

FRUIT AND VEGETABLE INTAKE AND MORTALITY IN ADULTS UNDERGOING MAINTENANCE HEMODIALYSIS

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SUPPLEMENTARY MATERIAL LEGEND

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Supplemental Table 1. Combined intake of fruits and vegetables by country

| Country | Servings per week Median (interquartile range) |
|----------------|---|
| Turkey | 4.0 (1.5-8.0) |
| Poland | 6.0 (3.5-10.0) |
| Romania | 7.5 (4.0-11.0) |
| Spain | 7.5 (3.0-11.0) |
| Argentina | 8.0 (4.0-14.0) |
| Hungary | 8.0 (4.0-12.5) |
| France | 8.5 (5.5-14.0) |
| Germany | 8.5 (6.0-14.0) |
| Portugal | 10.0 (6.0-17.0) |
| Sweden | 10.0 (7.5-14.0) |
| Italy | 12.5 (7.0-17.0) |

Supplemental Table 2. Adjusted association (hazard ratio, 95% confidence interval) between baseline clinical characteristics and fruits and vegetables intake with mortality (clustered by country)

| | Cardiovascular mortality | P value | Non-cardiovascular mortality | P value | All-cause mortality | P value |
|--|---------------------------------|----------------|-------------------------------------|----------------|----------------------------|----------------|
| Fruits and vegetables intake (tertiles) | | 0.14 | | 0.01 | | 0.002 |
| 0 to 5.5 | 1.00 | | 1.00 | | 1.00 | |
| 5.6 to 10 | 0.95 (0.81-1.11) | | 0.88 (0.76-1.02) | | 0.90 (0.81-1.00) | |
| >10 | 0.84 (0.70-1.00) | | 0.77 (0.66-0.91) | | 0.80 (0.71-0.91) | |
| Age (1 year increase) | 1.02 (1.01-1.03) | <.001 | 1.02 (1.01-1.03) | <.001 | 1.02 (1.02-1.03) | <.001 |
| Male | 1.20 (1.04-1.39) | 0.01 | 1.22 (1.07-1.39) | <.001 | 1.21 (1.09-1.34) | 0.001 |
| Current or former smoker* | 1.27 (1.08-1.49) | <.001 | 1.06 (0.92-1.24) | 0.27 | 1.16 (1.04-1.30) | 0.03 |
| Daily physical activity* | 0.85 (0.68-1.06) | 0.23 | 0.91 (0.73-1.14) | 0.69 | 0.86 (0.73-1.01) | 0.11 |
| Energy intake (1000 kcal/day increase)* | 1.00 (0.93-1.07) | 0.89 | 1.00 (0.97-1.10) | 0.30 | 1.01 (0.96-1.06) | 0.66 |
| Life partner | - | | - | | 0.87 (0.78-0.98) | 0.02 |
| Secondary education | 0.82 (0.70-0.95) | 0.01 | - | | 0.88 (0.79-0.97) | 0.03 |
| Diabetes | 1.26 (1.07-1.50) | 0.01 | - | | | |
| Myocardial infarction | 1.57 (1.30-1.90) | <.001 | 0.94 (0.79-1.13) | 0.01 | 1.20 (1.06-1.37) | <.001 |
| Arteriovenous fistula | 0.78 (0.67-0.92) | <.001 | 0.59 (0.52-0.68) | <.001 | 0.68 (0.62-0.76) | <.001 |
| Body-mass index (kg/m²) | | <.001 | | <.001 | | <.001 |
| Underweight (<18.5) | 1.00 | | 1.00 | | 1.00 | |
| Normal range (18.5-24.9) | 0.75 (0.55-1.01) | | 0.63 (0.50-0.81) | | 0.66 (0.55-0.80) | |
| Pre-obese (25.0-29.9) | 0.54 (0.40-0.74) | | 0.47 (0.36-0.60) | | 0.48 (0.40-0.59) | |
| Obese (≥30.0) | 0.49 (0.35-0.68) | | 0.43 (0.32-0.56) | | 0.43 (0.34-0.53) | |
| Albumin tertiles (g/L) | | <.001 | | <.001 | | <.001 |
| Lowest (<38) | 1.00 | | 1.00 | | 1.00 | |
| Middle (38-41) | 0.66 (0.54-0.79) | | 0.60 (0.51-0.70) | | 0.62 (0.55-0.70) | |

