<td>129±75</td>
<td>119±67</td>
<td>104±46</td>
<td>155±92</td>
<td>0.0010*</td>
</tr>
<tr>
<td>VFA/SFA ratio</td>
<td>1.14±0.81</td>
<td>0.91±0.90</td>
<td>1.28±0.71</td>
<td>1.10±0.85</td>
<td>0.17</td>
</tr>
<tr>
<td>Smoking status (current/ex/never)</td>
<td>11/35/87</td>
<td>2/5/16</td>
<td>3/14/37</td>
<td>6/16/34</td>
<td>0.82</td>
</tr>
<tr>
<td>Brinkman index</td>
<td>141±281</td>
<td>80.7±160</td>
<td>143±305</td>
<td>164±296</td>
<td>0.49</td>
</tr>
<tr>
<td>MetS</td>
<td>50 (38)</td>
<td>4 (17)</td>
<td>18 (33)</td>
<td>28 (50)</td>
<td>0.018*</td>
</tr>
<tr>
<td>Hypertension</td>
<td>84 (63)</td>
<td>9 (39)</td>
<td>31 (57)</td>
<td>44 (79)</td>
<td>0.0023*</td>
</tr>
<tr>
<td>Dyslipidemia</td>
<td>74 (56)</td>
<td>9 (39)</td>
<td>32 (59)</td>
<td>33 (59)</td>
<td>0.22</td>
</tr>
<tr>
<td>Total cholesterol (mg/dl)</td>
<td>197±40</td>
<td>186±39</td>
<td>193±36</td>
<td>205±43</td>
<td>0.10</td>
</tr>
<tr>
<td>LDL-C (mg/dl)</td>
<td>119±33</td>
<td>115±30</td>
<td>116±34</td>
<td>122±34</td>
<td>0.56</td>
</tr>
<tr>
<td>HDL-C (mg/dl)</td>
<td>51.6±13</td>
<td>55.0±14</td>
<td>52.4±12</td>
<td>49.5±14</td>
<td>0.20</td>
</tr>
<tr>
<td>Triglycerides (mg/dl)</td>
<td>140±103</td>
<td>117±73</td>
<td>125±60</td>
<td>164±138</td>
<td>0.065</td>
</tr>
<tr>
<td>Serum creatinine (mg/dl)</td>
<td>0.78±0.16</td>
<td>0.72±0.16</td>
<td>0.81±0.16</td>
<td>0.79±0.16</td>
<td>0.058</td>
</tr>
<tr>
<td>C-reactive protein (mg/dl)</td>
<td>0.01±0.19</td>
<td>0.13±0.32</td>
<td>0.063±0.12</td>
<td>0.15±0.17</td>
<td>0.049*</td>
</tr>
<tr>
<td>Serum adiponectin (μg/dl)</td>
<td>8.47±4.7</td>
<td>9.32±5.0</td>
<td>8.36±5.0</td>
<td>8.23±4.2</td>
<td>0.63</td>
</tr>
</tbody>
</table>

Data given as mean±SD or n (%). *P<0.05. Abdominal obesity was defined as WC ≥85cm in male subjects, ≥90cm in female subjects. Visceral obesity was defined as visceral fat area ≥100cm². AHI, apnea hypopnea index; BMI, body mass index; CVD, cardiovascular disease; DBP, diastolic blood pressure; HDL-C, high density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; MetS, metabolic syndrome; SBP, systolic blood pressure; SFA, subcutaneous fat area; VFA, visceral fat area; WC, waist circumference.

**Table 2. Polysomnography Data**

<table>
<thead>
<tr>
<th></th>
<th>All patients (n=133)</th>
<th>AHI &lt;15 (n=23)</th>
<th>15≤AHI&lt;30 (n=54)</th>
<th>30≤ AHI (n=56)</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>AHI (/h)</td>
<td>33.2±21.8</td>
<td>8.72±4.4</td>
<td>23.0±3.9</td>
<td>53.2±18.9</td>
<td>&lt;0.0001*</td>
</tr>
<tr>
<td>3% oxygen desaturation index (/h)</td>
<td>30.0±23.0</td>
<td>5.92±3.9</td>
<td>18.7±4.8</td>
<td>50.8±20.7</td>
<td>&lt;0.0001*</td>
</tr>
<tr>
<td>Arousal index (/h)</td>
<td>30.9±16.8</td>
<td>20.8±11.0</td>
<td>23.2±11.4</td>
<td>42.4±16.5</td>
<td>&lt;0.0001*</td>
</tr>
<tr>
<td>Sleep efficiency (%)</td>
<td>74.6±12.3</td>
<td>77.7±8.5</td>
<td>75.7±12.4</td>
<td>72.2±13.1</td>
<td>0.13</td>
</tr>
<tr>
<td>SpO₂ &lt;90% time per total sleep time (%)</td>
<td>13.6±21.3</td>
<td>0.96±1.5</td>
<td>3.51±3.7</td>
<td>28.5±26.1</td>
<td>&lt;0.0001*</td>
</tr>
<tr>
<td>Minimum SpO₂ (%)</td>
<td>78.6±11.6</td>
<td>87.6±4.3</td>
<td>81.0±9.2</td>
<td>72.5±12.4</td>
<td>&lt;0.0001*</td>
</tr>
</tbody>
</table>

