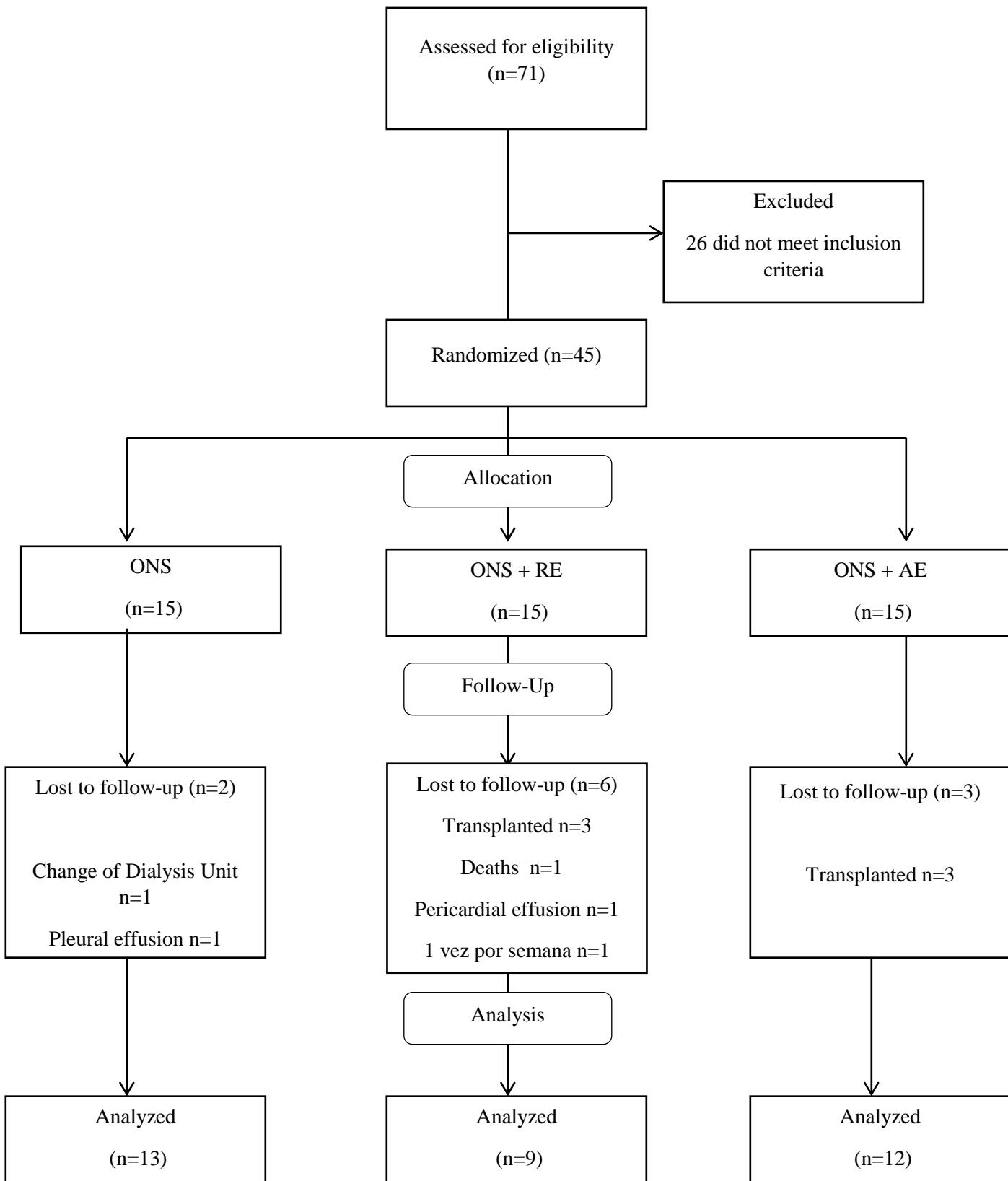


Participant flow

Figure 1. Screening, Randomization, and Follow-up according to the CONSORT diagram.



