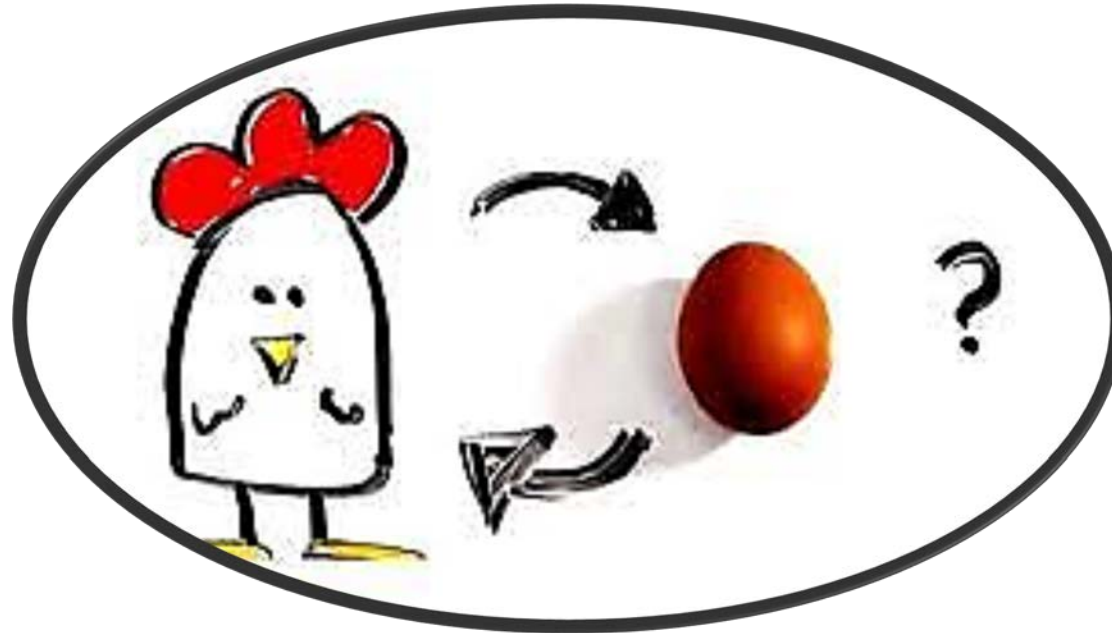


UROLITIASIS Y OSTEOPOROSIS ¿CUÁL ES LA RELACIÓN?



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Unidad de Investigación Epidemiológica, FALP

Unidad de Endocrinología

DECLARACION DE CONFLICTOS DE INTERES

No tengo conflictos de interés en relación al contenido de esta exposición

EPIDEMIOLOGÍA DE LA UROLITIASIS

| Sex | Prevalence | Incidence |
|-------|------------|------------------------------|
| Men | 10.6% | 140.6 per 100,000 population |
| Women | 7.1% | 65.8 per 100,000 population |

| Year following initial stone event | Risk of recurrence |
|------------------------------------|--------------------|
| 2 | 11% |
| 5 | 20% |
| 10 | 31% |
| 15 | 39% |

Ziemba, J.B. *Investig. Clin. Urol.* 2017, 58, 299–306

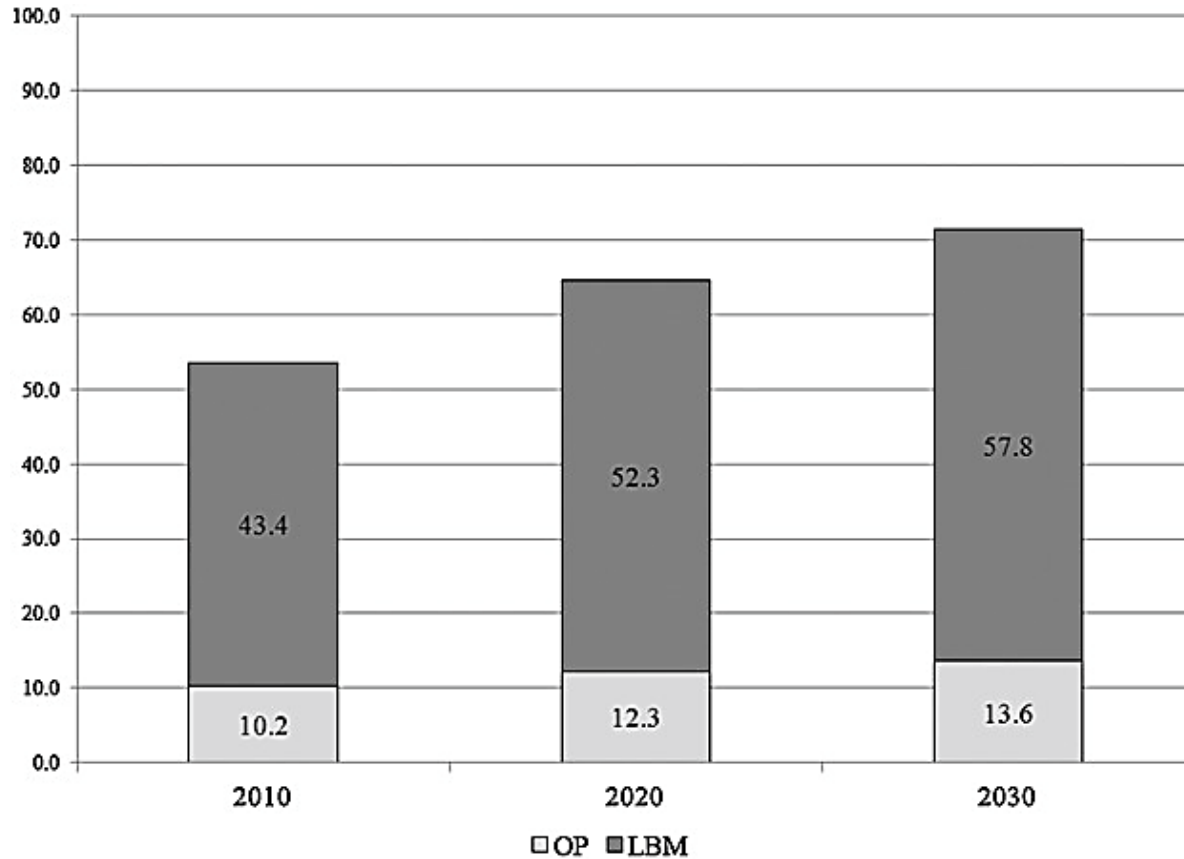
Scales CD Jr. *Eur Urol* 2012;62:160-5

| Characteristic | Odds ratio (95% CI) | <i>p</i> value |
|----------------------|---------------------|----------------|
| Age, yr | | |
| 20–39 | 1.00 (referent) | – |
| 40–59 | 1.83 (1.37–2.45) | <0.001 |
| ≥60 | 2.18 (1.74–2.73) | <0.001 |
| Female | 0.63 (0.52–0.75) | <0.001 |
| Race | | |
| White, non-Hispanic | 1.00 (referent) | – |
| Black, non-Hispanic | 0.37 (0.28–0.49) | <0.001 |
| Hispanic | 0.60 (0.49–0.73) | <0.001 |
| Other/multiracial | 0.57 (0.37–0.89) | 0.014 |
| BMI category | | |
| Normal | 1.00 (referent) | – |
| Overweight | 1.29 (0.96–1.72) | 0.0875 |
| Obese | 1.55 (1.25–1.94) | <0.001 |
| Household income, \$ | | |
| ≥75 000 | 1.00 (referent) | – |
| 35 000–74 999 | 1.49 (1.16–1.92) | 0.002 |
| 20 000–34 999 | 1.65 (1.27–2.15) | <0.001 |
| 0–19 999 | 1.57 (1.17–2.09) | 0.002 |
| Diabetes | 1.59 (1.22–2.07) | <0.001 |
| Gout | 1.92 (1.44–2.56) | <0.001 |

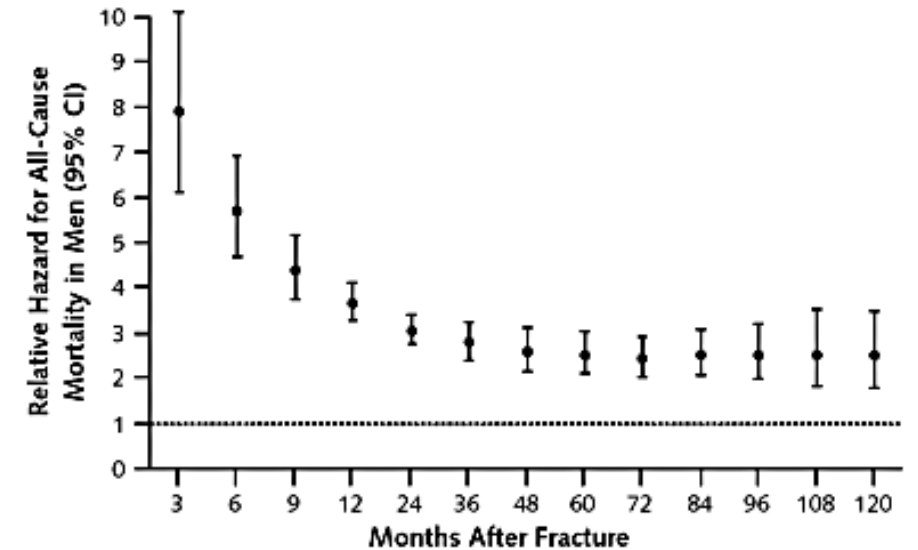
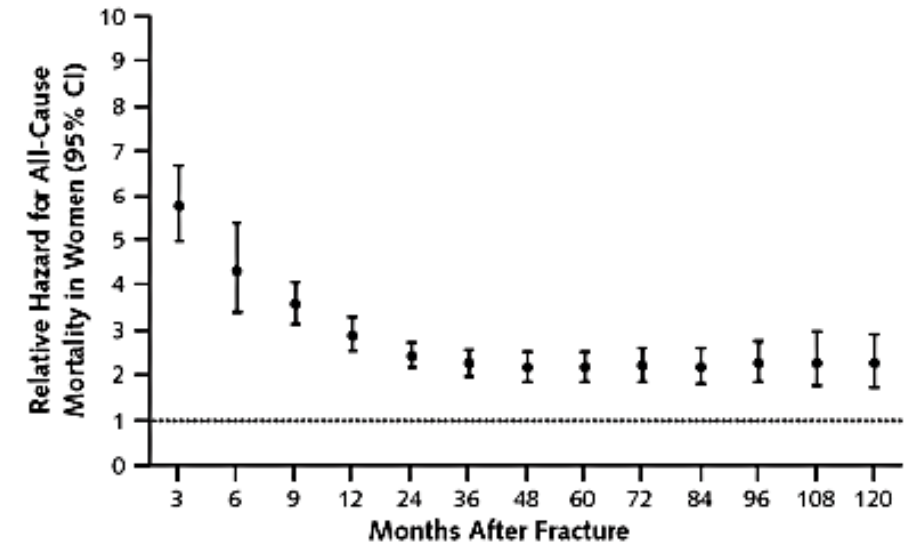
CI = confidence interval; BMI = body mass index.

EPIDEMIOLOGÍA DE LA OSTEOPOROSIS

Proyección de la prevalencia (%) de osteoporosis (OP) y masa ósea baja (LBM), Medida por DPX en CL o CF, en mayores de 50 años no institucionalizados de EEUU

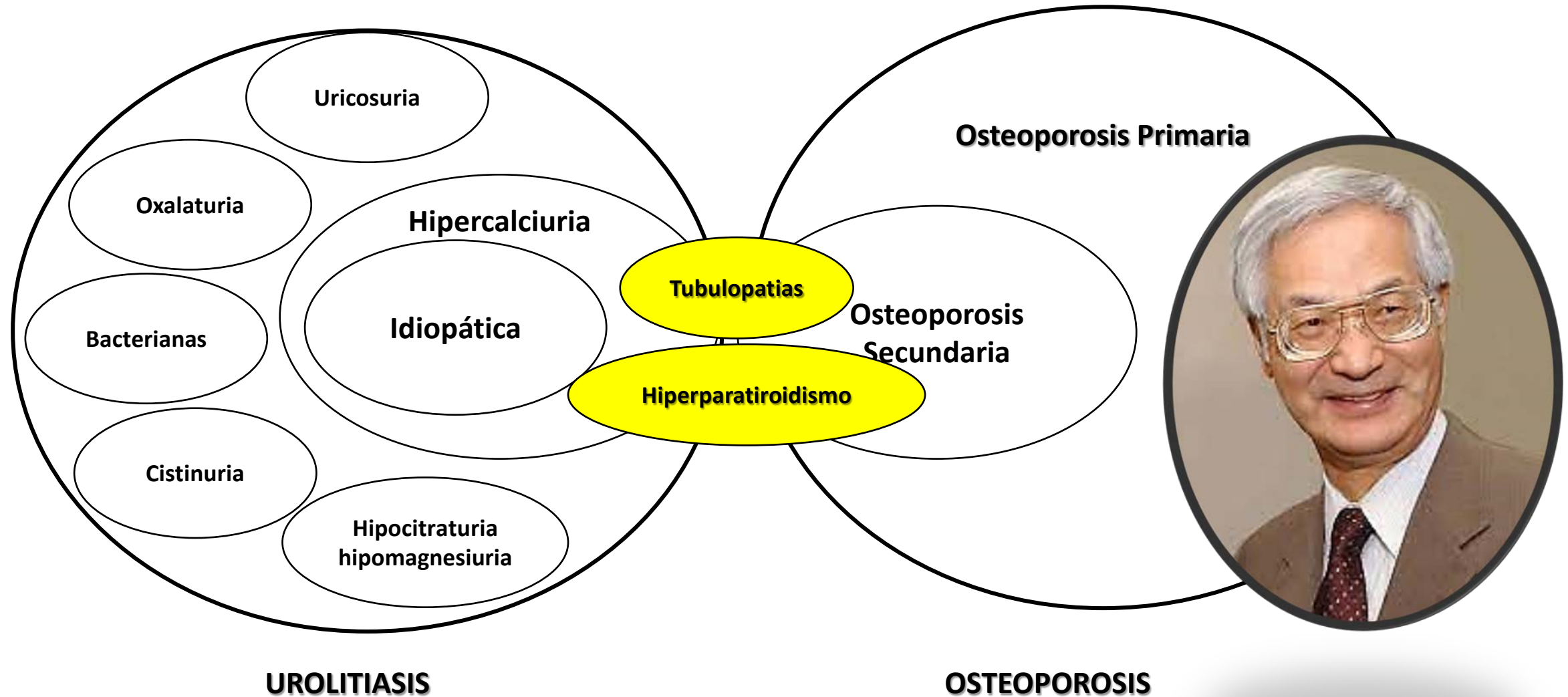


Wright NC. J Bone Miner Res. 2014. doi:10.1002/jbmr.2269



Haentjens P. Ann Intern Med. 2010; 152: 380-390

UROLITIASIS Y OSTEOPOROSIS ¿CUÁL ES LA RELACIÓN?



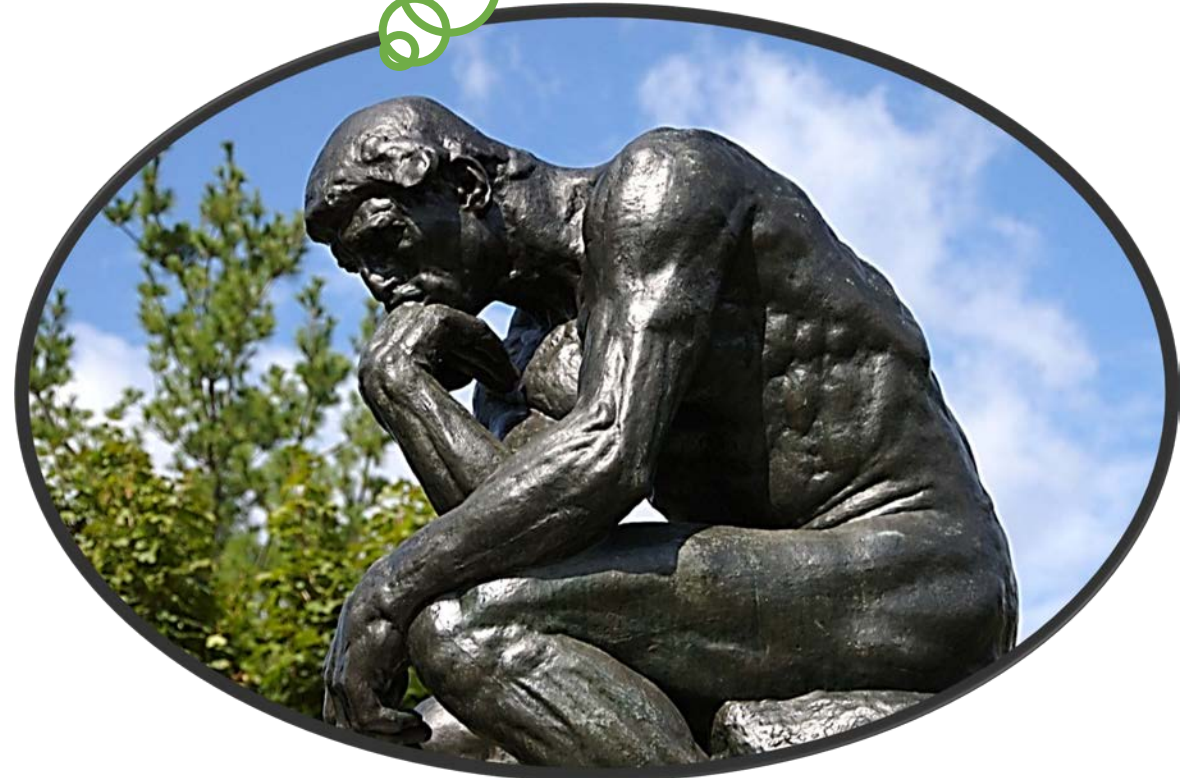
Características de los residentes de Rochester, Minnesota Con urolitiasis sintomática

| | <i>N</i> | <i>%</i> |
|---------------------------------|----------|----------|
| Type of stone | | |
| Calcium oxalate | 158 | 25.3 |
| Calcium phosphate | 15 | 2.4 |
| Mixed | 45 | 7.2 |
| Urate | 8 | 1.3 |
| Other | 1 | 0.2 |
| Unknown | 397 | 63.6 |
| Clinical diagnosis ^a | | |
| Hyperuricemia/gout | 146 | 23.4 |
| Idiopathic hypercalciuria | 47 | 7.5 |
| Hyperparathyroidism | 13 | 2.1 |
| Small bowel resection | 9 | 1.4 |
| Inflammatory bowel disease | 4 | 0.6 |
| Sarcoidosis | 1 | 0.2 |
| Uncertain | 428 | 68.6 |