| | Cardiovascular mortality | P value | Non-cardiovascular mortality | P value | All-cause mortality | P value |
|---|---------------------------------|----------------|-------------------------------------|----------------|----------------------------|----------------|
| Highest>41) | 0.56 (0.45-0.70) | | 0.45 (0.37-0.54) | | 0.49 (0.42-0.56) | |
| Charlson comorbidity score quartiles | | <.001 | | <.001 | | <.001 |
| 0-4 | 1.00 | | 1.00 | | 1.00 | |
| 43256 | 1.59 (1.11-2.26) | | 1.97 (1.35-2.87) | | 1.75 (1.35-2.27) | |
| 43319 | 2.20 (1.48-3.26) | | 2.67 (1.78-4.00) | | 2.40 (1.81-3.17) | |
| >8 | 3.26 (2.11-5.05) | | 4.57 (2.97-7.03) | | 3.90 (2.89-5.28) | |
| Hemoglobin (g/dL) | 0.92 (0.87-0.97) | 0.001 | 0.91 (0.87-0.96) | <.001 | 0.91 (0.88-0.94) | <.001 |
| Phosphorus (mg/dL) | 1.08 (1.03-1.14) | 0.002 | - | | 1.05 (1.01-1.09) | 0.01 |
| Calcium, (mg/dL) | 1.21 (1.10-1.33) | 0.001 | - | | 1.15 (1.07-1.22) | <.001 |
| Primary renal diagnosis | | | - | <.001 | | <.001 |
| Diabetes | - | | 1.00 | | 1.00 | |
| Hypertension | - | | 0.95 (0.79-1.14) | | 0.90 (0.79-1.03) | |
| Glomerulonephritis | - | | 0.85 (0.66-1.09) | | 0.81 (0.68-0.98) | |
| Other | - | | 0.97 (0.82-1.13) | | 0.84 (0.75-0.95) | |
| Wait-listed for transplant | - | | 0.57 (0.43-0.75) | <.001 | 0.76 (0.63-0.91) | 0.01 |
| Time on dialysis (year) | - | | - | | 1.01 (1.00-1.02) | 0.04 |
| Kt/V | - | | - | | 0.83 (0.72-0.97) | 0.02 |

*These were clinically relevant variables specified *a priori*. The remaining variables included in each model were selected by backward elimination because they significantly predicted mortality ($p < 0.05$) or had a clinically meaningful impact on the hazard ratio for mortality ($\geq 10\%$). Other variables tested as potential confounders were: race, occupation, hypertension, stroke, grams per day of nutrients (monounsaturated fatty acids, saturated fatty acids, *trans* fatty acids, n-3 long chain polyunsaturated fatty acids, n-6 polyunsaturated fatty acids, cereal fiber, protein, alcohol, and sugar), servings per week of foods (legumes, cereals, dairy, white meat, red and processed meat, fish, olive oil, sweets and sweet drinks), and Mediterranean diet score.

Supplemental Table 3. Adjusted mortality hazard ratio (95% confidence interval) by fruits and vegetables intake (tertiles of serving per week) and plant-based protein intake (one gram/day increase)

| | ^a Cardiovascular mortality | P value | ^b Non-cardiovascular mortality | P value | ^c All-cause mortality | P value |
|-------------------------------------|---------------------------------------|---------|---|-------------|----------------------------------|-------------|
| Fruits and vegetables intake | | | | | | |
| 0 to 5.5 | 1.00 | 0.13 | 1.00 | 0.04 | 1.00 | 0.01 |
| 5.6 to 10 | 0.94 (0.81-1.11) | | 0.89 (0.77-1.04) | | 0.91 (0.81-1.01) | |
| >10 | 0.83 (0.69-1.00) | | 0.81 (0.68-0.95) | | 0.81 (0.72-0.92) | |
| Plant based protein intake | 1.00 (0.99-1.02) | 0.65 | 0.98 (0.97-1.00) | 0.02 | 0.99 (0.98-1.00) | 0.21 |
| Fruits intake | | | | | | |
| 0 to 1 | 1.00 | 0.97 | 1.00 | 0.08 | 1.00 | 0.30 |
| 1.1 to 6 | 1.01 (0.84-1.21) | | 0.87 (0.73-1.03) | | 0.92 (0.81-1.05) | |
| >6 | 0.99 (0.83-1.19) | | 0.84 (0.71-0.98) | | 0.91 (0.81-1.03) | |
| Plant based protein intake | 1.00 (0.99-1.02) | 0.91 | 0.98 (0.97-1.00) | 0.01 | 0.99 (0.98-1.00) | 0.11 |
| Vegetables intake | | | | | | |
| 0 to 1 | 1.00 | 0.57 | 1.00 | 0.70 | 1.00 | 0.36 |
| 1.1 to 3 | 1.02 (0.86-1.21) | | 1.01 (0.86-1.17) | | 0.99 (0.88-1.11) | |
| >3 | 0.93 (0.78-1.12) | | 0.95 (0.80-1.11) | | 0.92 (0.82-1.04) | |
| Plant based protein intake | 1.00 (0.99-1.02) | 0.91 | 0.98 (0.97-1.00) | 0.01 | 0.99 (0.98-1.00) | 0.11 |