Baseline characteristics

Table 1. Baseline demographic, body composition, laboratory and physical function characteristics of the study population

| | TOTAL (n=45) | ONS (n=15) | ONS + RE (n=15) | ONS + AE (n=15) | p |
|--|-----------------|---------------|--------------------|--------------------|--------------|
| Age (years) | 29 ± 9.3 | 27.1 ± 8.1 | 30.3 ± 10.8 | 31.58 ± 9.5 | 0.487 |
| Sex (male) | 21 (46.7) | 9 (60) | 5 (33.3) | 7 (46.7) | 0.343 |
| Etiology | | | | | 0.740 |
| Unknown | 36 (80) | 12 (80) | 13 (87.7) | 11 (73.3) | |
| Diabetes mellitus | 2 (4.4) | 1 (6.7) | 1 (6.7) | 1 (6.7) | |
| Glomerulopathy | 2 (4.4) | 1 (6.7) | 0 (0) | 1 (6.7) | |
| Hypertension | 3 (6.7) | 0 (0) | 1 (6.7) | 2 (13.3) | |
| Other | 2 (4.4) | 1 (6.7) | 1 (6.7) | 0 (0) | |
| Frequency of dialysis | | | | | 0.562 |
| 2 times per week | 39 (86.7) | 13 (86.7) | 12 (80) | 14 (93.3) | |
| 3 times per week | 6 (13.3) | 2 (13.3) | 3 (20) | 1 (6.7) | |
| Dialysis vintage, months | 21 (6.5,38) | 28(8,48) | 19(8,36) | 24(4,36) | 0.487 |
| Comorbidities | | | | | 0.593 |
| Diabetes | 2 (4.4) | 1 (6.7) | 0 (0) | 1 (6.7) | |
| Hypertension | 45 (100) | 15 (100) | 15 (100) | 15 (100) | |
| Vascular access | | | | | 0.757 |
| Catheter | 27 (60) | 8 (53.3) | 10 (66.7) | 9 (60) | |
| AV fistula | 18 (40) | 7 (46.7) | 5 (33.3) | 6 (40) | |
| Body composition | | | | | |
| Weight (kg) | 52.4 ± 8.2 | 52 ± 9.7 | 53.3 ± 6 | 52.2 ± 85 | 0.939 |
| BMI (kg/m²) | 20.3 ± 2.4 | 19 ± 1.8 | 21.5 ± 1.9 | 19.7±3.1 | 0.040 |
| MAC (cm) | 24.5 ± 2.9 | 23.8 ± 2.6 | 25.3 ± 2.8 | 23.6 ± 3.3 | 0.313 |
| AMC (mm) | 216 ± 25.2 | 217 ± 29.9 | 216.2 ± 20.8 | 205.4 ± 26.4 | 0.788 |
| AMA (cm²) | 31.2 ± 8.8 | 31.6 ± 10.5 | 31 ± 7.5 | 27.6 ± 8.7 | 0.833 |
| FM% from anthropometry | 17.7 ± 7 | 14.3 ± 5.7 | 21.2 ± 6.1 | 19.6 ± 7.5 | 0.043 |
| Triceps skinfold thickness (mm) | 9.3 ± 4.6 | 6.7 ± 3 | 11.8 ± 5.8 | 10 ± 4.6 | 0.042 |
| Resistance (ohm) | 595 ± 127.4 | 601 ± 125.7 | 562.5 ± 135.7 | 622.7 ± 131.2 | 0.402 |
| Reactance(ohm) | 54.4 ± 19.8 | 52.6 ± 17.5 | 54.7 ± 23.5 | 63.2 ± 18 | 0.577 |
| Phase angle (°) | 5 ± 1.1 | 4.8 ± .86 | 5.4 ± 1.4 | 5.6 ± .79 | 0.866 |
| Laboratory tests | | | | | |