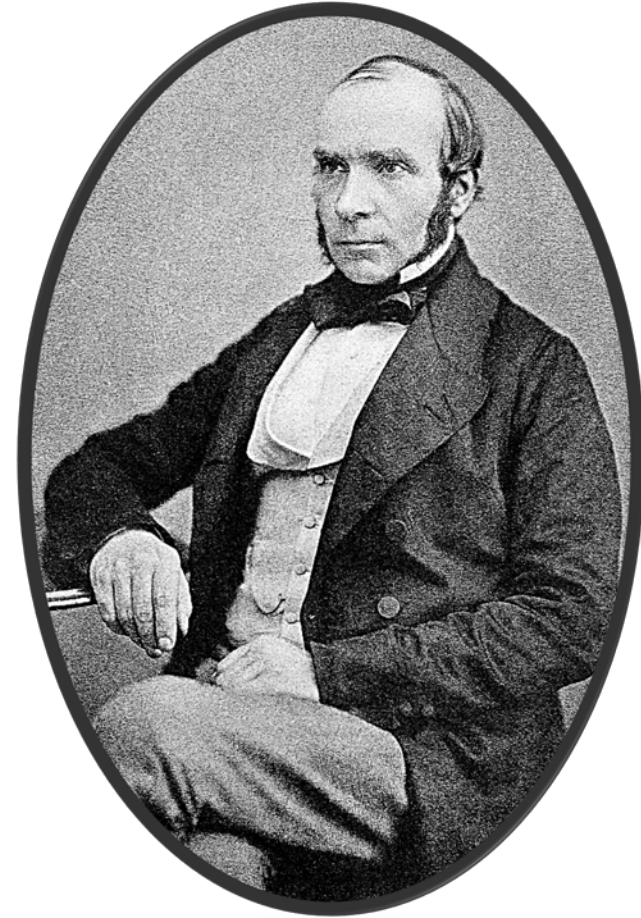
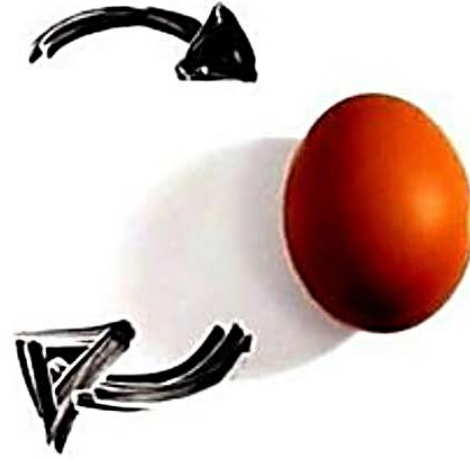
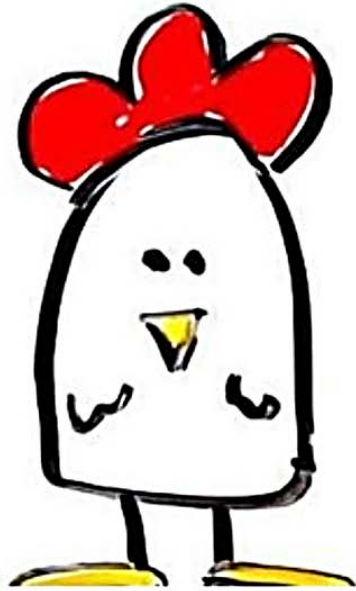
^a More than one diagnosis may have been reported

Melton LJ. *Kidney Int.* 1998;53(2):459–64

**...Hiperparatiroidismo
primario, tubulopatías...**



ASOCIACIÓN ENTRE UROLITIASIS Y OSTEOPOROSIS



ASOCIACIÓN ENTRE UROLITIASIS Y DMO

| Analysis | Number of studies | Number participants | | Meta-analysis | | | Heterogeneity I^2 |
|---------------------|-------------------|---------------------|------|---------------|---------|----------------|---------------------|
| | | Nephrolithiasis | HC | SMD | 95 % CI | <i>p</i> value | |
| Lumbar spine | | | | | | | |
| BMD | 15 | 1037 | 2584 | -0.58 | -0.91 | -0.25 | 0.0001 |
| T-score | 7 | 390 | 282 | -0.67 | -1.00 | -0.34 | <0.0001 |
| Z-score | 11 | 421 | | -0.41 | -0.85 | 0.03 | 0.07 |
| Total hip | | | | | | | |
| BMD | 15 | 1037 | 2584 | -0.52 | -0.82 | -0.22 | 0.04 |
| T-score | 7 | 390 | 282 | -0.62 | -1.11 | -0.52 | <0.0001 |
| Z-score | 11 | 421 | 182 | -0.82 | -1.10 | -0.54 | <0.0001 |
| Forearm | | | | | | | |
| BMD | 8 | 523 | 537 | -0.35 | -0.66 | -0.03 | 0.03 |
| T-score | 6 | 360 | 252 | -0.67 | -1.00 | -0.34 | <0.0001 |
| Z-score | 8 | 347 | 380 | -0.41 | -0.85 | 0.03 | 0.07 |

**24 estudios de casos y controles.
1595 pacientes con nefrolitiasis y 3402 controles sanos.
Sólo se excluyó HPP**

ASOCIACIÓN ENTRE UROLITIASIS Y FRACTURAS

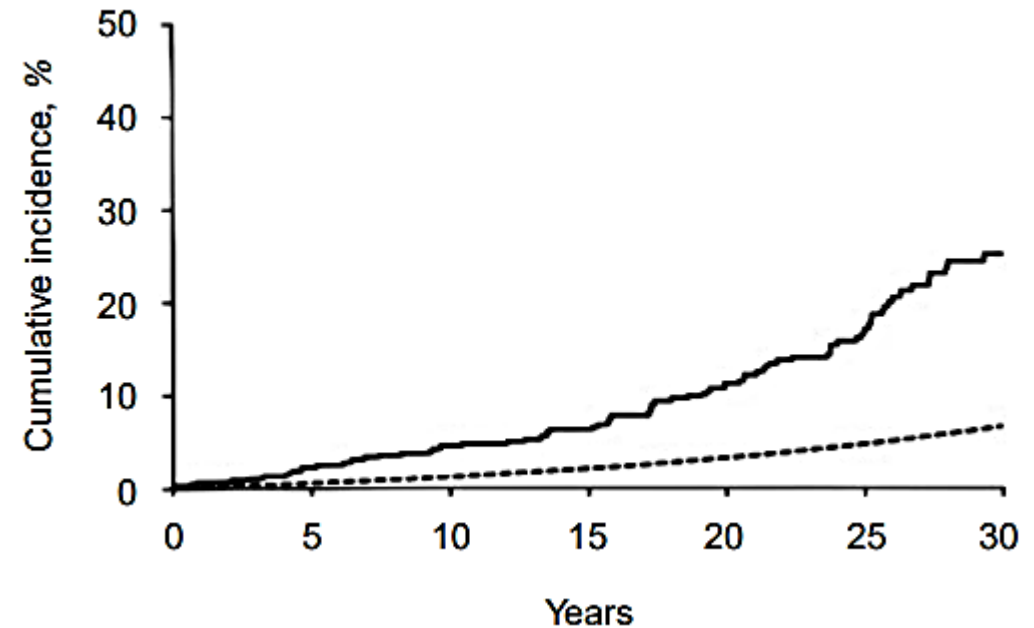
| <i>Fracture site</i> | <i>Men</i> | <i>Women</i> |
|----------------------|------------------|------------------|
| Wrist | 1.68 (1.02–2.76) | 1.13 (0.73–1.76) |
| Spine | 2.32 (1.04–5.18) | 1.75 (0.58–5.29) |

OR de prevalencia e IC 95% para fracturas de columna y muñeca en personas con antecedentes de urolitiasis en relación al control, ajustado por edad y raza/etnicidad (NHANES III)

Lauderdale DS. J Bone Miner Res. 2001;16(10):1893–8

| Clinical diagnosis ^a | | |
|---------------------------------|-----|------|
| Hyperuricemia/gout | 146 | 23.4 |
| Idiopathic hypercalciuria | 47 | 7.5 |
| Hyperparathyroidism | 13 | 2.1 |
| Small bowel resection | 9 | 1.4 |
| Inflammatory bowel disease | 4 | 0.6 |
| Sarcoidosis | 1 | 0.2 |
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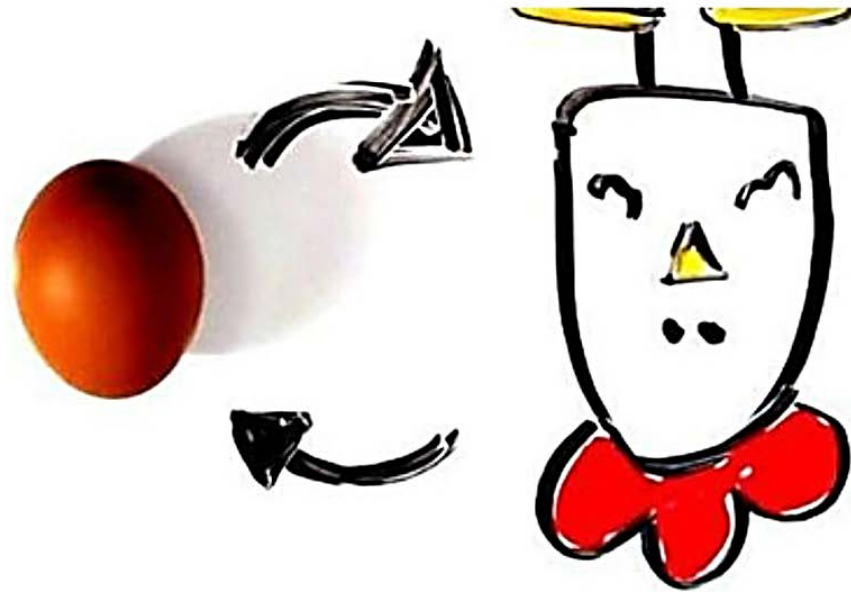
^a More than one diagnosis may have been reported



Incidencia acumulada de fracturas vertebrales observada (línea continua) y esperada (línea discontinua) entre los residentes de Rochester, Minnesota después del episodio inicial de urolitiasis sintomática

Melton LJ. Kidney Int. 1998;53(2):459–64

ASOCIACIÓN ENTRE OSTEOPOROSIS Y UROLITIASIS

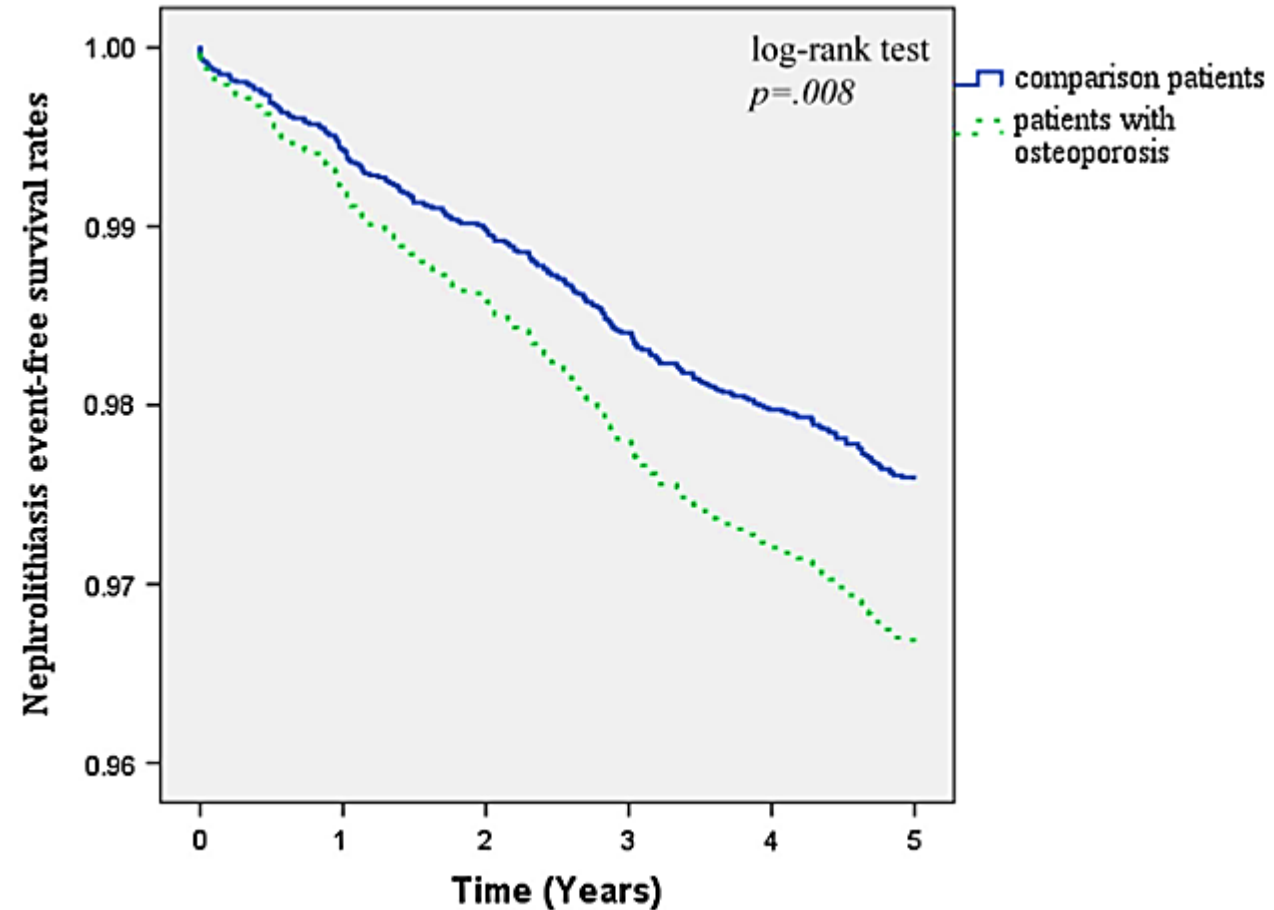


ASOCIACIÓN ENTRE OSTEOPOROSIS Y UROLITIASIS

HR de nefrolitiasis sintomática en pacientes con osteoporosis durante el período de seguimiento de 5 años

| Development of symptomatic nephrolithiasis | Total | | Patients with osteoporosis | | Patients without osteoporosis | |
|--|-------|------|----------------------------|------|-------------------------------|------|
| | No. | (%) | No. | (%) | No. | (%) |
| 5-year follow-up period | | | | | | |
| Yes | 225 | 2.8 | 60 | 3.7 | 165 | 2.5 |
| No | 7945 | 97.2 | 1574 | 96.3 | 6371 | 97.5 |
| Crude HR (95 % CI) | | | 1.48 (1.10–1.99)** | | 1 | |
| Adjusted HR (95 % CI) | | | 1.38 (1.03–1.86)* | | 1 | |