^aAdjusted for country (fixed effect in competing risk analysis and random effect in the others), gender, smoking (current or former versus never), daily physical activity, life partner, education (secondary versus none/primary), primary renal diagnosis, myocardial infarction, vascular access type (fistula versus graft/catheter), body mass index (categories according to WHO), albumin (tertiles), Charlson comorbidity score (quartiles), wait-listed for transplantation, age, phosphorus, calcium, hemoglobin, Kt/V, time on dialysis, and total energy intake (1000 kcal day increase). Analyses for fruits are adjusted for vegetables and vice versa.

Supplemental Table 4. All-cause mortality hazard ratios (95% confidence interval) for fruits and vegetables intake

| Servings per week | Univariable random effect | P value | *Multivariable competing risk, fixed effect | P value | *Multivariable case-complete, random effect | P value |
|------------------------------|---------------------------|-------------|---|--------------|---|------------------|
| Fruits and vegetables | | | | | | |
| 1 serving increase | 0.99 (0.99-1.00) | 0.01 | 0.99 (0.98-0.99) | 0.001 | 0.98 (0.97-0.99) | <0.001 |
| Tertiles | | | | | | |
| 0 to 5.5 | 1.00 | 0.03 | 1.00 | 0.003 | 1.00 | <.0001 |
| 5.6 to 10 | 0.93 (0.84-1.03) | | 0.91 (0.82-1.02) | | 0.79 (0.67-0.94) | |
| >10 | 0.86 (0.77-0.96) | | 0.81 (0.71-0.91) | | 0.65 (0.54-0.78) | |
| Fruits | | | | | | |
| 1 serving increase | 0.99 (0.98-1.00) | 0.09 | 0.99 (0.98-1.00) | 0.02 | 0.97 (0.96-0.99) | <0.001 |
| Tertiles | | | | | | |
| 0 to 1 | 1.00 | 0.22 | 1.00 | 0.30 | 1.00 | 0.04 |
| 1.1 to 6 | 0.91 (0.81-1.02) | | 0.93 (0.82-1.05) | | 0.86 (0.70-1.05) | |
| >6 | 0.93 (0.83-1.03) | | 0.91 (0.81-1.03) | | 0.80 (0.67-0.95) | |
| Vegetables | | | | | | |
| 1 serving increase | 0.99 (0.97-1.00) | 0.02 | 0.99 (0.98-1.00) | 0.11 | 0.98 (0.96-1.00) | 0.10 |
| Tertiles | | | | | | |
| 0 to 1 | 1.00 | 0.16 | 1.00 | 0.23 | 1.00 | 0.17 |
| 1.1 to 3 | 1.02 (0.91-1.13) | | 0.98 (0.88-1.10) | | 0.94 (0.79-1.12) | |
| >3 | 0.92 (0.83-1.03) | | 0.91 (0.80-1.02) | | 0.84 (0.70-1.01) | |

* Adjusted for country (fixed effect in competing risk analysis and random effect in the others), gender, smoking (current or former versus never), daily physical activity, life partner, education (secondary versus none/primary), primary renal diagnosis, myocardial infarction, vascular access type (fistula versus graft/catheter), body mass index (categories according to WHO), albumin (tertiles), Charlson comorbidity score (quartiles), wait-listed for transplantation, age, phosphorus, calcium, hemoglobin, Kt/V, time on dialysis, and total energy intake (1000 kcal day increase). Analyses for fruits are adjusted for vegetables and vice versa.

Supplemental Table 5. Cardiovascular mortality hazard ratios (95% confidence interval) for fruits and vegetables intake