| | | | | | |
|---|----------------|---------------|---------------|----------------|--------------|
| Haemoglobin (g/dl) | 9.9 ± 2.3 | 10.8 ± 2.3 | 10.1 ± 1.4 | 10.2 ± 2.1 | 0.636 |
| Total lymphocytes count (cells/mm³) | 1187 ± 405 | 1100 ± 360 | 1231.5 ± 515 | 1249 ± 328 | 0.586 |
| Creatinine mg/dl | 12.9 ± 3.2 | 13.7 ± 3.3 | 12.8 ± 3.2 | 12.1 ± 3.2 | 0.422 |
| Albumin (g/dL) | 3.7 ± .45 | 3.7 ± .48 | 3.7 ± .56 | 3.6 ± .31 | 0.888 |
| Phosphorus (mg/dl) | 5.7 (4.7,6.7) | 5.4 (4.7,3) | 6.3 (5.3,7.2) | 5.6 (4.6,5.9) | 0.549 |
| Potassium (mmol/L) | 5.5 ± .85 | 5.5 ± .81 | 5.5 ± .72 | 5.4 ± 1 | 0.895 |
| CRP (mg/L) | 4.2 (2.8,10.5) | 3.6 (2.8,5.2) | 4.6 (2.3,12) | 7.2 (3.1,13.8) | 0.781 |
| Physical Function | | | | | |
| Six-minute walk (m) | 435 ± 61 | 463.8 ± 51.7 | 424.7 ± 61.7 | 419.2 ± 62.7 | 0.090 |
| Six-minute walk (m/s) | 1.2 ± .16 | 1.2 ± .14 | 1.1 ± .17 | 1.1 ± .17 | 0.086 |
| Time Up and Go (s) | 7.3 ± 1.3 | 7 ± 1.2 | 7.5 ± 1.6 | 7.5 ± 1.3 | 0.671 |
| Sit to stand (s) | 10.16 ± 2.9 | 9.4 ± 2.1 | 9.7 ± 2.3 | 11.2 ± 3.8 | 0.284 |
| Handgrip strength (kg) | 22.5 ± 7.4 | 25 ± 8.9 | 21.6 ± 7.9 | 20.8 ± 4.6 | 0.279 |
| Physical Activity (Kcal from PAQ) | 2211.9 ± 591 | 2256.6 ± 722 | 2223.7 ± 502 | 2154.7 ± 539 | 0.914 |

Data are indicated as absolute number (percentage), mean ± SD or median (first and third quartiles). ONS, oral nutritional supplementation; ONS + RE, oral nutritional supplementation plus resistance exercise; ONS + AE, oral nutritional supplementation + aerobic exercise; BMI, body mass index; MAC, midarm circumference; AMC, arm muscle circumference; AMA, arm muscle area; FM%, fat mass as a percentage of body weight; PAQ, physical activity questionnaire.

Outcome measures

Table 2. Effect size (Cohen's-d) calculation for physical function tests

| | ONS | ONS + RE | ONS + AE |
|---------------------------------|------|----------|----------|
| Six-minute walk test (m) | 0.35 | 0.94 | 1.11 |
| Sit to stand test (s) | 0.52 | 0.81 | 1.20 |
| Timed up and go test (s) | 0.91 | 1.04 | 1.6 |
| Handgrip strength (kg) | 0.11 | 1.01 | 0.60 |

Cohen's -d was calculated considering repeated measures. ONS, oral nutritional supplementation; ONS + RE; oral nutritional supplementation plus resistance exercise; ONS + AE; oral nutritional supplementation plus aerobic exercise.