Total sample number = 8170



Chou, P.S. Calcif. Tissue Int. 2014, 95, 317– 322

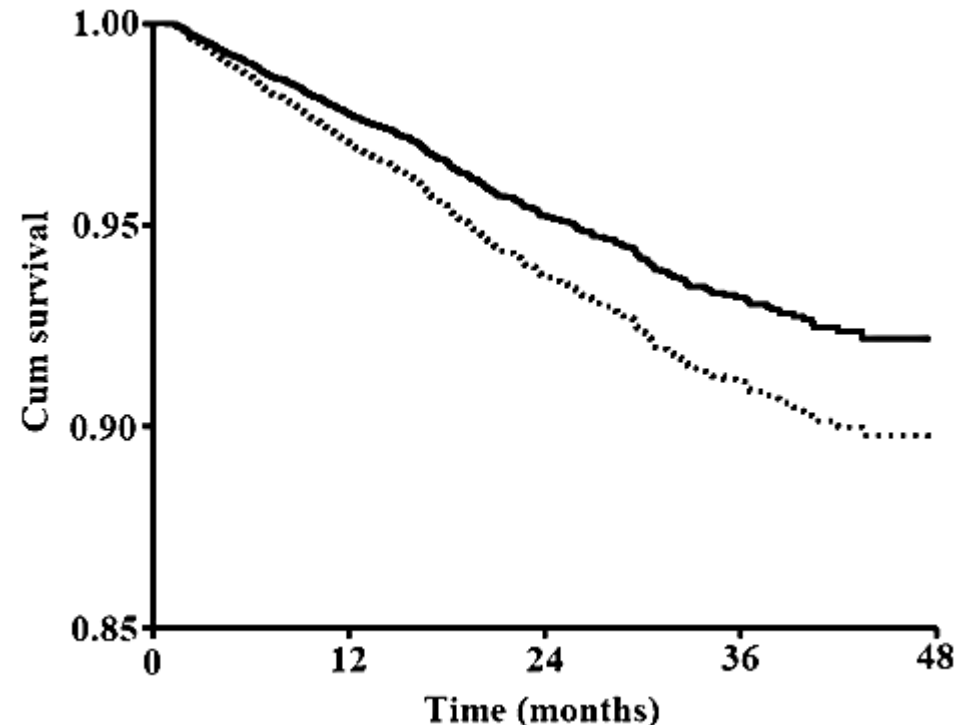
ASOCIACIÓN ENTRE OSTEOPOROSIS Y UROLITIASIS

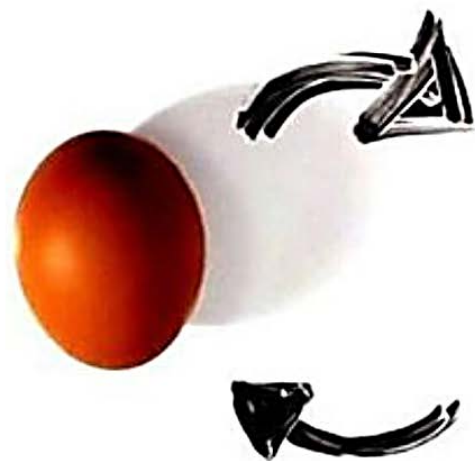
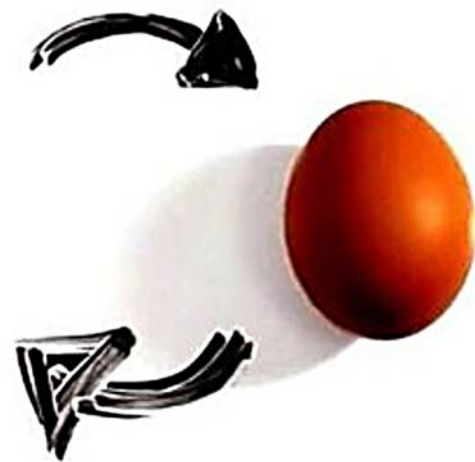
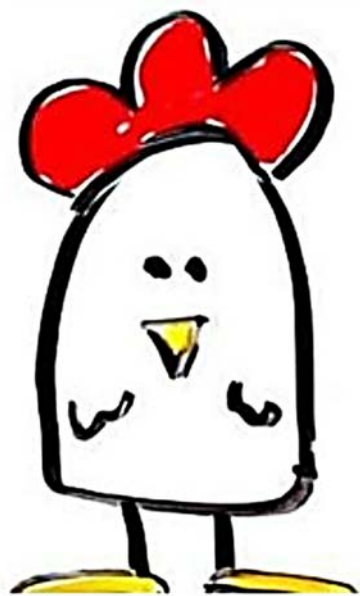
Factores de riesgo para la urolitiasis incidente en sujetos del sur de Italia mayores de 40 años

| | <i>p</i> | HR | 95% CI | |
|------------------------------|----------|------|--------|------|
| Osteoporosis diagnosis (y/n) | 0.04 | 1.33 | 1.01 | 1.74 |
| Age (years) | 0.02 | 0.99 | 0.98 | 1.00 |
| Sex (m/f) | 0.52 | 0.87 | 0.56 | 1.34 |
| BMI (Kg/m ²) | 0.22 | 0.99 | 0.97 | 1.01 |
| Osteoporosis treatment (y/n) | 0.39 | 1.15 | 0.84 | 1.56 |
| Current smokers (y/n) | 0.18 | 1.17 | 0.93 | 1.48 |
| Previous smokers (y/n) | 0.68 | 1.08 | 0.73 | 1.60 |

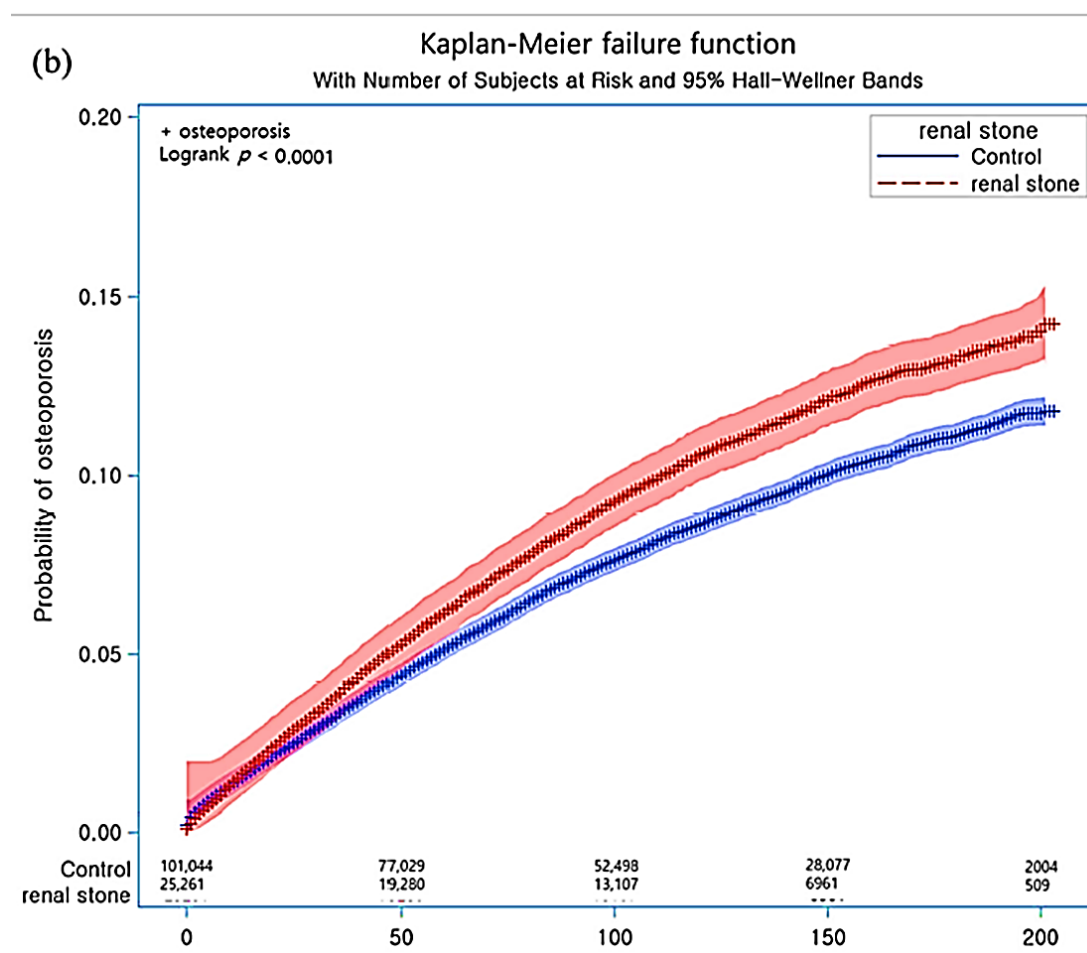
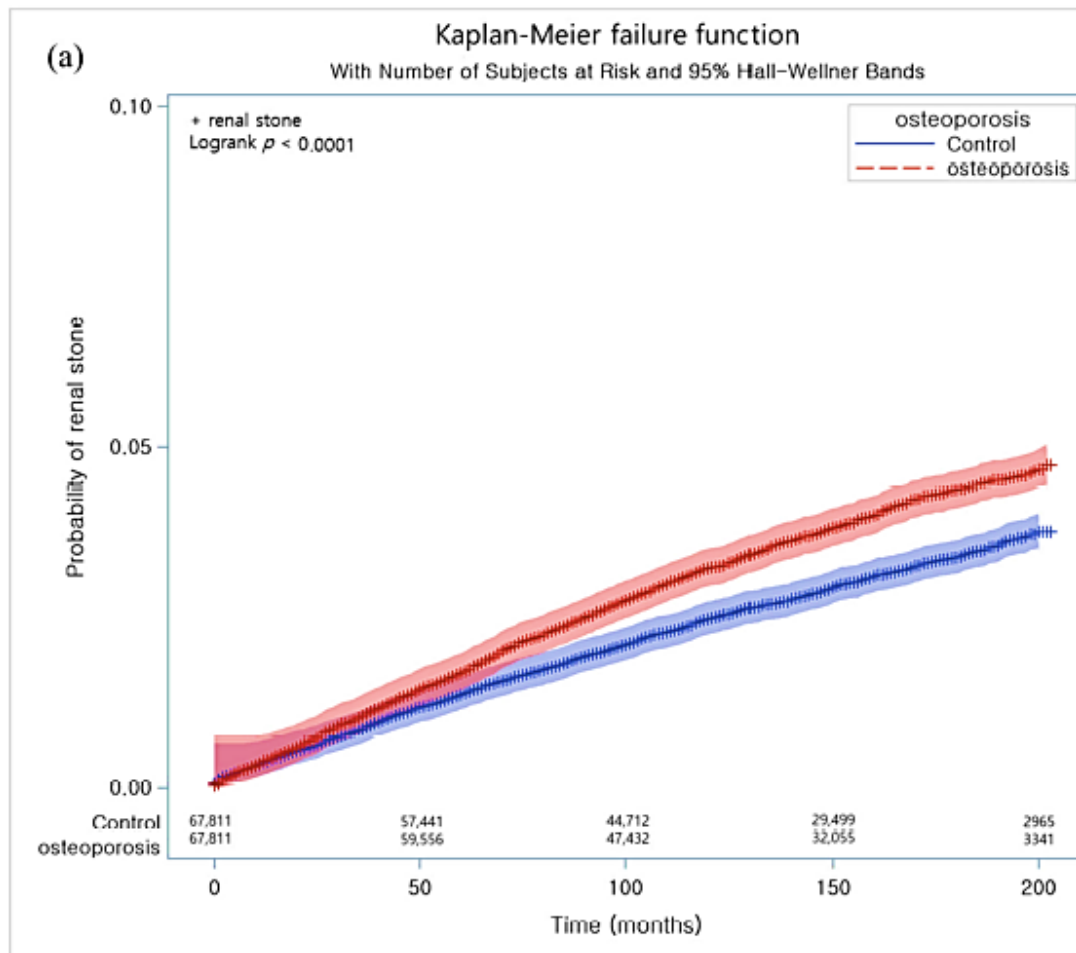
*Se excluyó HPP

Probabilidad dependiente del tiempo de aparición de nefrolitiasis. La línea punteada y la línea continua indican pacientes con y sin osteoporosis





RELACIÓN RECÍPROCA ENTRE LA OSTEOPOROSIS Y LOS CÁLCULOS RENALES



So Young Kim. J. Clin. Med. 2022, 11,6614.<https://doi.org/10.3390/jcm11226614>



HIPERCALCIURIA Y DMO

| Outcome | Number comparisons | β | 95 % CI | | <i>p</i> value | R^2 |
|-------------------|--|----------------|---------------|----------------|----------------|-------------|
| Lumbar BMD | 7 | 0.002 | -0.005 | 0.009 | 0.63 | 0.00 |
| Lumbar T-score | 5 | 0.0002 | -0.003 | 0.003 | 0.90 | 0.00 |
| Lumbar Z-score | 7 | 0.002 | -0.009 | 0.14 | 0.68 | 0.00 |
| Total hip BMD | Only 2 studies | | | | | |
| Total hip T-score | 4 | -0.006 | -0.02 | 0.01 | 0.49 | 0.00 |
| Total hip Z-score | 4 | -0.006 | -0.02 | 0.01 | 0.49 | 0.00 |
| Neck BMD | Only 3 studies with data about calciuria | | | | | |
| Neck T-score | 4 | -0.0003 | -0.005 | -0.0001 | 0.05 | 1.00 |
| Neck Z-score | 5 | -0.006 | -0.010 | -0.002 | 0.04 | 0.41 |

Bold values represent significant results (*p*-values <0.05)

β unstandardized regression coefficient, *CI* confidence interval, *BMD* bone mineral density

Asociación entre la diferencia en calcio en orina de 24 h (calculado como nefrolitiasis menos controles) y estimaciones de salud ósea. Se excluyó HPP

Lucato, P. Osteoporos. Int. 2016, 27, 3155–3164

UROLITIASIS NORMOCALCIURICA Y OSTEOPOROSIS

| BMD (gr/cm ²) | | | |
|---------------------------|---------------------------|---------------------------|----------|
| | Group 1 (BMD) | Group 2 (BMD) | <i>P</i> |
| Total femur | 0.897 ± 0.132 | 0.945 ± 0.170 | <0.05 |
| Femur neck | 0.759 ± 0.132 | 0.887 ± 0.135 | <0.001 |
| Lumbal Spine (L2-L4) | 0.874 ± 0.130 | 1.021 ± 0.301 | <0.001 |
| <i>t</i> score | | | |
| | Group 1 (<i>t</i> score) | Group 2 (<i>t</i> score) | <i>P</i> |
| Total femur | -0.795 ± 0.689 | -0.588 ± 2.416 | >0.05 |
| Femur neck | -0.883 ± 0.749 | 0.435 ± 0.547 | <0.001 |
| Lumbal spine (L2-L4) | -1.509 ± 1.224 | -0.225 ± 0.854 | <0.001 |

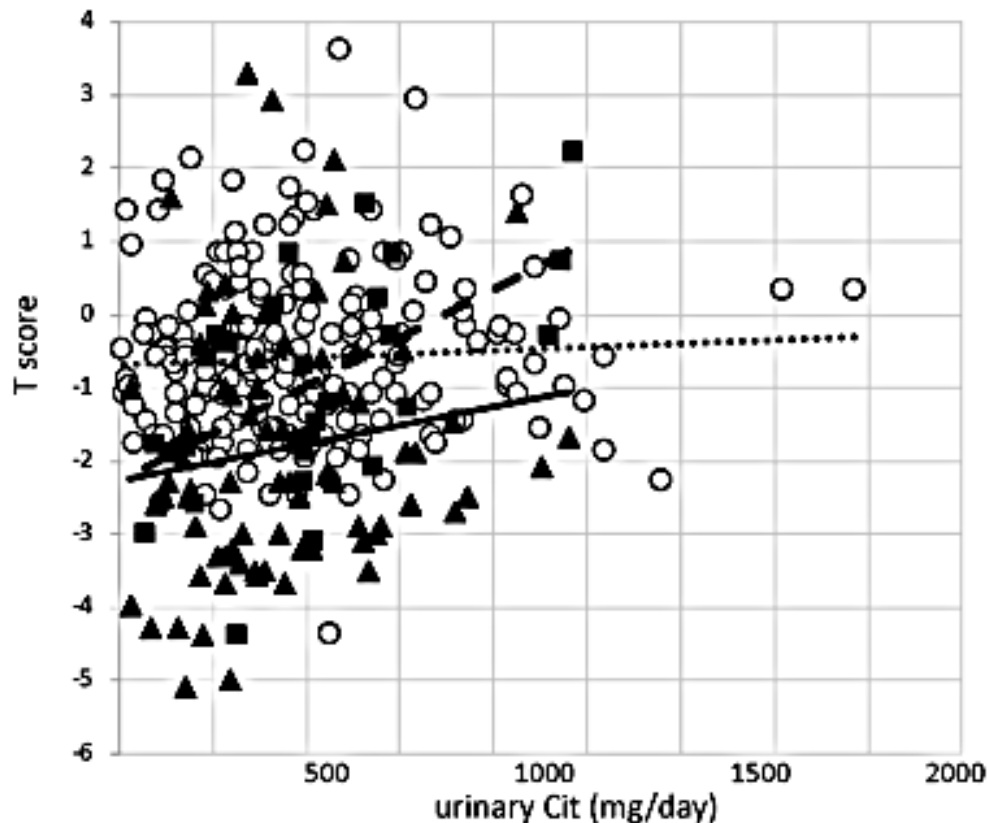
Means ± SD are shown

P values were calculated by unequal variance (Students's *t* tests)

Grupo 1: Al menos 2 episodios de urolitiasis y normocalciuria (N= 155)

Grupo 2: Controles sin historia personal ni familiar de urolitiasis (N=60)

HIPOCITRATURIA, HIPEROXALURIA Y DMO

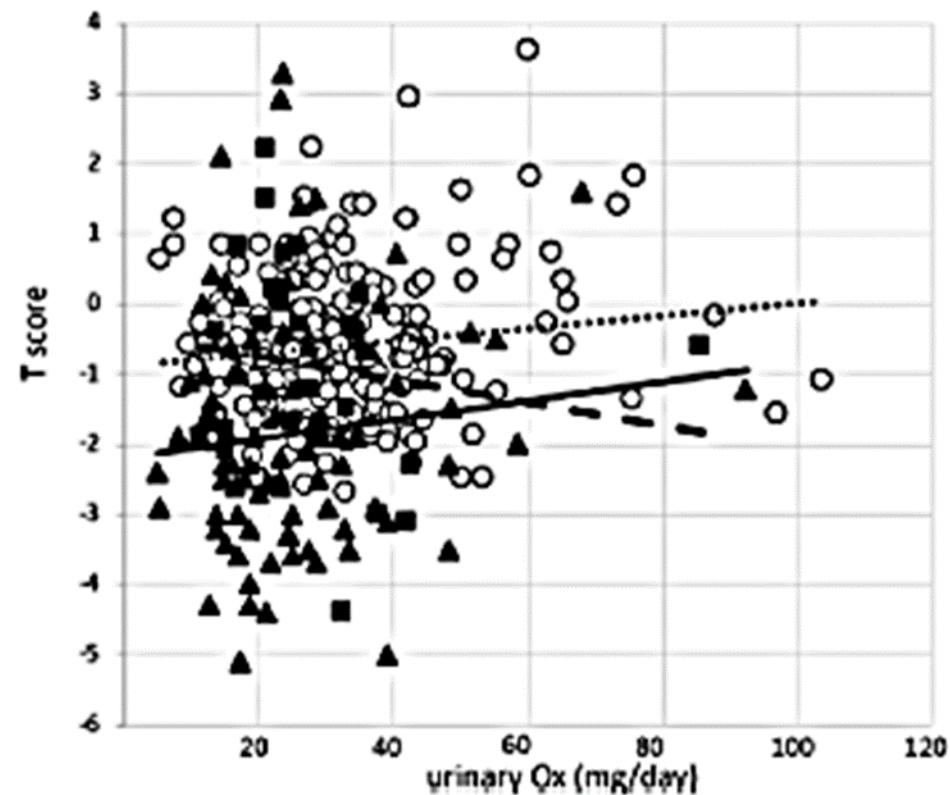


Pruebas de correlación de rango de Spearman entre Tscore y citrato en orina de 24 horas (N= 370)