Data given as mean±SD. *P<0.05. SpO₂, arterial oxygen saturation. Other abbreviation as in Table 1.
Impact of Severe OSA on Endothelial Dysfunction

(2) severe OSA was associated with endothelial dysfunction independently of visceral fat area and serum adiponectin level; and (3) endothelial function in patients with severe OSA improved to that for none-moderate OSA following 3 months of CPAP treatment, although serum adiponectin and WC did not improve.

Endothelial Function in the Presence of Visceral Obesity and OSA

Visceral obesity is an important risk factor for CVD. Visceral fat gain evaluated on abdominal CT is significantly correlated with impairment of endothelial function. The relationship between obesity and endothelial inflammation or increased vascular oxidative stress has also been reported. These reports,

Changes in Clinical Parameters After 3 Months of CPAP

The mean usage time for CPAP was 4.43±1.8 h/day. After CPAP for 3 months, RHI in severe OSA patients improved significantly (P=0.013; Table 4). RHI between patients with none-moderate OSA and those with severe OSA after CPAP treatment did not differ significantly (2.07±0.55 vs. 2.00±0.53, P=0.45), while the AHI following CPAP treatment significantly decreased (P<0.0001). Serum adiponectin, WC, BMI, and waist hip ratio were not significantly improved (Table 4).

Discussion

This is the first study using RH-PAT to investigate the impact of OSA on endothelial function with or without visceral obesity. The main findings were as follows: (1) the coexistence of severe OSA significantly impaired endothelial function in patients with visceral obesity; (2) severe OSA was associated with endothelial dysfunction independently of visceral fat area and serum adiponectin level; and (3) endothelial function in patients with severe OSA improved to that for none-moderate OSA following 3 months of CPAP treatment, although serum adiponectin and WC did not improve.

Figure 2. Reactive hyperemia index (RHI) vs. (A) apnea hypopnea index (AHI), (B) waist circumference, (C) visceral fat area, (D) subcutaneous fat area and (E) serum adiponectin. *P<0.05. r, correlation coefficient.
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Endothelial Function, Serum Adiponectin and OSA

Adiponectin is secreted by adipocytes and has a multiplicity of actions affecting the cardiovascular system, such as insulin resistance, atherosclerosis, intimal hyperplasia, endothelial dysfunction, and pathological cardiac hypertrophy. Low adiponectin is one of the mechanisms explaining the association between obesity and CVD. Also, adiponectin was shown to be associated with endothelial dysfunction evaluated using FMD. Similarly, in this study, serum adiponectin was positively correlated with endothelial dysfunction evaluated on RH-PAT. On multivariate analysis, however, a relationship between serum adiponectin level and endothelial function was not shown. The present results suggest that severe OSA might have a greater association with endothelial dysfunction than adiponectin.

In this study, serum adiponectin was not associated with OSA. Lower adiponectin in patients with OSA than in control subjects has been reported. Adiponectin, however, was associ-
Impact of Severe OSA on Endothelial Dysfunction

Several reports using FMD showed that OSA was associated with endothelial dysfunction and that treatment of OSA with CPAP significantly improved endothelial function.\(^{11,12,37,38}\) The present results were similar to that of these reports. FMD reflects endothelial function in larger vessels such as the brachial artery; in contrast, RH-PAT reflects endothelial function in the smaller vessels in the end of the finger. Using a forearm skin biopsy sample, it was suggested that patients with OSA with low cardiovascular risk had endothelial dysfunction in the microcirculation, which improved with CPAP.\(^{39}\) The present results supported this finding that OSA patients had endothelial dysfunction in the microcirculation. A recent review suggested that microvascular endothelial function may be an earlier indicator of CVD risk than macrovascular endothelial function.\(^{40}\) Using RH-PAT, we might be able to detect early cardiovascular risk in patients with OSA.