Table 3. Changes in body composition measured by anthropometrics and bioelectric impedance analysis

| Variables | ONS | | | ONS + RE | | | ONS + AE | | | |
|--|-----------------|---------------|--------------|----------------|--------------|--------------|-----------------|---------------|--------------|----------------|
| | Baseline (n=13) | Final (n=13) | p | Baseline (n=9) | Final (n=9) | p | Baseline (n=12) | Final (n=12) | p | P [†] |
| Weight (kg) | 52 ± 9.7 | 53 ± 9.3 | 0.032 | 53.3 ± 6 | 54.9 ± 5.1 | 0.006 | 52.2 ± 8.5 | 52.8 ± 8.3 | 0.097 | 0.216 |
| BMI (kg/m²) | 19.4 ± 1.6 | 19.8 ± 1.7 | 0.035 | 21.4 ± 1.2 | 22.1 ± 1 | 0.006 | 20.8 ± 2.8 | 21 ± 2.6 | 0.123 | 0.209 |
| MAC (cm) | 24.1 ± 2.3 | 24.4 ± 2.1 | 0.179 | 25.8 ± 2.1 | 26.2 ± 2.1 | 0.209 | 25.3 ± 2.8 | 25.5 ± 2.7 | 0.590 | 0.843 |
| AMC (mm) | 217.9 ± 27.6 | 219.9 ± 24 | 0.410 | 224.4 ± 23.1 | 225.1 ± 23.4 | 0.839 | 220.6 ± 24.7 | 219 ± 25 | 0.666 | 0.685 |
| AMA (cm²) | 31.8 ± 9.7 | 34.4 ± 8.4 | 0.500 | 33.9 ± 8.5 | 34.2 ± 8.6 | 0.821 | 32.7 ± 8.8 | 32.4 ± 8.8 | 0.684 | 0.752 |
| FM % from anthropometry | 14.3 ± 5.7 | 15.1 ± 5.6 | 0.082 | 21.2 ± 6.1 | 22.4 ± 6.2 | 0.031 | 19.6 ± 7.5 | 20.3 ± 7.7 | 0.316 | 0.797 |
| Triceps skinfold thickness (mm) | 7.3 ± 3 | 7.8 ± 2.6 | 0.139 | 10.8 ± 3.2 | 11.7 ± 3.2 | 0.035 | 10.5 ± 4.3 | 11.2 ± 4.6 | 0.222 | 0.780 |
| R (ohm) | 563.8 ± 120.2 | 577.2 ± 115.8 | 0.547 | 634.5 ± 118.6 | 647 ± 114.5 | 0.583 | 597.5 ± 148.2 | 598.1 ± 156.3 | 0.982 | 0.488 |
| Xc (ohm) | 52.6 ± 14.7 | 56.2 ± 14.6 | 0.410 | 61 ± 17.6 | 63.9 ± 15.1 | 0.489 | 58 ± 26.3 | 60.6 ± 23.4 | 0.511 | 0.983 |
| Phase angle (°) | 5.3 ± 1 | 5.5 ± 1.3 | 0.613 | 5.3 ± 3.7 | 5.6 ± .93 | 0.289 | 5.1 ± 1.3 | 5.6 ± 1.3 | 0.067 | 0.776 |
| Physical activity (Kcal from PAQ) | 2130 ± 586.3 | 2067 ± 502 | 0.569 | 2186 ± 523.6 | 2258 ± 254 | 0.695 | 2151.4 ± 565 | 2021.5 ± 368 | 0.322 | 0.591 |

Anthropometric indicators are mean ± SD or median (first and third quartiles).