Varones: coef=0.18, p=0.01

Mujeres premenopausicas: coef=0.57 p=0.003

Mujeres postmenopausicas: coef=0.22, p=0.12

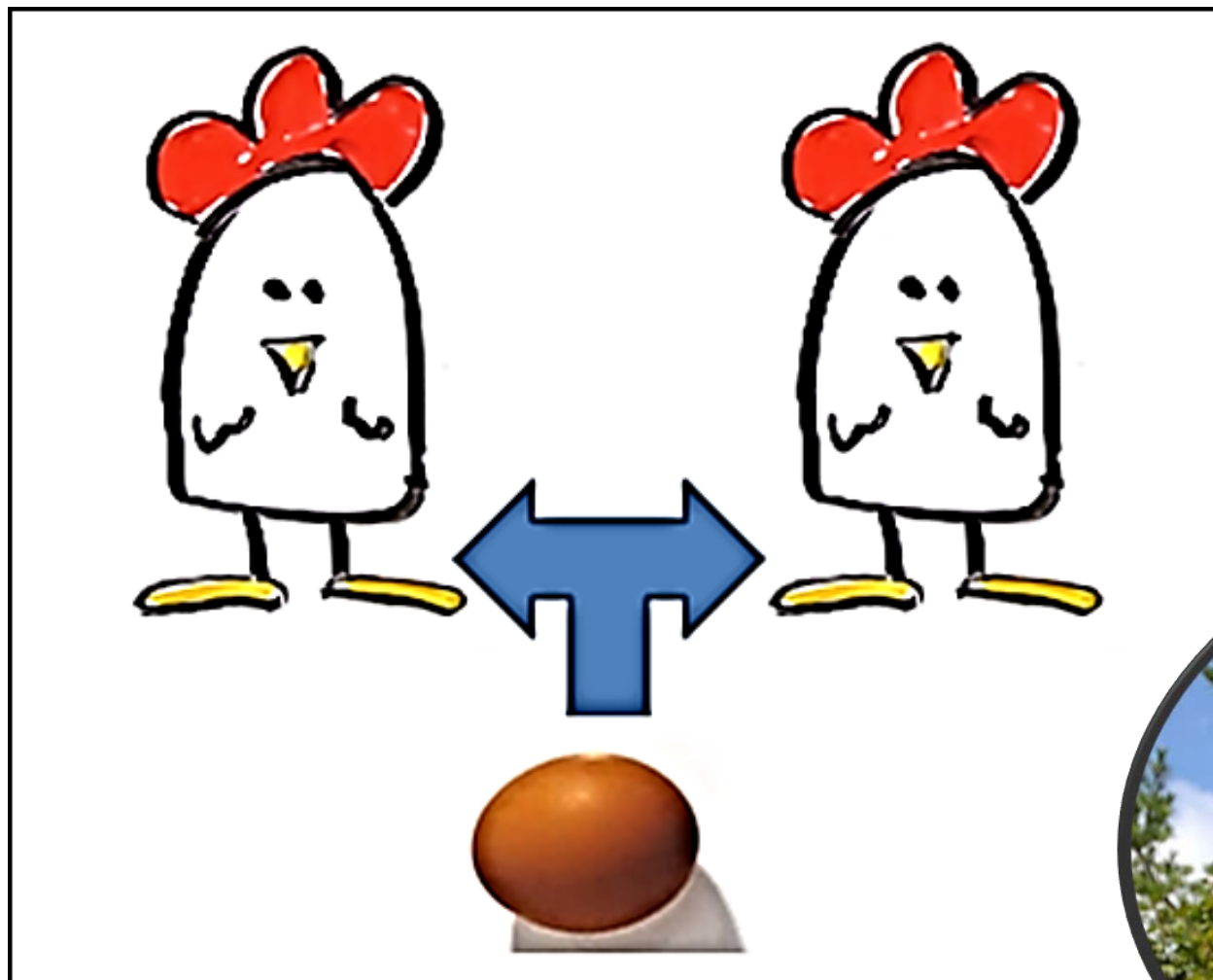


Pruebas de correlación de rango de Spearman entre Tscore y oxalato en orina de 24 horas (N= 370)

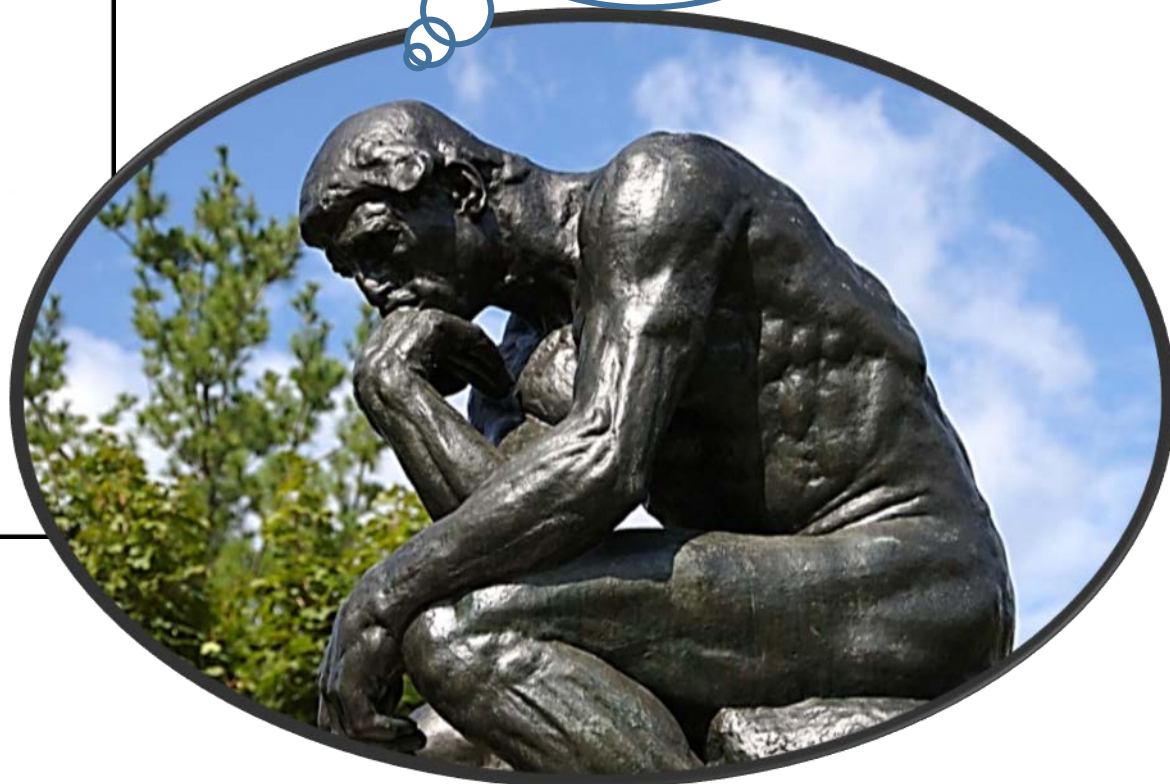
Varones: coef=0.14, p=0.05

Mujeres premenopausicas: coef=0.21 p=0.31

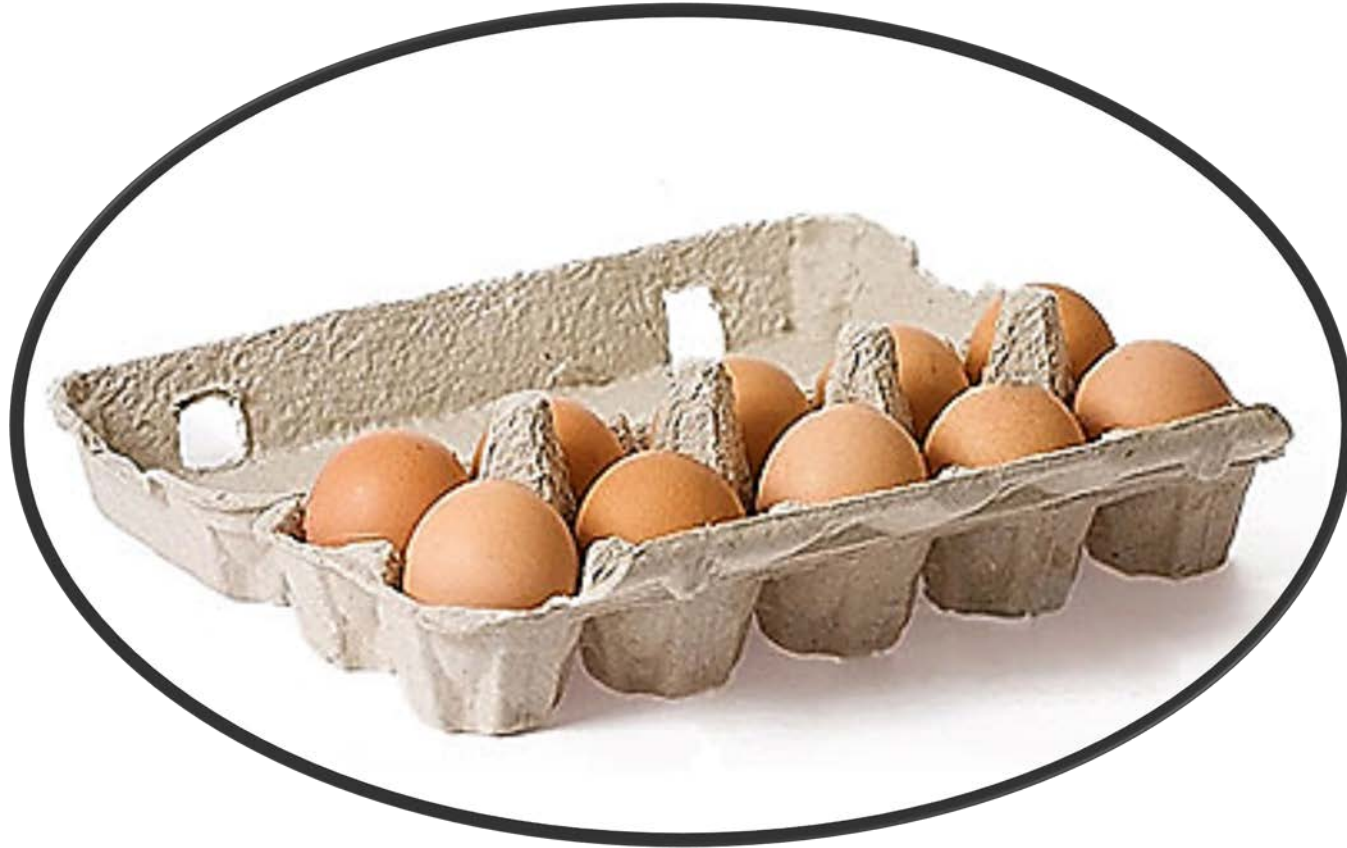
Mujeres postmenopausicas: coef=0.11, p=0.44



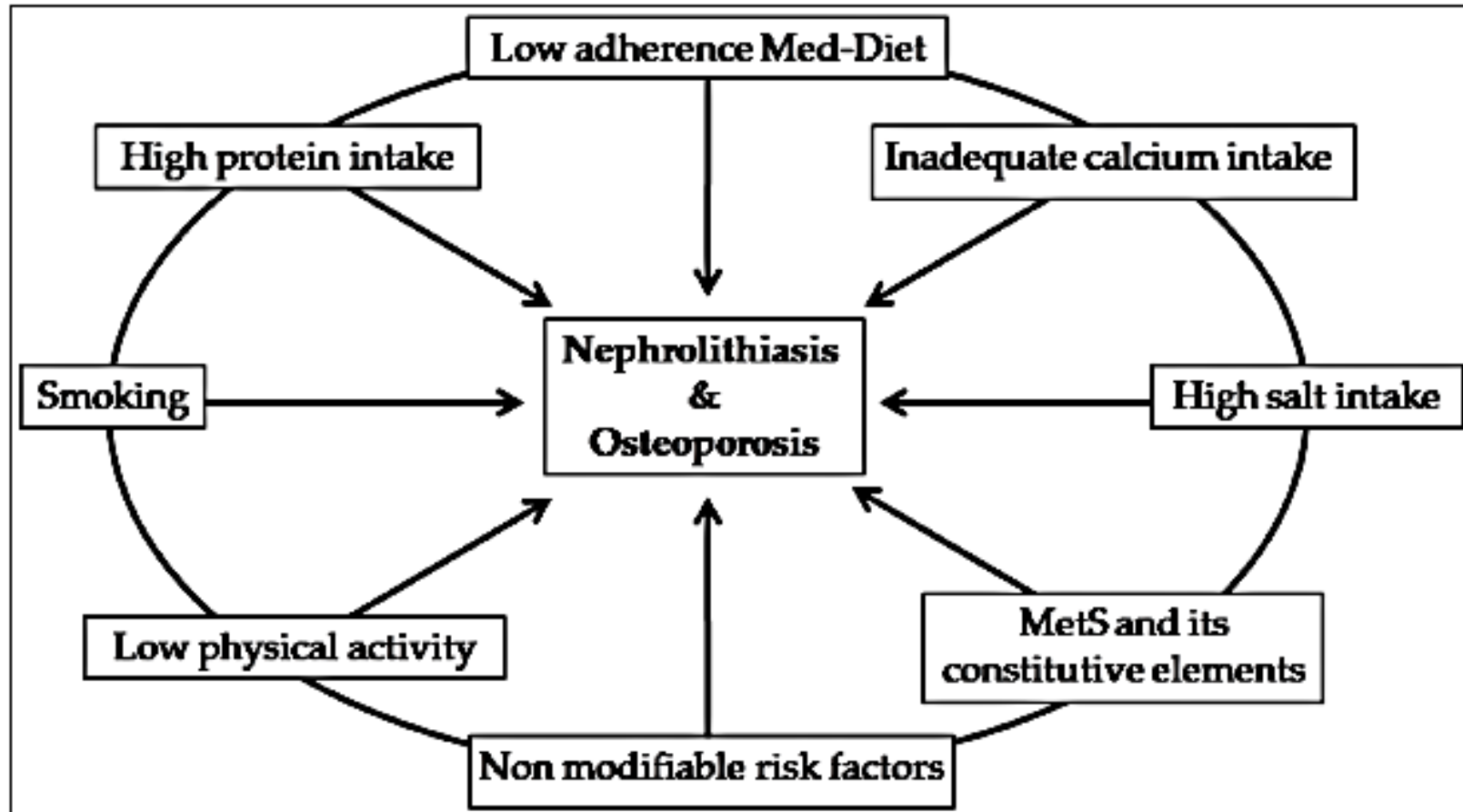
¿Y cuál sería el huevo?



FACTORES PATOGENICOS COMUNES

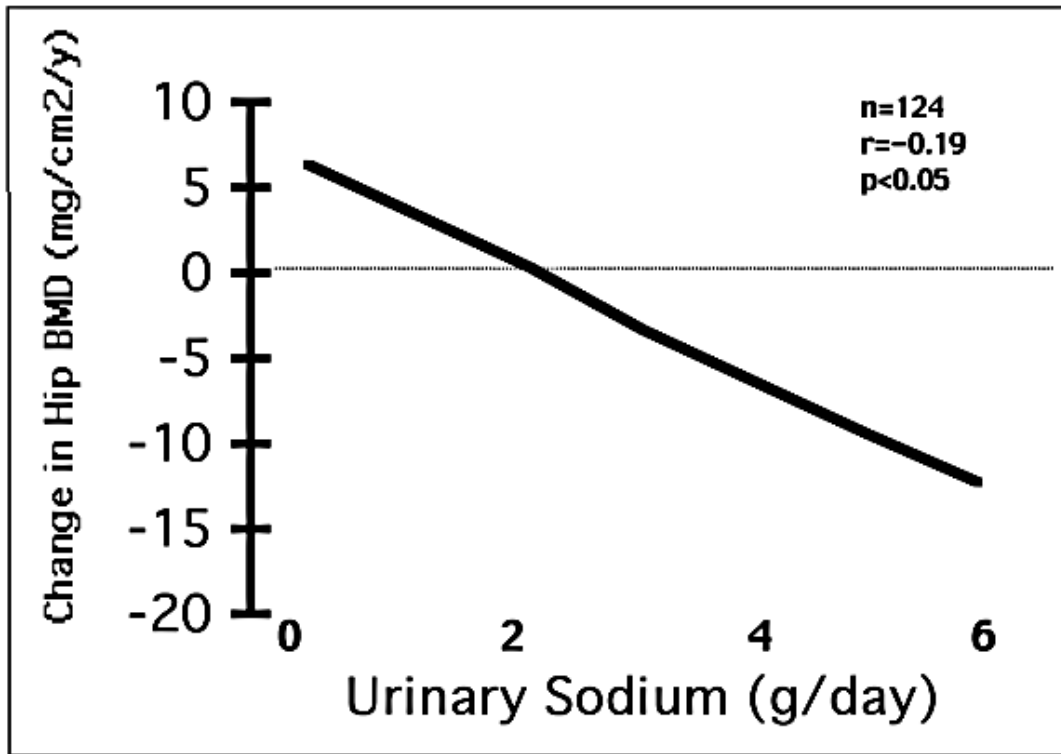


FACTORES PATOGENÉTICOS COMUNES



Rendina D. *Int. J. Mol. Sci.* 2020, 21, 8183; doi:10.3390/ijms21218183

INGESTA AUMENTADA DE SAL



| Nutrient | Group | | P value |
|--------------|-----------------|-----------------|---------|
| | IH (n = 55) | NC (n = 50) | |
| Calcium (mg) | 520.13 ± 245.62 | 531.44 ± 299.48 | 0.832 |
| Salt (g)* | 16.47 ± 5.93 | 11.79 ± 3.87 | < 0.001 |