In 2 previous large epidemiological studies, including 1 that used RH-PAT, a significant association between OSA and endothelial function was shown in female subjects only.\(^{41,42}\) Although those studies had greater numbers of female participants than the present study, the prevalence of severe OSA was low. In addition to the present study, endothelial dysfunction was also seen in severe OSA patients in other studies, including 1 that used RH-PAT.\(^{43-45}\) In the present multivariate analysis, severe OSA was related to RHI independently of sex. Indeed, most of the patients who did not have hypertension, dyslipidemia, or hyperglycemia had moderate or severe OSA. Therefore, in this cohort, it might be difficult to detect differences in endothelial function between patients with and without hypertension, dyslipidemia, or hyperglycemia independent of the effect of OSA. Third, similar to the present results, in the Framingham Heart Study, which is an epidemiological study using RH-PAT, an association between hypertension and endothelial function was not shown.\(^{52}\) Conversely, in the other report using FMD, this association was found.\(^{46}\) Because RH-PAT and FMD focused on different vessels, these reports suggested that hypertension might have limited effects on the function of the small vessels evaluated using RH-PAT.\(^{52}\) Therefore, the difference in methodology might have influenced the present results. And fourth, the size of the present study cohort might not be sufficient to detect associations between endothelial dysfunction and hypertension, dyslipidemia or hyperglycemia. For reasons already noted, it was difficult to discuss fully the differences in results between the present study and the previous ones.\(^{46-48}\) Therefore, further studies with larger subject groups are needed to investigate these differences.

**Study Limitations**

The present study had a few limitations. First, CPAP was not allocated randomly. Unmeasured factors may have changed during CPAP therapy and could have contributed to the observed improvement in endothelial function. The reversal of endothelial dysfunction in the present study, however, suggested that OSA was mainly responsible for the change in endothelial function. We could not show causality or identify the mechanism because this was an observational study. We should confirm these results in a future randomized clinical trial. Second, we did not verify the RH-PAT results. The measurement of endothelial function as a clinical tool in daily practice is not well

**Table 4. Effect of 3-Month CPAP Treatment in Severe OSA (n=44)**

<table>
<thead>
<tr>
<th>Measure</th>
<th>Baseline</th>
<th>After 3 months of CPAP</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>RHI</td>
<td>1.78±0.40</td>
<td>2.00±0.53</td>
<td>0.013*</td>
</tr>
<tr>
<td>Serum adiponectin (µg/ml)</td>
<td>8.74±4.1</td>
<td>8.73±4.4</td>
<td>0.95</td>
</tr>
<tr>
<td>AHI (/h)</td>
<td>53.2±17.8</td>
<td>4.18±4.0</td>
<td>&lt;0.0001*</td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>27.4±6.1</td>
<td>27.5±5.9</td>
<td>0.84</td>
</tr>
<tr>
<td>WC (cm)</td>
<td>95.9±10.6</td>
<td>95.0±11.7</td>
<td>0.18</td>
</tr>
<tr>
<td>Waist hip ratio</td>
<td>0.99±0.041</td>
<td>0.97±0.038</td>
<td>0.076</td>
</tr>
<tr>
<td>SBP (mmHg)</td>
<td>131±18</td>
<td>123±15</td>
<td>&lt;0.0001*</td>
</tr>
<tr>
<td>DBP (mmHg)</td>
<td>83±11</td>
<td>76±12</td>
<td>&lt;0.0001*</td>
</tr>
<tr>
<td>C-reactive protein (mg/dl)</td>
<td>0.14±0.18</td>
<td>0.082±0.099</td>
<td>0.018*</td>
</tr>
<tr>
<td>Total cholesterol (mg/dl)</td>
<td>199±42</td>
<td>191±47</td>
<td>0.046*</td>
</tr>
<tr>
<td>LDL-C (mg/dl)</td>
<td>117±32</td>
<td>111±29</td>
<td>0.12</td>
</tr>
<tr>
<td>HDL-C (mg/dl)</td>
<td>49.7±11</td>
<td>50.5±12</td>
<td>0.32</td>
</tr>
<tr>
<td>Triglycerides (mg/dl)</td>
<td>159±148</td>
<td>148±141</td>
<td>0.079</td>
</tr>
</tbody>
</table>

Data given as mean±SD. *P<0.05. CPAP, continuous positive airway pressure; OSA, obstructive sleep apnoea. Other abbreviations as in Tables 1, 3.
established. A further study would be needed with both RH-PAT and another method such as FMD to acquire comprehensive vascular information on OSA patients.

Conclusions

Severe OSA, but not visceral fat area or serum adiponectin level, was independently associated with endothelial function according to RHI. The coexistence of severe OSA impaired endothelial function in patients with visceral obesity. In addition, the disturbance in endothelial function improved significantly after 3 months of CPAP treatment.

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Disclosures

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References


Supplementary Files

Supplementary File 1
Table S1. Subject characteristics vs. follow-up PSG
Table S2. Subject characteristics vs. sex
Please find supplementary file(s); http://dx.doi.org/10.1253/circj.CJ-14-1303