ONS, oral nutritional supplementation; ONS + RE, oral nutritional supplementation plus resistance exercise; ONS +AE, oral nutritional supplementation plus aerobic exercise;

BMI, body mass index; MAC, midarm circumference; AMC, arm muscle circumference; AMA, arm muscle area; FM%, fat mass; PAQ, physical activity questionnaire

as percentage of body weight; R, resistance at 50 kHz; Xc, reactance at 50 kHz.

p[†]=Repeated Measures ANOVA (3 groups x 2 times)

Table 4. Changes in biochemical indicators

| | ONS | | | ONS + RE | | | ONS + AE | | | <i>p</i> | <i>p</i> [†] |
|--|-----------------|---------------|--------------|----------------|---------------|--------------|-----------------|-------------------------|--------------|--------------|-----------------------|
| | Baseline (n=13) | Final (n=13) | <i>p</i> | Baseline (n=9) | Final (n=9) | <i>p</i> | Baseline (n=12) | Final (n=12) | <i>p</i> | | |
| Hemoglobin (g/dl) | 10.8 ± 2.3 | 10.6 ± 1.8 | 0.083 | 10.1 ± 1.4 | 11.7 ± 1.7 | 0.673 | 10.2 ± 2.1 | 9.8 ± 2.3 | 0.866 | 0.210 | |
| Total lymphocytes count (cell/mm³) | 1134 ± 375 | 1443 ± 506 | 0.001 | 1260.6 ± 486.8 | 1327 ± 431.7 | 0.003 | 1215 ± 347 | 1128.3 ± 399 | 0.048 | 0.009 | |
| Creatinine (mg/dl) | 14 ± 3.5 | 12.3 ± 3.4 | 0.078 | 12.6 ± 3.1 | 11.9 ± 3.9 | 0.095 | 12.6 ± 3.3 | 11.7 ± 3.6 | 0.037 | 0.732 | |
| Albumin (g/dL) | 3.8 ± 0.47 | 3.9 ± 0.40 | 0.028 | 3.8 ± 0.52 | 3.8 ± 0.44 | 0.037 | 3.6 ± .32 | 3.4 ± 0.55 ^a | 0.561 | 0.423 | |
| Phosphorus (mg/dl) | 5.4 (4.2,7.5) | 5.8 (3.6,7.9) | 0.780 | 6 (4.7,6.4) | 7 (6.8,7) | 0.123 | 5.6 (5.2,6.3) | 5.5(4.6,7.8) | 0.814 | 0.234 | |
| Potassium (mmol/L) | 5.6 ± 0.86 | 5.2 ± 0.92 | 0.024 | 5.6 ± 0.6 | 5.6 ± 0.58 | 0.788 | 5.4 ± 1.1 | 5.5 ± .74 | 0.656 | 0.405 | |
| CRP (mg/L) | 3.5 (2.4,4.4) | 3.4 (1.9,6.1) | 0.834 | 3.4 (2.6,9.2) | 6.2 (3.8,9.3) | 0.110 | 7.2 (3.3,12.6) | 5.4 (2.8,17.1) | 0.784 | 0.441 | |

Biochemical indicators are indicated as mean ± SD or median (first and third quartiles). CRP: C-Reactive Protein; ONS, oral nutritional supplementation; ONS + RE; oral nutritional supplementation plus resistance exercise; ONS + AE; oral nutritional supplementation plus aerobic exercise.

p[†]=Repeated Measures ANOVA (3 groups x 2 times)