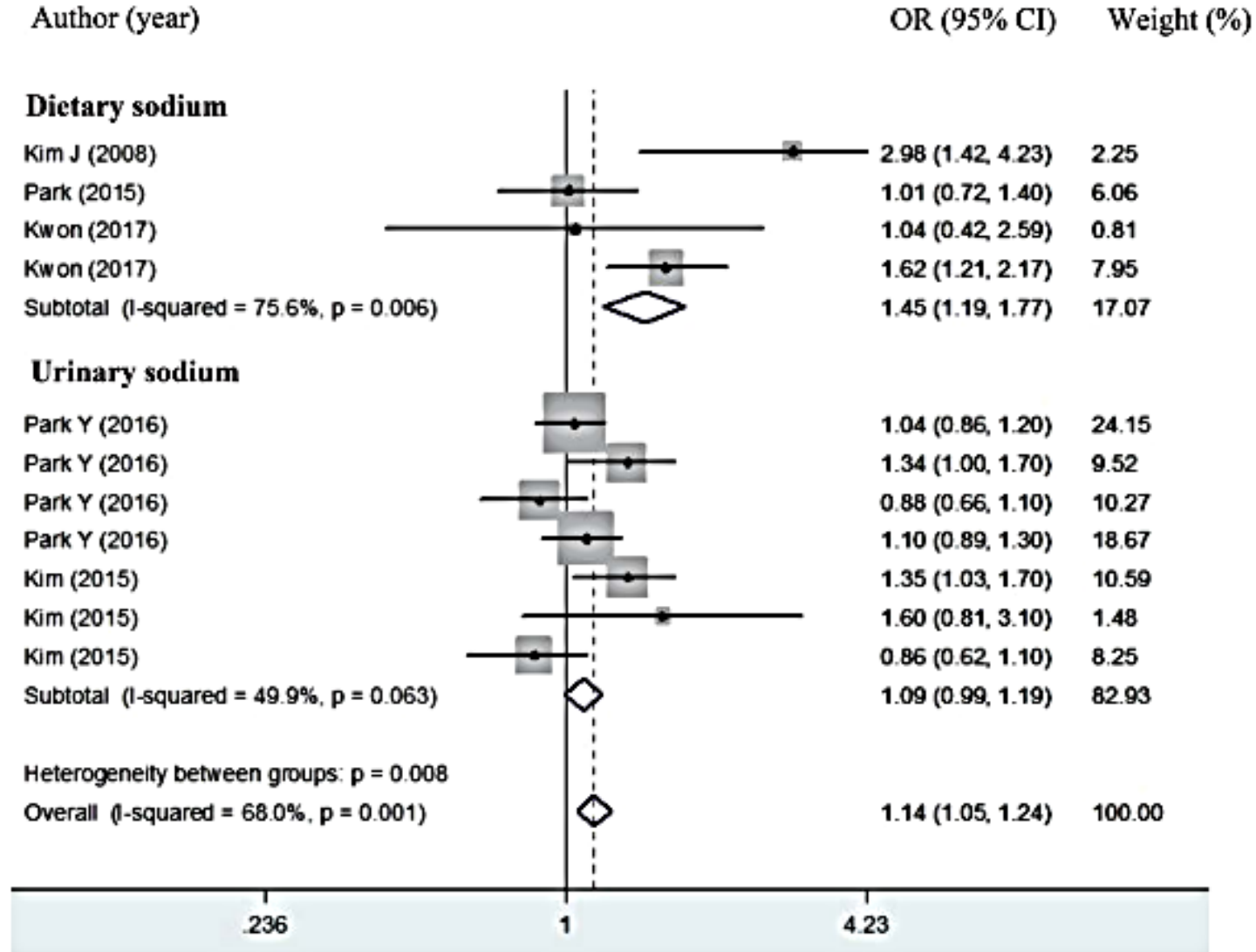
* Based on 24-hour urine

Descripción de la mediana y DS para la composición de la ingesta dietética diaria en sujetos hipercalciuricos y controles

Cappuccio, F.P.J. *Nephrol.* 2000, 13, 169–177

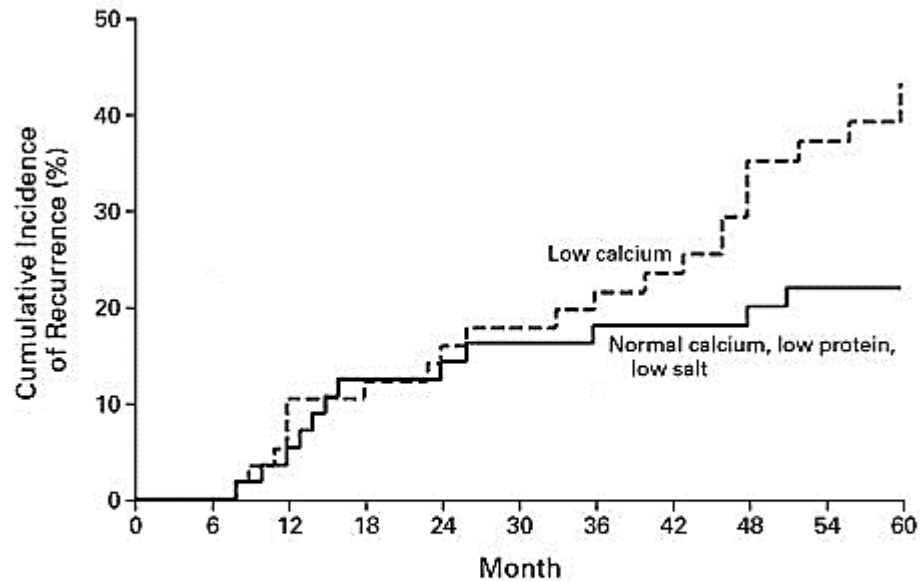
Timio, F. *Blood Press.* 2003, 12, 122–127

INGESTA DE SAL Y RIESGO DE FRACTURAS



INGESTA DE CALCIO Y RIESGO DE UROLITIASIS

| VARIABLE* | DIETARY CALCIUM† | | | | | CHI (P FOR TREND)‡ |
|---------------------------------|-----------------------|-----------------------|-----------------------|-----------------------|-----------------------|--------------------|
| | GROUP 1 (N = 8861) | GROUP 2 (N = 9029) | GROUP 3 (N = 9106) | GROUP 4 (N = 9184) | GROUP 5 (N = 9330) | |
| Dietary calcium intake (mg/day) | <605 | 605–722 | 723–848 | 849–1049 | ≥1050 | — |
| Incidence/100,000 person-yr | 435 | 310 | 279 | 266 | 243 | — |
| No. of cases | 139 | 102 | 93 | 89 | 82 | |
| Age-adjusted RR | 1.0 | 0.71 | 0.64 | 0.61 | 0.56 | –4.37 (<0.001) |
| 95% CI | | 0.55–0.92 | 0.50–0.83 | 0.47–0.80 | 0.43–0.73 | |
| Multivariate RR | 1.0 | 0.74 | 0.68 | 0.68 | 0.66 | –2.38 (0.018) |
| 95% CI | | 0.57–0.97 | 0.52–0.90 | 0.51–0.90 | 0.49–0.90 | |



| No. AT RISK | 0 | 6 | 12 | 18 | 24 | 30 | 36 | 42 | 48 | 54 | 60 |
|---|----|----|----|----|----|----|----|----|----|----|----|
| Low calcium | 60 | 59 | 51 | 49 | 46 | 44 | 42 | 39 | 33 | 31 | 28 |
| Normal calcium, low protein, low salt | 60 | 57 | 53 | 47 | 46 | 45 | 44 | 43 | 41 | 40 | 40 |

Curhan, G.C. N. Engl. J. Med. 1993, 328, 833–838

Borghi, L. N. Engl. J. Med. 2002, 346, 77–84

INGESTA DE AZUCARES SIMPLES

OR ajustadas para el consumo de refrescos de cola y riesgo de fractura de muñeca y antebrazo

| | Odds ratios (95% CI): per unit increase Times of drinking cola/week |
|---------------------|---|
| Crude odds ratio | 1.39 (1.01, 1.91)* |
| Step 1 ^a | 1.43 (1.03, 1.97)* |
| Step 2 ^a | 1.33 (0.96, 1.83) |
| Step 3 ^a | 1.32 (0.96, 1.82) |
| Step 4 ^a | 1.31 (0.94, 1.83) |

^a Step 1: adjusted for milk intake; step 2: television, computer and video watching only; step 3: bone mineral density only; step 4: all factors including milk intake, television, computer and video watching, lumbar spine bone apparent mineral density, and metacarpal morphometry
 $P < 0.05$

Ma, D. Calcif. Tissue Int. 2004, 75, 286–291

Riesgo relativo de cálculos renales según categorías de consumo de diferentes bebidas

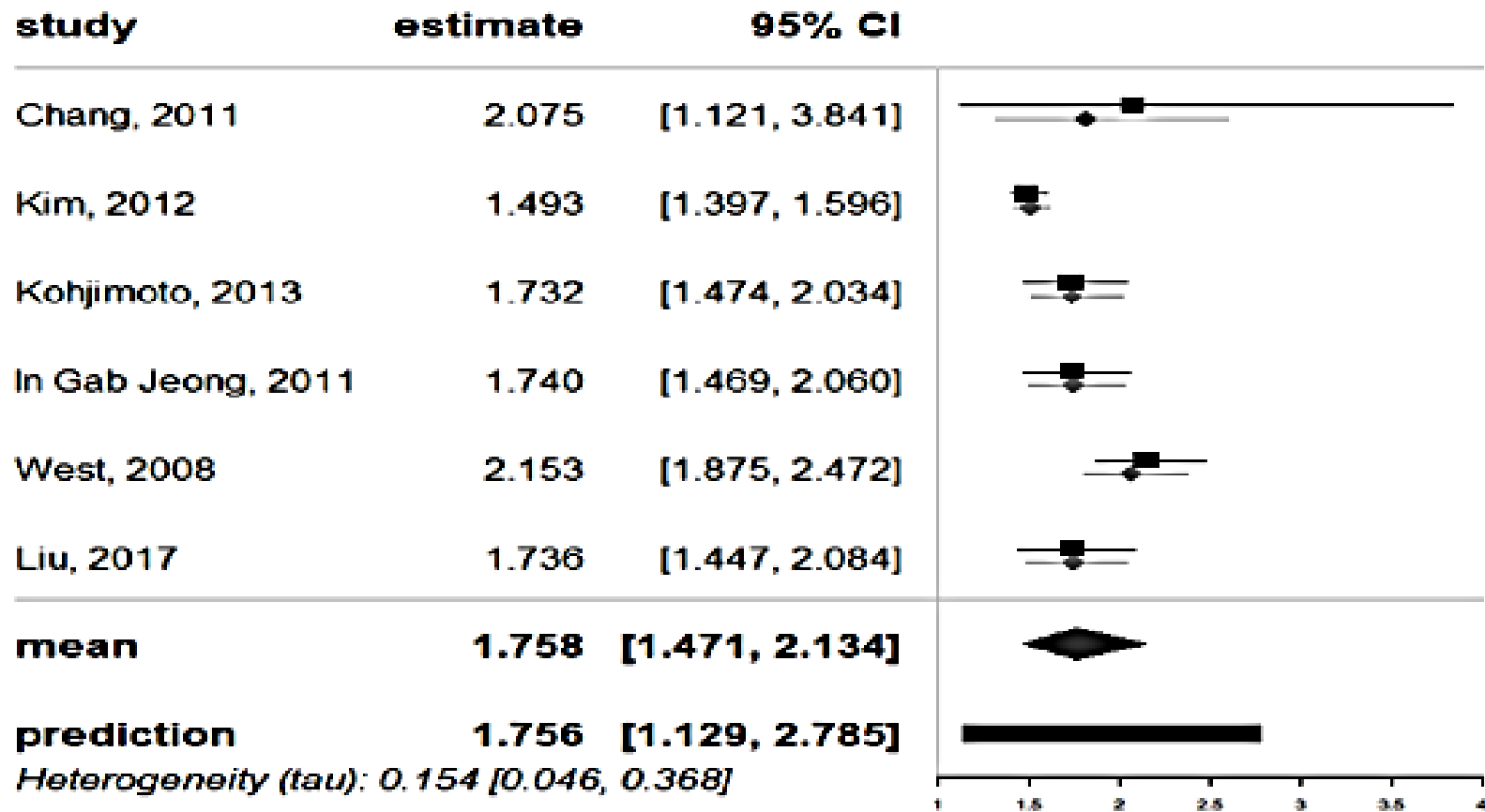
| | | | | | | Trend |
|-----------------------------|------------------|---------------------|---------------------|---------------------|---------------------|--------|
| Sugar-sweetened cola | | | | | | |
| Cases (IR) | 2962 (149) | 555 (193) | 443 (236) | 117 (262) | 385 (287) | |
| Age-adjusted HR (95% CI) | 1.00 (reference) | 1.20 (1.00 to 1.44) | 1.41 (1.27 to 1.57) | 1.52 (1.18 to 1.95) | 1.77 (1.40 to 2.22) | <0.001 |
| Multivariate HR (95% CI) | 1.00 (reference) | 1.07 (0.89 to 1.27) | 1.19 (1.06 to 1.34) | 1.28 (1.09 to 1.50) | 1.23 (0.98 to 1.55) | 0.02 |
| Artificially sweetened cola | | | | | | |
| Cases (IR) | 2381 (174) | 454 (166) | 564 (170) | 181 (174) | 882 (158) | |
| Age-adjusted HR (95% CI) | 1.00 (reference) | 0.93 (0.75 to 1.14) | 0.98 (0.82 to 1.17) | 0.94 (0.80 to 1.12) | 0.87 (0.76 to 1.00) | 0.05 |
| Multivariate HR (95% CI) | 1.00 (reference) | 1.01 (0.85 to 1.21) | 1.05 (0.95 to 1.17) | 1.03 (0.87 to 1.21) | 0.91 (0.78 to 1.06) | 0.08 |

Ferraro, P.M. Clin. J. Am. Soc. Nephrol. 2013, 8, 1389–1395

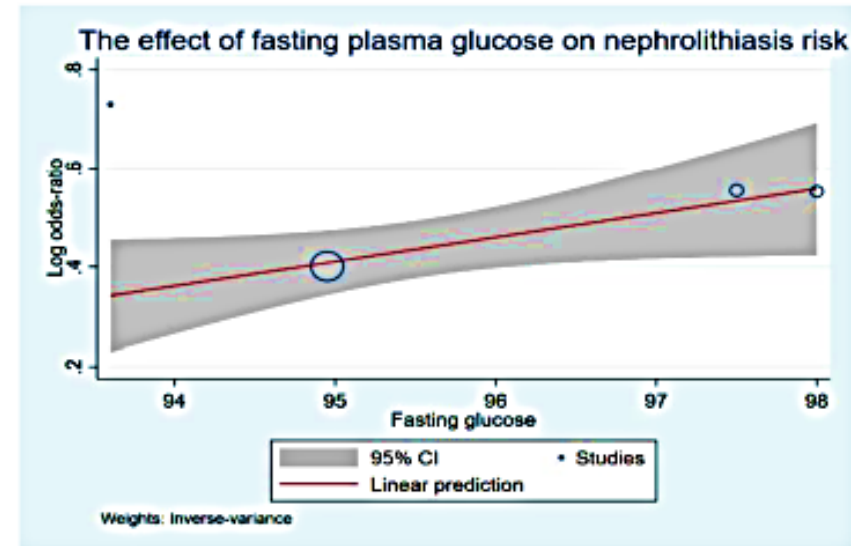
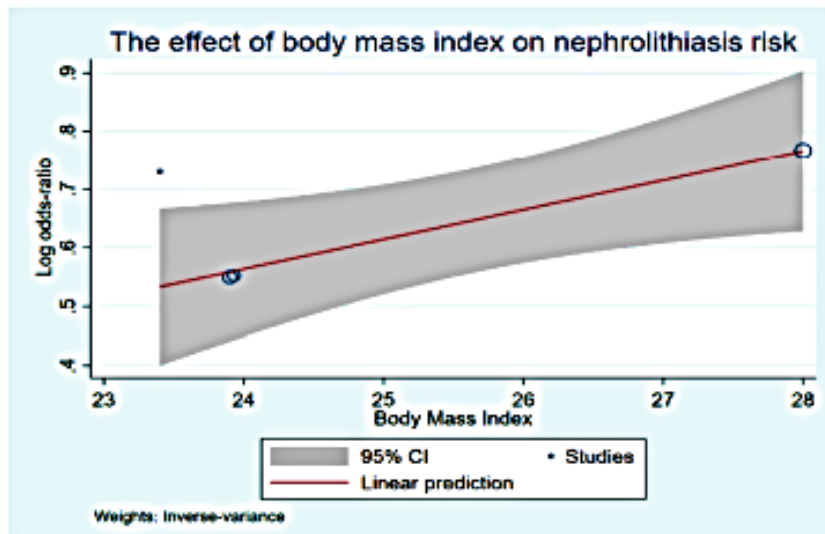
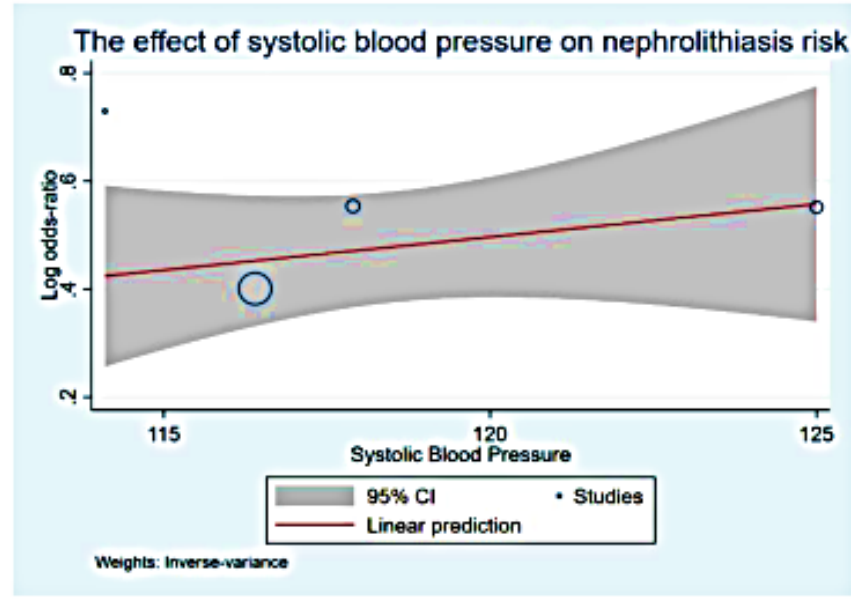
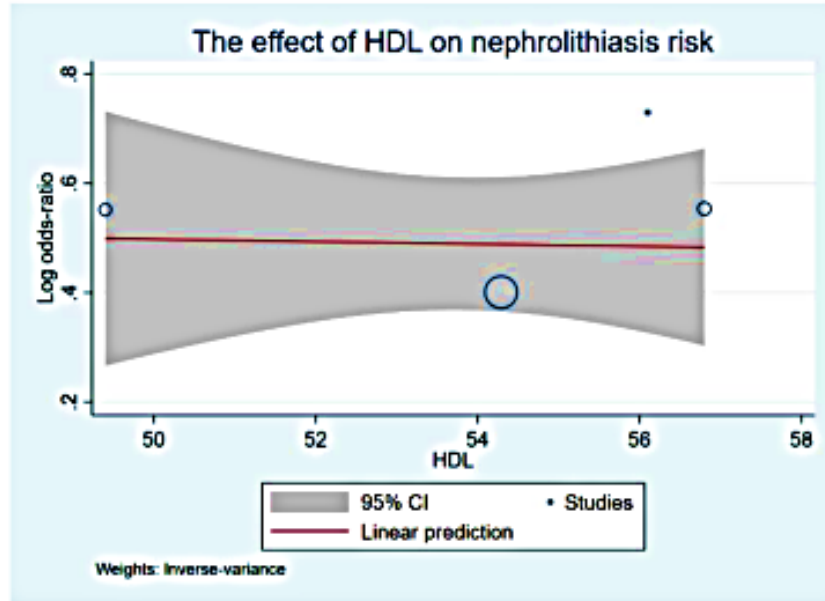
SINDROME METABOLICO