| Servings per week | Univariate random effect | P value | *Multivariable competing risk, fixed effect | P value | *Multivariable case-complete, random effect | P value |
|------------------------------|---------------------------------|----------------|--|----------------|--|----------------|
| Fruits and vegetables | | | | | | |
| 1 serving increase | 0.99 (0.98-1.00) | 0.06 | 0.99 (0.98-1.00) | 0.13 | 0.98 (0.97-1.00) | 0.02 |
| Tertiles | | | | | | |
| 0 to 5.5 | 1.00 | 0.15 | 1.00 | 0.31 | 1.00 | 0.06 |
| 5.6 to 10 | 0.92 (0.79-1.07) | | 0.96 (0.81-1.12) | | 0.85 (0.66-1.10) | |
| >10 | 0.85 (0.72-1.00) | | 0.87 (0.73-1.04) | | 0.72 (0.54-0.94) | |
| Fruits | | | | | | |
| 1 serving increase | 0.99 (0.98-1.01) | 0.18 | 0.99 (0.97-1.01) | 0.22 | 0.98 (0.95-1.00) | 0.04 |
| Tertiles | | | | | | |
| 0 to 1 | 1.00 | 0.77 | 1.00 | 0.98 | 1.00 | 0.65 |
| 1.1 to 6 | 0.95 (0.80-1.12) | | 1.02 (0.85-1.22) | | 0.93 (0.69-1.26) | |
| >6 | 0.95 (0.81-1.11) | | 1.01 (0.85-1.21) | | 0.88 (0.66-1.16) | |
| Vegetables | | | | | | |
| 1 serving increase | 0.99 (0.97-1.01) | 0.18 | 1.00 (0.98-1.02) | 0.78 | 0.99 (0.96-1.02) | 0.53 |
| Tertiles | | | | | | |
| 0 to 1 | 1.00 | 0.45 | 1.00 | 0.71 | 1.00 | 0.39 |
| 1.1 to 3 | 1.04 (0.88-1.21) | | 1.02 (0.86-1.21) | | 1.04 (0.80-1.36) | |
| >3 | 0.94 (0.80-1.10) | | 0.95 (0.80-1.13) | | 0.87 (0.65-1.15) | |

* Adjusted for country (fixed effect in competing risk analysis and random effect in the others) gender, smoking (current or former versus never), daily physical activity, education (secondary versus none/primary), diabetes, myocardial infarction, vascular access type (fistula versus graft/catheter), body mass index (categories according to WHO), albumin (tertiles), Charlson comorbidity score (quartiles), age, phosphorus, calcium, hemoglobin, and total energy intake (1000 kcal day increase). Analyses for fruits are adjusted for vegetables and vice versa.

Supplemental Table 6. Non-cardiovascular mortality hazard ratios (95% confidence interval) for fruits and vegetables intake

| Servings per week | Univariable random effect | P value | *Multivariable competing risk, fixed effect | P value | *Multivariable case-complete, random effect | P value |
|------------------------------|---------------------------|---------|---|-------------|---|--------------|
| Fruits and vegetables | | | | | | |
| 1 serving increase | 0.99 (0.98-1.00) | 0.05 | 0.99 (0.98-1.00) | 0.01 | 0.98 (0.97-0.99) | 0.001 |
| Teriles | | | | | | |
| 0 to 5.5 | 1.00 | 0.20 | 1.00 | 0.03 | 1.00 | 0.001 |
| 5.6 to 10 | 0.93 (0.81-1.07) | | 0.92 (0.79-1.07) | | 0.81 (0.67-0.99) | |
| >10 | 0.87 (0.75-1.01) | | 0.80 (0.68-0.95) | | 0.67 (0.54-0.82) | |
| Fruits | | | | | | |
| 1 serving increase | 0.99 (0.98-1.01) | 0.30 | 0.99 (0.97-1.00) | 0.08 | 0.98 (0.96-1.00) | 0.01 |
| Teriles | | | | | | |
| 0 to 1 | 1.00 | 0.22 | 1.00 | 0.12 | 1.00 | 0.07 |
| 1.1 to 6 | 0.87 (0.74-1.03) | | 0.90 (0.76-1.07) | | 0.87 (0.69-1.09) | |
| >6 | 0.91 (0.79-1.05) | | 0.84 (0.72-0.99) | | 0.79 (0.64-0.97) | |
| Vegetables | | | | | | |
| 1 serving increase | 0.98 (0.97-1.00) | 0.05 | 0.99 (0.97-1.00) | 0.11 | 0.98 (0.96-1.01) | 0.19 |
| Teriles | | | | | | |
| 0 to 1 | 1.00 | 0.33 | 1.00 | 0.54 | 1.00 | 0.30 |
| 1.1 to 3 | 1.00 (0.87-1.16) | | 0.97 (0.83-1.13) | | 0.97 (0.80-1.19) | |
| >3 | 0.91 (0.79-1.05) | | 0.91 (0.77-1.08) | | 0.86 (0.69-1.06) | |

* Adjusted for country (fixed effect in competing risk analysis and random effect in the others), gender, smoking (current or former versus never), daily physical activity, primary renal diagnosis, myocardial infarction, vascular access type (fistula versus graft/catheter), body mass index (categories according to WHO), albumin (teriles), Charlston comorbidity score (quartiles), wait-listed for transplantation, age, hemoglobin, and total energy intake (1000 kcal day increase). Analyses for fruits are adjusted for vegetables and vice versa.