Table 5. Changes in quality of life

| | ONS | | ONS + RE | | ONS + AE | |
|--------------------------------------|--------------|--------------|--------------|---------------------------|--------------|------------------------|
| | Pre | Post | Pre | Post | Pres | Post |
| Specific part | | | | | | |
| Symptoms | 80.80 ± 8.9 | 76.87 ± 13.4 | 77.39 ± 15.6 | 83.94 ± 12.5 | 76.33 ± 13.6 | 80.75 ± 11.7 |
| Effects of Kidney disease | 72.92 ± 21.8 | 64.59 ± 18.1 | 56.70 ± 19.9 | 76.79 ± 23.8 | 73.06 ± 13.7 | 73.06 ± 17.0 |
| Burden of kidney disease | 50.01 ± 16.2 | 56.96 ± 19.6 | 46.43 ± 11.9 | 50.90 ± 19.2 | 55.48 ± 16.8 | 64.86 ± 17.9 |
| Work status | 61.11 ± 33.3 | 50.00 ± 35.4 | 28.57 ± 26.7 | 42.86 ± 34.4 | 31.25 ± 45.8 | 37.50 ± 44.3 |
| Cognitive function | 20.00 ± 16.3 | 24.46 ± 27.3 | 8.557 ± 12.6 | 20.00 ± 21.4 | 26.66 ± 16.3 | 27.51 ± 33.3 |
| Quality of social interaction | 25.19 ± 15.2 | 25.92 ± 25.3 | 17.14 ± 16.3 | 11.43 ± 13.8 | 27.50 ± 18.3 | 31.66 ± 23.3 |
| Sexual function | 29.17 ± 44.1 | 8.333 ± 25.0 | 23.21 ± 41.1 | 41.07 ± 51.4 | 54.69 ± 47.2 | 29.69 ± 42.2 |
| Sleep | 73.06 ± 13.1 | 69.72 ± 6.9 | 69.29 ± 9.4 | 77.50 ± 10.7 [‡] | 69.06 ± 5.5 | 72.50 ± 8.4 |
| Social Support | 72.21 ± 20.4 | 68.53 ± 10.0 | 78.57 ± 12.6 | 76.19 ± 16.2 | 66.68 ± 12.6 | 77.08 ± 21.7 |
| Dialysis staff encouragement | 81.94 ± 11.0 | 80.56 ± 12.6 | 83.93 ± 17.2 | 83.93 ± 15.7 | 75.00 ± 21.1 | 82.81 ± 16.3 |
| Patient satisfaction | 53.33 ± 15.0 | 52.22 ± 13.4 | 68.57 ± 3.8 | 57.14 ± 17.1 | 52.50 ± 13.4 | 60.00 ± 10.7 |
| Generic part | | | | | | |
| Physical function | 83.33 ± 12.7 | 84.44 ± 22.7 | 75.00 ± 20 | 80.00 ± 13.2 | 74.38 ± 16.1 | 74.38 ± 14.7 |
| Physical role | 61.11 ± 46.9 | 88.89 ± 33.3 | 46.43 ± 44.3 | 85.71 ± 37.8 | 34.38 ± 44.2 | 71.88 ± 45.2 |
| Pain | 73.61 ± 27.6 | 85.00 ± 18.3 | 88.93 ± 15.9 | 86.79 ± 14.6 | 63.44 ± 17.6 | 71.88 ± 29.3 |
| General Health perceptions | 52.78 ± 17.1 | 53.33 ± 17.5 | 40.00 ± 17.5 | 52.86 ± 18.2 [‡] | 52.50 ± 17.3 | 54.38 ± 18.2 |
| Emotional well-being | 74.22 ± 10.2 | 75.11 ± 24.7 | 82.86 ± 15.4 | 82.29 ± 8.5 | 65.00 ± 18.8 | 73.00 ± 12.4 |
| Emotional role | 77.78 ± 44.1 | 92.59 ± 22.2 | 66.67 ± 47.1 | 90.49 ± 16.2 | 54.18 ± 46.9 | 100.0 ± 0 [‡] |
| Social function | 84.72 ± 16.2 | 90.28 ± 19.5 | 75.00 ± 20.4 | 96.43 ± 9.4 [‡] | 70.31 ± 25.8 | 85.94 ± 26.2 |
| Energy/fatigue | 69.44 ± 16.8 | 72.78 ± 22.6 | 71.43 ± 71.4 | 70.00 ± 9.5 | 62.50 ± 24.6 | 49.38 ± 14.0 |

Data are mean ± SD. ONS, oral nutritional supplementation; ONS + RE, oral nutritional supplementation plus resistance exercise; ONS + AE, oral nutritional supplementation plus aerobic exercise

p[‡]<0.05, Students T-test for comparison with the baseline measurement. Group x time interaction *p*=N.S

Adverse events

There were no adverse events associated with this trial