The risk of Nephrolithiasis in Patients with Metabolic Syndrome

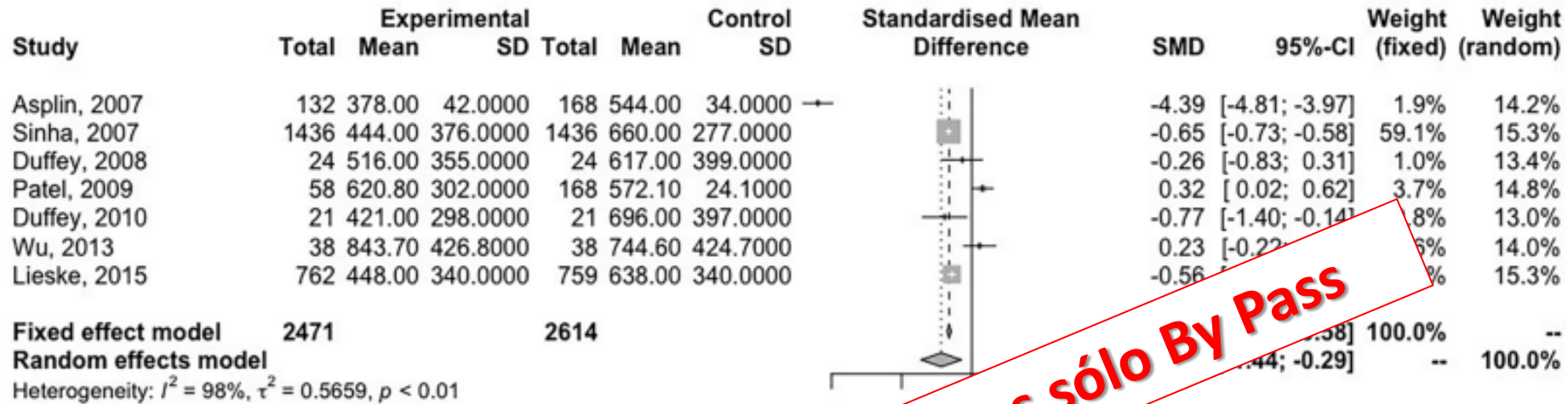
■ quoted estimate ◆ shrinkage estimate



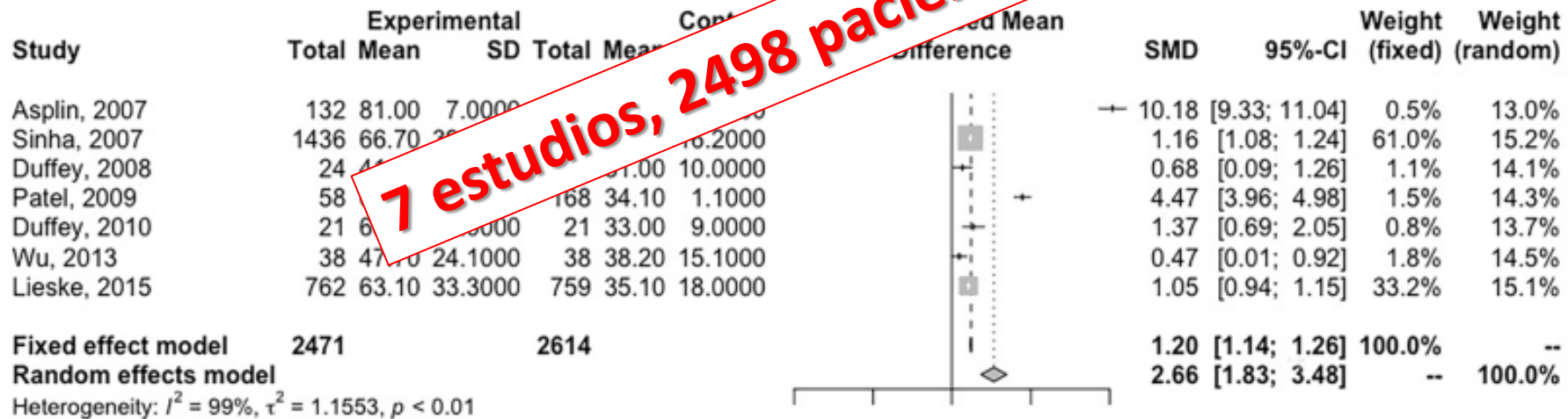
SINDROME METABOLICO



CIRUGIA BARIATRICA Y UROLITIASIS

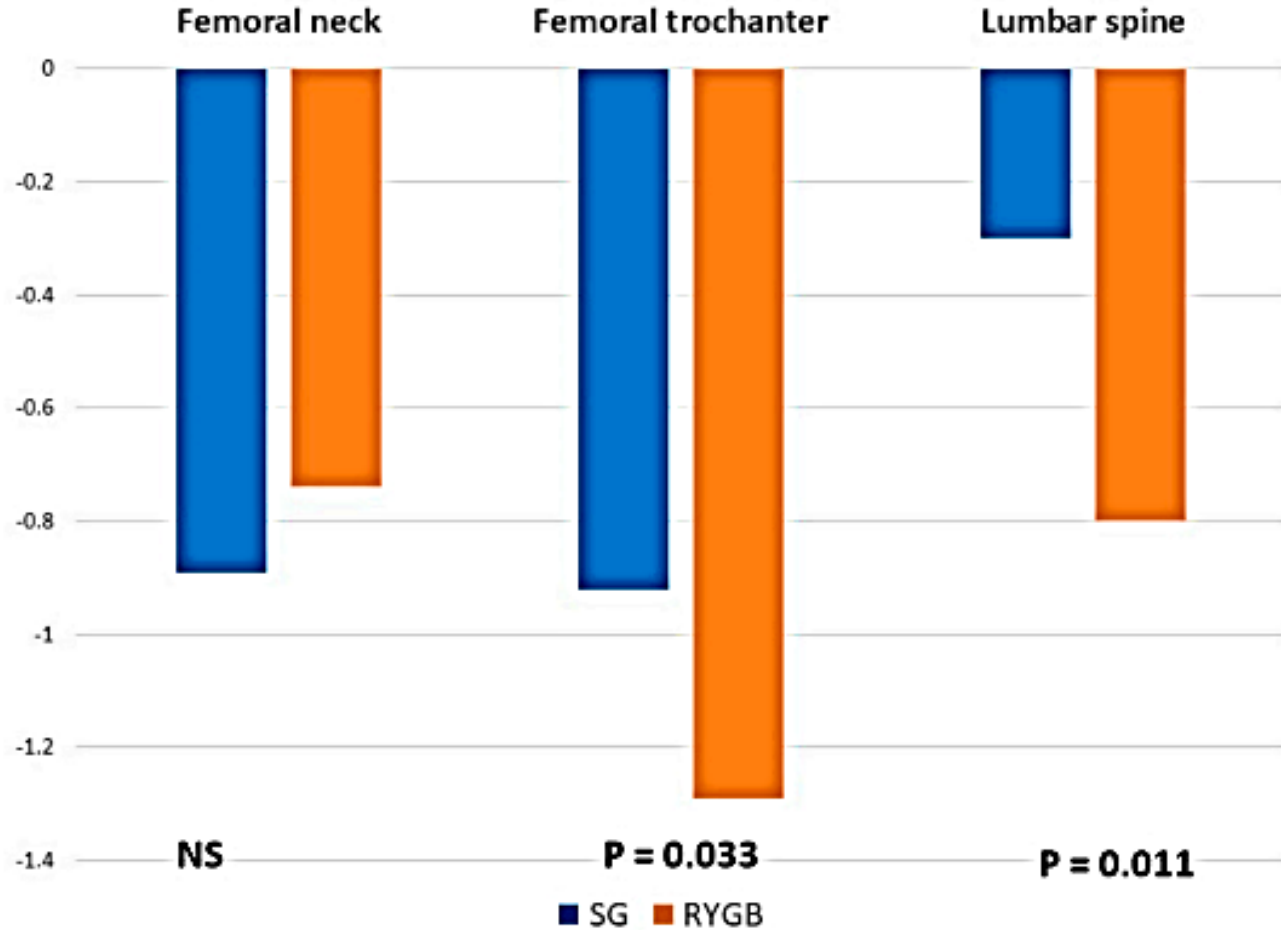


Citrato orina 24 h



Oxalato orina 24 h

CIRUGIA BARIATRICA Y DMO



Diferencias medias entre los valores basales y postoperatorios de densidad mineral del hueso

PATRONES DE ALIMENTACION ACEITE DE OLIVA, VINO y UROLITIASIS

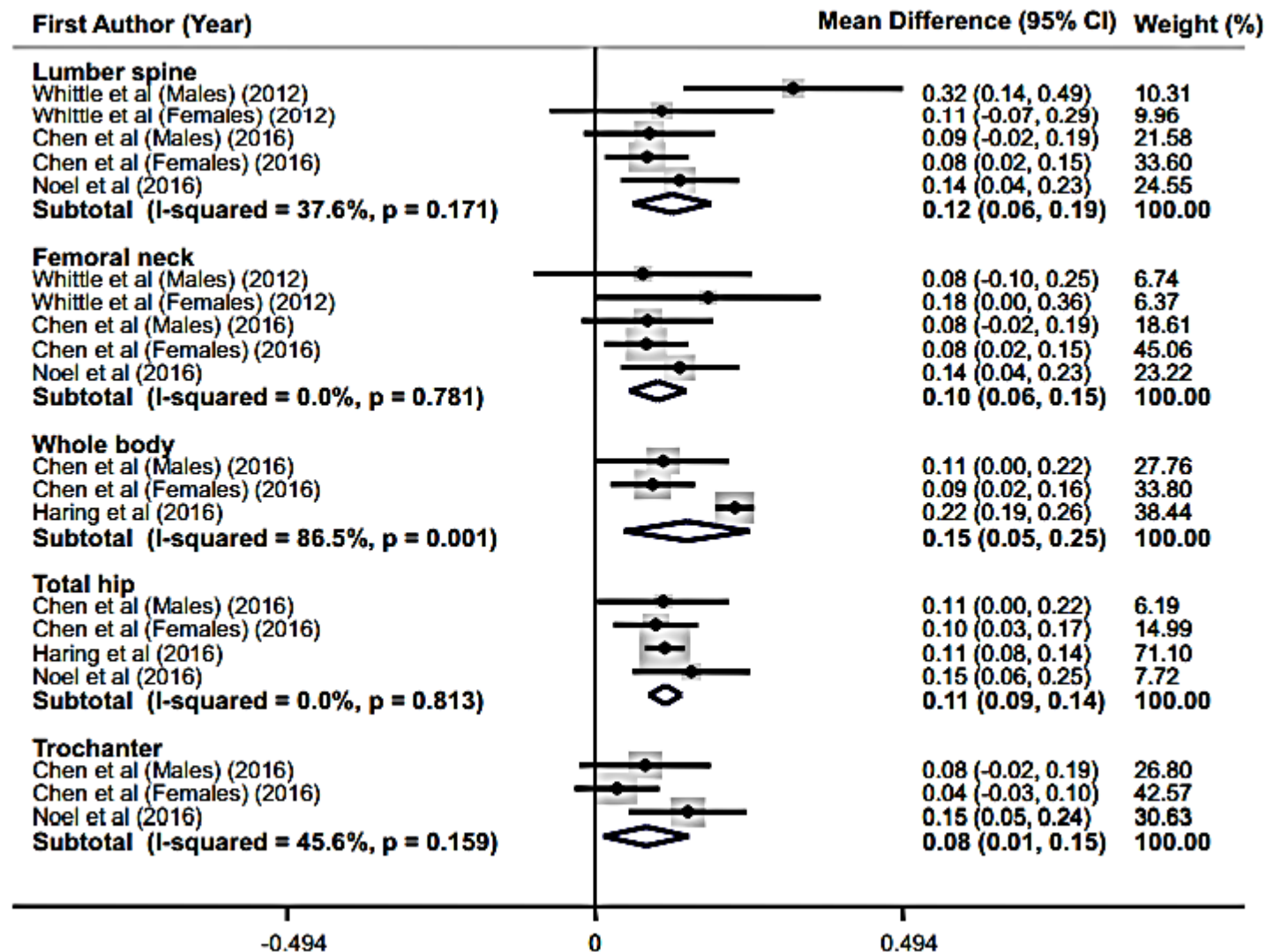
| | Score 1 <1 spoon/day | Score 2 1-3 spoons/day | Score 3 or 4 ≥4 spoons/day | |
|-------------------|----------------------|------------------------|----------------------------|-----------------|
| Olive oil | | | | |
| Stone formers | 6 (13.3) | 28 (62.2) | 11 (24.4) | |
| Non-stone formers | 33 (7.5) | 221 (50) | 188 (42.6) | |
| OR | 1 | 0.7 | 0.32 | |
| 95% CI | | 0.3-1.8 | 0.1-0.9 | |
| p | | 0.46 | 0.036 | |
| | Score 1 | Score 2 | Score 3 | Score 4 |
| | No consumption | 1-2 glasses/week | 3-7 glasses/week | >7 glasses/week |
| Wine | | | | |
| Stone formers | 13 (30.2) | 4 (9.3) | 16 (37.2) | 10 (23.3) |
| Non-stone formers | 189 (42.9) | 115 (26.1) | 83 (18.8) | 54 (12.2) |
| OR | 1 | 0.51 | 2.8 | 2.7 |
| 95% CI | | 0.2-1.6 | 1.3-6.1 | 1.1-6.5 |
| p | | 0.24 | 0.009 | 0.027 |

Análisis de regresión multinomial del consumo de aceite de oliva y vino en formadores y no formadores de cálculos

Soldati, L. J. Transl. Med. 2014, 12, 34

PATRONES DE ALIMENTACION DIETA MEDITERRANEA y OSTEOPOROSIS

Diagrama de bosque de las diferencias medias en la densidad mineral ósea (DMO) en cinco sitios diferentes, comparando la adherencia más alta y más baja a la dieta mediterránea



