

Cocoa flavanol supplementation preserves early and late radial artery function after transradial catheterization

Running title: Cocoa flavanols preserve artery function

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Supplementary Information (SI):

SI Table 1:

| | Placebo (n = 18) | | | | Flavanol (n = 18) | | | |
|---|------------------|-------------|-------------|-------------|-------------------|-------------|-------------|-------------|
| | <i>baseline</i> | -2 h | +2 h | + 6 M | <i>baseline</i> | -2 h | +2 h | + 6 M |
| primary endpoint | | | | | | | | |
| intima-media thickness (mm) | 0.37 ± 0.01 | 0.36 ± 0.01 | 0.37 ± 0.01 | 0.44 ± 0.01 | 0.36 ± 0.01 | 0.36 ± 0.01 | 0.36 ± 0.01 | 0.37 ± 0.01 |
| intima-media thickness contralateral (mm) | 0.35 ± 0.01 | 0.36 ± 0.01 | 0.36 ± 0.01 | 0.35 ± 0.01 | 0.36 ± 0.01 | 0.36 ± 0.01 | 0.37 ± 0.01 | 0.37 ± 0.01 |
| secondary endpoint | | | | | | | | |
| FMD (%) | 8.4 ± 0.3 | 8.4 ± 0.5 | 4.3 ± 0.6 | 7.7 ± 0.4 | 9.2 ± 0.3 | 11.2 ± 0.5 | 5.9 ± 0.6 | 8.9 ± 0.4 |
| Fdc (%) | 2.2 ± 0.3 | 2.0 ± 0.3 | 1.3 ± 0.3 | 1.9 ± 0.3 | 1.8 ± 0.3 | 3.2 ± 0.3 | 1.7 ± 0.3 | 2.7 ± 0.3 |
| FMD contralateral (%) | 8.6 ± 0.3 | 8.5 ± 0.4 | 8.0 ± 0.4 | 8.2 ± 0.3 | 8.7 ± 0.3 | 10.9 ± 0.4 | 8.8 ± 0.4 | 8.9 ± 0.3 |
| tertiary endpoint | | | | | | | | |
| inner diameter (mm) | 2.02 ± 0.09 | 2.06 ± 0.09 | 2.25 ± 0.14 | 1.95 ± 0.10 | 1.83 ± 0.08 | 1.79 ± 0.07 | 1.96 ± 0.10 | 1.77 ± 0.08 |
| outer diameter (mm) | 2.76 ± 0.09 | 2.79 ± 0.08 | 2.98 ± 0.13 | 2.84 ± 0.10 | 2.56 ± 0.09 | 2.52 ± 0.08 | 2.68 ± 0.13 | 2.52 ± 0.10 |

SI Table 1: Vascular structural and functional parameters at four time points (baseline, -2 h, +2 h, and +6 M) for the flavanol and placebo group. Values are reported as mean ± SEM; (d = days; h = hours, M = months).

SI Table 2:

| quaternary endpoint | Placebo (n = 13) | | | Flavanol (n = 13) | | |
|---|------------------|--------------|--------------|-------------------|--------------|--------------|
| | Baseline | -2 h | +2 h | Baseline | -2 h | +2 h |
| EMP CD31 ⁺ 41 ⁻ (events/ μ l) | 595 \pm 28 | 585 \pm 26 | 675 \pm 27 | 607 \pm 28 | 465 \pm 26 | 628 \pm 27 |
| EMP CD144 ⁺ (events/ μ l) | 713 \pm 24 | 746 \pm 19 | 802 \pm 35 | 749 \pm 24 | 665 \pm 19 | 745 \pm 35 |
| EMP CD65E ⁺ (events/ μ l) | 618 \pm 21 | 601 \pm 25 | 687 \pm 26 | 611 \pm 21 | 539 \pm 25 | 651 \pm 26 |

SI Table 2: Endothelial microparticle concentrations at three time points (baseline, -2 h, and +2 h) for the flavanol and placebo group. Values are reported as mean \pm SEM; (d = days; h = hours).

SI Table 3:

| Parameter | Placebo | Cocoa flavanol |
|--|----------------|----------------|
| Daily serving size | 4 capsules/day | 4 capsules/day |
| Total cocoa flavanols (DP 1-7) ² (mg) | nd | 1,000 |
| Total flavanol monomers ³ (mg) | nd | 220 |
| (-)-epicatechin ³ (mg) | nd | 160 |
| (+)-catechin ³ (mg) | nd | 6.0 |
| (-)-catechin ³ (mg) | nd | 50 |
| Calories (kcal) | <5 | <5 |
| Total fat (g) | <0.2 | <0.2 |
| Saturated fat (g) | <0.15 | <0.15 |
| Total carbohydrates (g) | <0.5 | <0.5 |
| Sugars (g) | 0 | 0 |
| Fiber (g) | <0.5 | <0.5 |
| Protein (g) | <0.1 | <0.1 |
| Theobromine (mg) | 100 | 100 |
| Caffeine (mg) | 30 | 30 |

SI Table 3: Composition of daily cocoa flavanol and placebo interventions. The participants consumed 2 of capsules in the morning with breakfast and 2 at night with dinner. nd = not detected.

¹ Test material is not a significant source (≤ 1 mg/serving) of sodium, potassium, iron, magnesium, copper, manganese, phosphorous, or calcium.

² Analysis based on AOAC 2020.05. Cocoa flavanols includes flavanol monomers and procyanidins with a degree of polymerization (DP) up to 7 units.

³ Analysis based on AOAC official method. AOAC 2020.05-2020 Flavanol and Procyanidin (by Degree of Polymerization).

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