PATRONES DE ALIMENTACION DIETA MEDITERRANEA y FRACTURAS

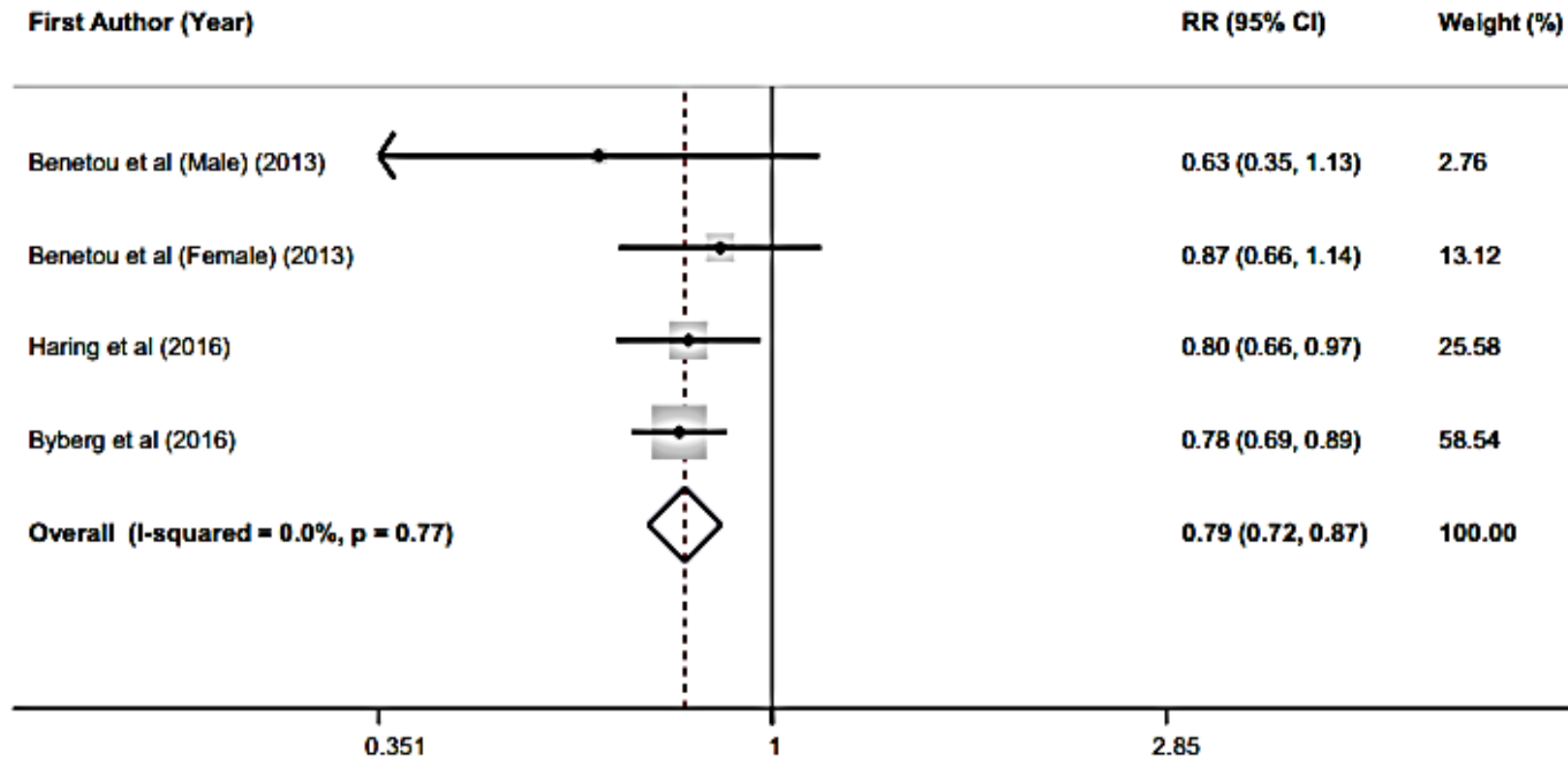


Diagrama de bosque de la asociación entre la adherencia a la dieta mediterránea y el riesgo de fractura de cadera

Malmir, H. Eur. J. Nutr. 2018, 57, 2147–2160

EJERCICIO FISICO

| | Physical activity (METs/week) | | | | | p-value for trend |
|------------------------|-------------------------------|-------------------|-------------------|-------------------|-------------------|-------------------|
| | < 5 | 5 – 9.9 | 10 – 19.9 | 20 – 29.9 | ≥30 | |
| HPFS | | | | | | |
| cases | 367 | 205 | 356 | 250 | 663 | |
| person-years | 136,789 | 79,675 | 130,804 | 99,102 | 263,469 | |
| age-adjusted HR | 1.00 | 0.94 (0.79, 1.12) | 0.99 (0.86, 1.15) | 0.93 (0.79, 1.09) | 0.92 (0.80, 1.04) | 0.26 |
| first multivariate HR | 1.00 | 0.94 (0.79, 1.12) | 0.99 (0.85, 1.14) | 0.92 (0.78, 1.08) | 0.93 (0.81, 1.06) | 0.38 |
| second multivariate HR | 1.00 | 0.96 (0.81, 1.15) | 1.03 (0.88, 1.19) | 0.97 (0.82, 1.15) | 1.00 (0.87, 1.14) | 0.94 |
| NHS I | | | | | | |
| cases | 563 | 243 | 275 | 141 | 217 | |
| person-years | 456,604 | 213,348 | 264,921 | 153,582 | 218,050 | |
| age-adjusted HR | 1.00 | 0.92 (0.79, 1.07) | 0.84 (0.73, 0.98) | 0.75 (0.62, 0.90) | 0.81 (0.69, 0.95) | 0.003 |
| first multivariate HR | 1.00 | 0.96 (0.82, 1.11) | 0.91 (0.79, 1.05) | 0.84 (0.69, 1.01) | 0.94 (0.80, 1.10) | 0.27 |
| second multivariate HR | 1.00 | 0.99 (0.85, 1.15) | 0.95 (0.82, 1.10) | 0.89 (0.74, 1.08) | 1.01 (0.85, 1.19) | 0.88 |
| NHS II | | | | | | |
| cases | 600 | 382 | 441 | 245 | 407 | |
| person-years | 383,275 | 251,592 | 321,629 | 185,592 | 310,258 | |
| age-adjusted HR | 1.00 | 0.94 (0.82, 1.07) | 0.85 (0.75, 0.96) | 0.82 (0.71, 0.95) | 0.81 (0.71, 0.92) | 0.001 |
| first multivariate HR | 1.00 | 1.00 (0.88, 1.13) | 0.95 (0.84, 1.07) | 0.95 (0.81, 1.10) | 0.99 (0.87, 1.12) | 0.81 |
| second multivariate HR | 1.00 | 1.01 (0.89, 1.15) | 0.97 (0.86, 1.10) | 0.98 (0.84, 1.14) | 1.03 (0.90, 1.18) | 0.64 |

HR e IC 95 % de cálculos renales incidentes por categoría de actividad física en 3 estudios

TABAQUISMO, ALCOHOL Y OBESIDAD

| Characteristics | N of Nephrolithiasis (Exposure/Total, %) | N of Control (Exposure/Total, %) | Odd Ratios for Nephrolithiasis (95% Confidence Interval) | | | |
|-----------------------|--|--|--|-----------------|------------------------|-----------------|
| | | | Crude [†] | <i>p</i> -Value | Adjusted ^{†‡} | <i>p</i> -Value |
| Smoking status | 4887/28,395 (17.2) | 18,944/113,580 (16.7) | 1.04 (1.01–1.08) | 0.023 * | 1.03 (1.00–1.07) | 0.102 |
| Alcohol consumption | 9955/28,395 (35.1) | 42,361/113,580 (37.3) | 0.90 (0.87–0.92) | <0.001 * | 0.89 (0.86–0.92) | <0.001 * |
| Obesity | | | | <0.001 * | | <0.001 * |
| <18.5 (underweight) | 392/28,395 (1.4) | 2691/113,580 (2.4) | 0.70 (0.63–0.78) | <0.001 * | 0.70 (0.63–0.78) | <0.001 * |
| 18.5 to 23 (normal) | 8016/28,395 (28.2) | 38,969/113,580 (34.3) | 1.00 | | 1.00 | |
| 23 to 25 (overweight) | 8189/28,395 (28.8) | 31,561/113,580 (27.8) | 1.27 (1.22–1.31) | <0.001 * | 1.27 (1.22–1.31) | <0.001 * |
| 25 to 30 (obese I) | 10,728/28,395 (37.8) | 37,052/113,580 (32.6) | 1.42 (1.37–1.46) | <0.001 * | 1.42 (1.37–1.46) | <0.001 * |
| 30 (obese II) | 1070/28,395 (3.8) | 3307/113,580 (2.9) | 1.58 (1.47–1.70) | <0.001 * | 1.59 (1.47–1.71) | <0.001 * |

Razones de probabilidad brutas y ajustadas (intervalos de confianza del 95%) de tabaquismo, consumo de alcohol, y obesidad por nefrolitiasis.

VITAMINA D

| | Group 1 | Group 2 | <i>p</i> |
|-------------------------|---------------|---------------|----------|
| <i>Serum parameters</i> | | | |
| Creatinine | 0.85 ± 0.15 | 0.87 ± 0.23 | n.s. |
| Sodium | 141.81 ± 2.07 | 141.58 ± 2.24 | n.s. |
| Potassium | 4.45 ± 0.41 | 4.52 ± 0.41 | n.s. |
| Chloride | 103.74 ± 2.61 | 102.37 ± 2.97 | 0.0001 |
| Uric acid | 4.76 ± 1.27 | 5.11 ± 1.34 | 0.01 |
| Calcium | 9.43 ± 0.41 | 9.58 ± 0.42 | 0.001 |
| Phosphorus | 3.35 ± 0.54 | 3.03 ± 0.51 | 0.0001 |
| iPTH | 50.21 ± 17.68 | 48.78 ± 22.21 | n.s. |
| 25-OH-vitamin D | 28.48 ± 9.49 | 25.71 ± 10.18 | 0.02 |

Group 1: 127 controls with no history of kidney stones

Group 2: 239 patients with a history of calcium kidney stones

Girón-Prieto, M.S. Int. Urol. Nephrol. 2016, 48, 1243–1246

FACTORES DE RIESGO NO MODIFICABLES

| <i>Gene</i> | <i>HGNC Symbol</i> | <i>Location</i> | <i>Ref.</i> |
|---|--------------------|---------------------|------------------|
| <i>Calcium-sensing receptor</i> | <i>CASR</i> | <i>3q13.3-q21.1</i> | <i>[124–126]</i> |
| <i>Vitamin D receptor</i> | <i>VDR</i> | <i>12q13.11</i> | <i>[127–132]</i> |
| <i>Alkaline phosphatase</i> | <i>ALPL</i> | <i>1p36.12</i> | <i>[133,134]</i> |
| <i>Osteopontin</i> | <i>SPP1</i> | <i>4q22.1</i> | <i>[135,136]</i> |
| <i>Claudin 14</i> | <i>CLDN14</i> | <i>21q22.13</i> | <i>[137–139]</i> |
| <i>Type 2a sodium–phosphate cotransporter</i> | <i>SLC34A1</i> | <i>5q35.3</i> | <i>[140]</i> |
| <i>Fibroblast growth factor 23</i> | <i>FGF23</i> | <i>12p13.32</i> | <i>[141]</i> |
| <i>25(OH)D-24-hydroxylase</i> | <i>CYP24A1</i> | <i>20q13.2</i> | <i>[142]</i> |

HGNC: Human Genome Organisation (HUGO) Gene Nomenclature Committee.

Genes asociado con la ocurrencia de osteoporosis y urolitiasis

Howles, S.A. Nat. Rev. Urol. 2020, 17, 407–421

FACTORES DE RIESGO NO MODIFICABLES POLIMORFISMOS DE CASR Y OSTEOPOROSIS

Comparación de las características clínicas y bioquímicas entre pacientes con genotipos AA y AS/SS del SNP A986S

| Patients characteristics | A986S | | <i>p</i> ^a |
|---|----------------------|--------------------------|-----------------------|
| | AA (<i>n</i> = 206) | AS + SS (<i>n</i> = 38) | |
| Testosterone (nM) | 15.72 ± 4.95 | 15.02 ± 4.46 | 0.362 |
| Serum creatinine (μmol/L) | 82.48 ± 20.41 | 88.33 ± 20.20 | 0.210 |
| Serum calcium (mM) | 2.36 ± 0.08 | 2.44 ± 0.12 | 0.001 |
| Serum phosphorus (mM) | 0.87 ± 0.15 | 0.87 ± 0.15 | 0.298 |
| Urinary calcium (mmol 24 h) | 8.54 ± 3.30 | 8.12 ± 3.15 | 0.883 |
| Urinary phosphate (mmol 24 h) | 22.46 ± 11.79 | 22.11 ± 12.11 | 0.491 |
| 25(OH)-vitamin D (nM) | 47.81 ± 19.13 | 43.76 ± 18.93 | 0.363 |
| PTH (ng/dL) | 53.11 ± 19.11 | 67.43 ± 14.12 | 0.007 |
| BMD LS (g/cm ²) | 0.95 ± 0.17 | 0.91 ± 0.15 | 0.366 |
| BMD FN (g/cm ²) | 0.94 ± 0.15 | 0.90 ± 0.18 | 0.048 |
| Osteoporosis, <i>n</i> (%) ^b | 16 (7.8%) | 10 (23.8%) | 0.005 ^c |

FACTORES DE RIESGO NO MODIFICABLES POLIMORFISMOS DE CASR Y UROLITIASIS

| SNP | Allele | Genotype | Controls n (%) | Stone Formers n (%) | Odds Ratio (95% CI) | P |
|-----------|--------|----------|----------------|---------------------|---------------------|-------|
| rs7652589 | G | | 305 (73.3) | 204 (61.8) | 1 | |
| | A | | 111 (26.7) | 126 (38.2) | 1.69 (1.2–2.3) | .0012 |
| rs7648041 | C | | 291 (69.9) | 232 (70.3) | 1 | |
| | T | | 125 (30.1) | 98 (29.7) | 1.02 (0.7–1.4) | .906 |
| rs7648044 | C | | 361 (86.8) | 269 (81.5) | 1 | |
| | T | | 55 (13.2) | 61 (18.5) | 1.49 (1–2.2) | .049 |
| rs6776158 | A | | 306 (73.6) | 205 (62.1) | 1 | |
| | G | | 110 (26.4) | 125 (37.9) | 1.7 (1.2–2.3) | .0008 |
| rs1048213 | T | | 310 (76) | 262 (79.4) | 1 | |
| | C | | 98 (24) | 68 (20.6) | 0.82 (0.6–1.2) | .269 |
| rs1501899 | G | | 312 (75) | 204 (61.8) | 1 | |
| | A | | 104 (25) | 126 (38.2) | 1.87 (1.4–2.6) | .0001 |

Frecuencia de alelos y genotipos en los SNP del gen CaSR ubicados en la región reguladora de genes en formadores de cálculos y controles sanos

Vezzoli, G. J. Clin. Endocrinol. Metab. 2013, 98, 3839–3847

FACTORES DE RIESGO NO MODIFICABLES POLIMORFISMOS DE VDR Y UROLITIASIS

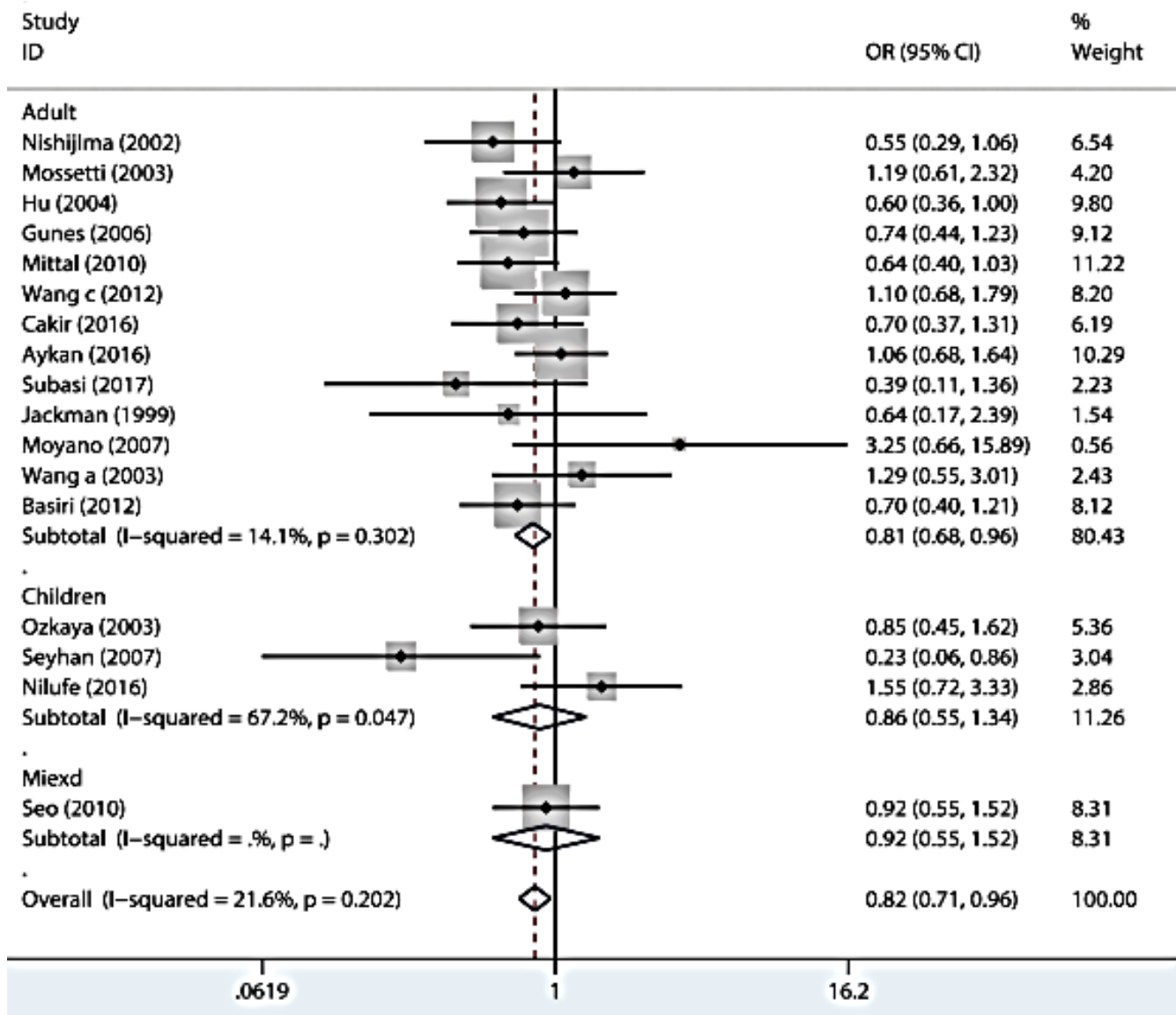
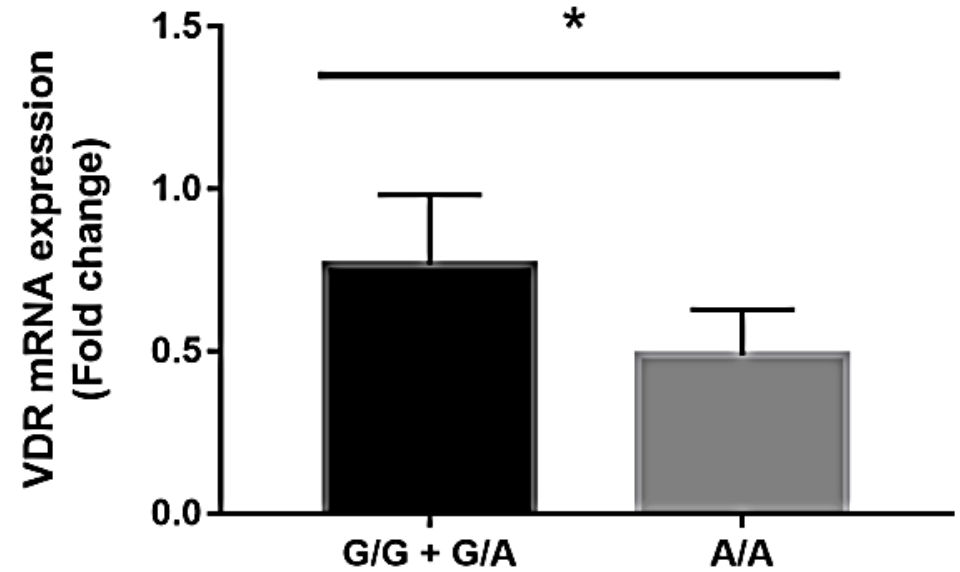


Diagrama de bosque de variante VDR TaqI y urolitiasis: TT vs. Tt+tt.

FACTORES DE RIESGO NO MODIFICABLES POLIMORFISMOS DE VDR Y OSTEOPOROSIS

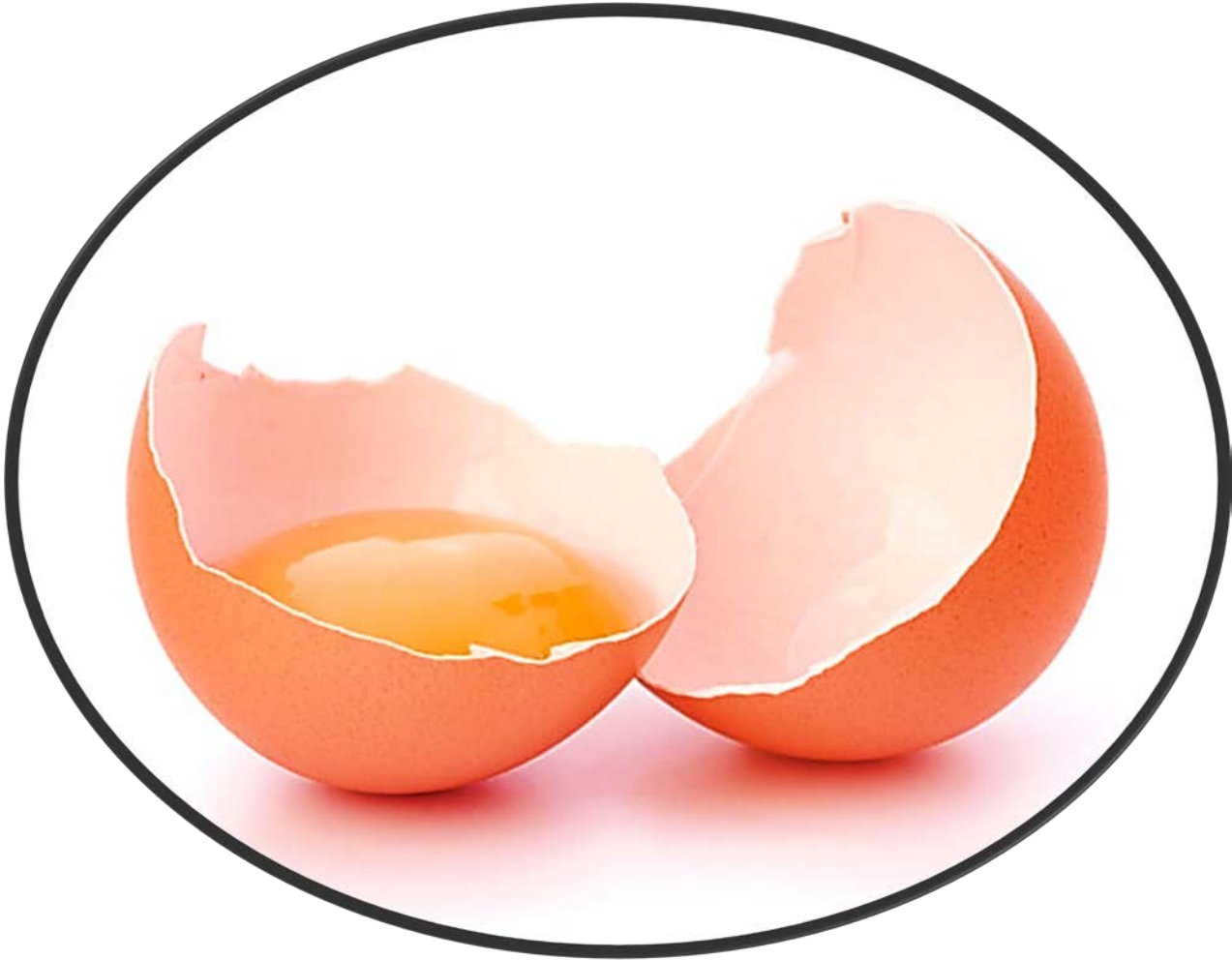
| rs731236 A > G <i>Taq1</i> | TOT | T/T | T/C | C/C | <i>p</i> ^a | OR (95% CI) |
|------------------------------|-----|-----|-----|-----|-----------------------|------------------|
| OP | 139 | 41 | 69 | 29 | 0.48 | 1.25 (0.68–2.28) |
| OPE | 54 | 16 | 26 | 12 | 0.58 | 1.24 (0.58–2.64) |
| CTR | 73 | 25 | 34 | 14 | | |
| rs2228570 C > T <i>Fok1</i> | TOT | C/C | C/T | T/T | <i>p</i> ^a | OR (95% CI) |
| OP | 135 | 54 | 62 | 23 | 0.54 | 1.25 (0.68–2.28) |
| OPE | 54 | 17 | 29 | 8 | 0.8 | 1.24 (0.58–2.64) |
| CTR | 73 | 25 | 34 | 14 | | |
| rs11568820 G > A <i>Cdx2</i> | TOT | G/G | G/A | A/A | <i>p</i> ^a | OR (95% CI) |
| OP | 139 | 78 | 52 | 9 | 0.05 | 1.81 (0.99–3.31) |
| OPE | 54 | 35 | 18 | 1 | 0.55 | 1.26 (0.6–2.66) |
| CTR | 73 | 51 | 19 | 3 | | |

p^a value represents the analysis based on the dominant model (heterozygous and homozygous variant genotypes were grouped in a single class). OR: odds ratio; CI: confidence interval. Significant associations are reported in italics.

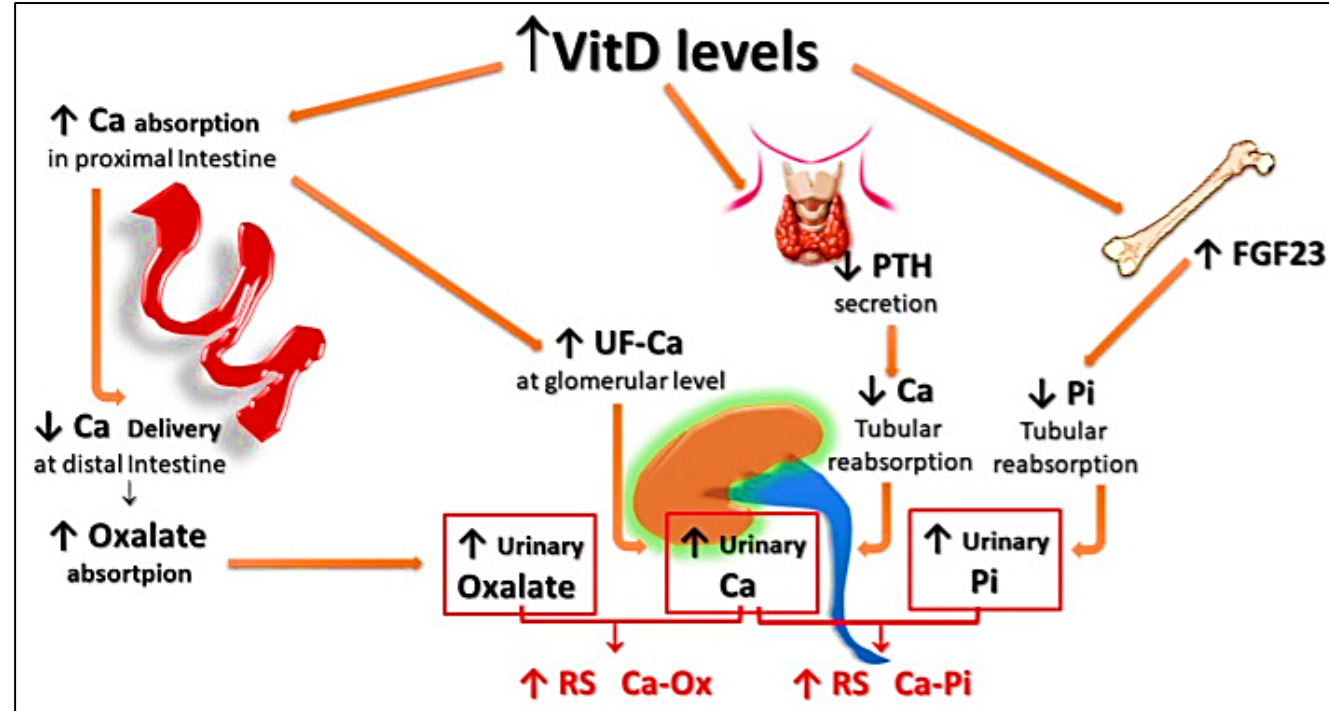
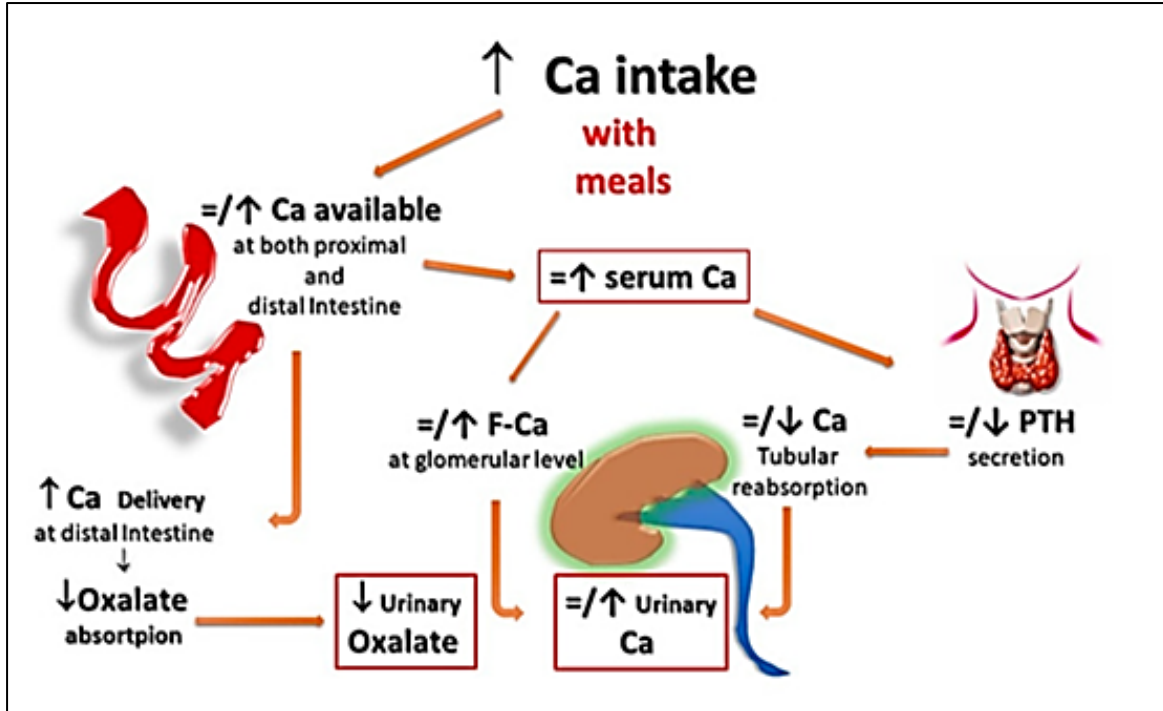


Distribución de genotipos de polimorfismos VDR

ASPECTOS CONTROVERSIALES



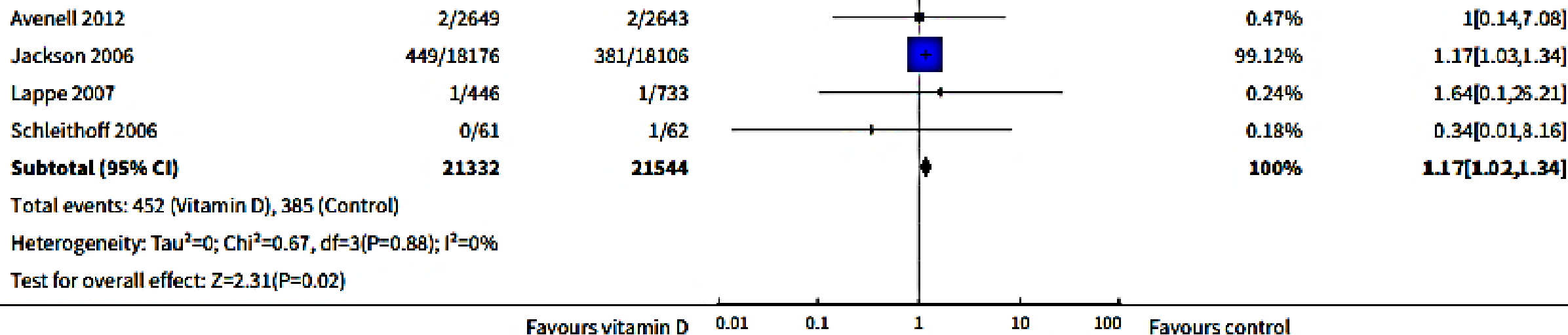
SUPLEMENTACIÓN CON VITAMINA D-CALCIO Y UROLITIASIS



Messa P. Nutrients. 2023 Mar 31;15(7):1724. doi: 10.3390/nu15071724

SUPLEMENTACIÓN CON CALCIO Y VITAMINA D/UROLITIASIS

1.26.3 Nephrolithiasis in trials using vitamin D3 combined with calcium



Bjelakovic, G. Cochrane Database Syst. Rev. 2014, 10, CD007470

SUPLEMENTACIÓN CON VITAMINA D/UROLITIASIS

| | Baseline | Follow-Up | P Value |
|------------------------------|---------------|---------------|---------|
| 24-hour urine studies | | | |
| calcium (mg) | 257±54 | 255±88 | 0.91 |
| oxalate (mg) | 42±18 | 41±15 | 0.84 |
| citrate (mg) | 696±383 | 701±320 | 0.92 |
| uric acid (g) | 0.77±0.26 | 0.77±0.20 | 0.96 |
| pH | 6.3±0.5 | 6.2±0.4 | 0.31 |
| sodium (mmol) | 228±94 | 202±65 | 0.23 |
| potassium (mmol) | 72±33 | 68±28 | 0.53 |
| phosphorus (g) | 1.0±0.3 | 1.0±0.3 | 0.91 |
| urea nitrogen (g) | 13±4 | 13±4 | 0.69 |
| sulfate (mEq) | 47±19 | 47±19 | 0.38 |
| creatinine (mg) | 1896±479 | 1896±479 | 0.74 |
| volume (L) | 1.7 (1.5–2.1) | 1.7 (1.5–2.1) | 0.50 |
| Supersaturation | | | |
| calcium oxalate | | 7±3 | 0.34 |
| calcium phosphate | | 1.2 (0.8–1.6) | 0.17 |
| uric acid | | 0.6 (0.3–0.8) | 0.20 |
| Serum studies | | | |
| 25(OH)D | 17±6 | 35±10 | <0.001 |
| calcium | 9.3±0.4 | 9.4±0.4 | 0.69 |
| PTH (pM) | 43 (27–60) | 39 (28–59) | 0.71 |

**29 sujetos con urolitiasis, con calciuria entre 150-400 mg/día.
Recibieron ergocalciferol 50.000 U semanales por 8 semanas**

Values shown are mean ± SD for normally distributed data and median (interquartile range) for skewed data. 25(OH)D, 25-hydroxyvitamin D; PTH, parathyroid hormone.

CONCLUSIONES

- La evidencia epidemiológica respalda una asociación entre urolitiasis y osteoporosis.
- Ambos trastornos comparten factores patogénicos ambientales y genéticos.
- Se podría considerar que en muchos casos la urolitiasis y la osteoporosis son expresiones de un síndrome clínico único.

RECOMENDACIONES

- Evaluación de factores de riesgo metabólicos para la nefrolitiasis en pacientes con osteoporosis.

Medición de calcio en orina de 24 horas (UK clinical guideline for the prevention and treatment of osteoporosis)

- Evaluación de DMO por DPX en pacientes con antecedentes de urolitiasis.

Considere la posibilidad de determinar densidad mineral ósea (DMO) en formadores de cálculos con evidencia de hipercalciuria y/o acidosis tubular renal distal (NE 2–3, Recomendación de grado C) (Canadian Urological Association Guideline, 2022)

- Realizar ultrasonido de abdomen después del diagnóstico de osteoporosis para evaluar la presencia de cálculos renales.

