

Epidemiología nutricional

Nutrición óptima \leftrightarrow Salud óptima



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Tema 5

- **Revisiones sistemáticas y meta-análisis.**
Planificación, diseño e interpretación.
Evidencia científica y valoración.

<https://www.ucm.es/innovadieta/nube>
<https://www.ucm.es/innovadieta/documentos-consenso>

Evidencia científica

Información contrastada para la toma de decisiones

"No hay parte de la medicina más mudable ni asentada sobre cimientos más movedizos, que la ciencia de la dietética; no pasa año que no cambie algo fundamental"

Gregorio Marañón, 1920

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Evidencia científica

Información contrastada para la toma de decisiones

- La vitamina C previene el resfriado y la gripe
- El consumo de aperitivos contribuye al incremento de peso
- El consumo de una copa de vino al día previene la ECV
- El consumo de antioxidantes aumenta la fertilidad en mujeres
-



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Evidencia científica

Información contrastada para la toma de decisiones

“Separar el grano de la paja” y sacar conclusiones

Importancia de la síntesis de la investigación

Necesidad de evidencia para práctica clínica y salud pública

¿Qué dieta es la mejor?

¿Qué le recomiendo al paciente o a la población?

Esto que dicen, ¿es cierto? ¿pueden decirlo?

¿Es cierto lo que dice esta etiqueta del lácteo?

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“Los hombres ciegos y el elefante”



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¡Demasiada información! “paradoja de la información”

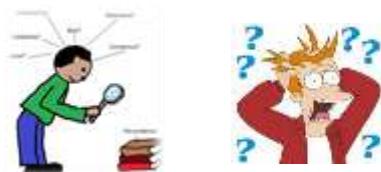
La práctica clínica tiene que estar basada en la evidencia, pero, ¿en cuál?

1940 → 2.300 revistas biomédicas

2000 → > 30.000 revistas biomédicas

2016 → Cada año se publican más de 2 millones de artículos biomédicos
(Pérez-Rodrigo, Rev Esp Comun Salud 2016, S1, S43-S51)

CÓMO ESTAR AL DÍA



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¡Demasiada información! “paradoja de la información”

- “Los profesionales de la salud se enfrentan a enormes dificultades para mantener actualizados sus conocimientos y para acceder a la información que necesitan al tomar decisiones”.
- “Los profesionales sanitarios se ven desbordados por una cantidad de información imposible de manejar e interpretar”.
- “Need to read 17 articles a day, 365 days a year!”
- Información contradictoria/no unanimidad.

(Madhukar Pai)

Sackett DL, WM Rosenberg, JA Gray, RB Haynes and WS Richardson 1996. Evidence based medicine: what it is and what it isn't. BMJ; 312:71-72. Disponible en URL: <http://bmj.bmjjournals.com/cgi/content/full/312/7023/71>

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Epidemiología nutricional. OBJETIVO FINAL

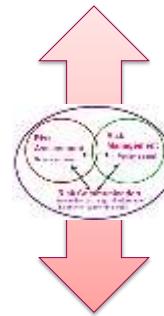


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La **evidencia científica** debe ser un marco de referencia clave para los grupos de interés, los responsables políticos, la comunidad y otras partes interesadas en el diseño, la implementación y la evaluación de **políticas públicas** para generar cambios en estilo de vida que permitan la prevención de enfermedades crónicas (Rigotti, 2013)



Los científicos producen la **Mejor evidencia posible** (*Research Epidemiology*)



Tomar decisiones: **Política nutricional → Recomendaciones dietéticas** (*Public Health Epidemiology*)

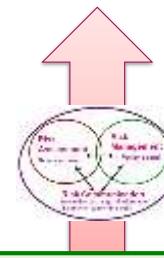
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Desarrollo de Políticas Públicas para Promoción de Estilo de Vida Saludable

Se requieren **iniciativas estratégicas colaborativas e integradas** de centros académicos, grupos privados (industria), gobiernos, grupos de interés y la comunidad, que permitan **un traspaso efectivo de la evidencia científica hacia medidas de salud pública** que promuevan hábitos de vida saludables en la población (Rigotti, 2013)



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Summary of strength of evidence on lifestyle factors and risk of developing cardiovascular diseases (WHO, 2003)				
Evidence	Decreased risk	No relationship	Increased risk	
Convincing	Regular physical activity Linoleic acid Fish and fish oils (EHA and DHA) Vegetables and fruits (including berries) Potassium Low to moderate alcohol intake (for coronary heart disease)	Vitamin E supplements	Myristic and palmitic acids Trans fatty acids High sodium intake Overweight High alcohol intake (for stroke)	
Probable	α-Linolenic acid Oleic acid NSP (fibra) Wholegrain cereals Nuts (unsalted) Plant sterols/stanols Folate	Linoleic acid	Dietary cholesterol Unfiltered boiled coffee	
Possible	Flavonoids Soy products			Los científicos producen la Mejor evidencia posible (Research Epidemiology) 
Inufficient	Calcium Magnesium Vitamin C			↑ Plant foods: wholegrain cereals, fruits and vegetables: • ≥ 400 g fruits and vegetables a day
'5 a day' programmes in Germany, Poland, Spain, Sweden, the United Kingdom, '6 a day' in Denmark and '10 a day' in France and equivalent marketing-based initiatives in other countries (e.g., '3 a day' in Hungary).				
• < 10% of daily energy intake from saturated fatty acids • < 1% of daily energy intake from trans fatty acids • < 10% of daily energy intake from free sugars • < 5 g a day of salt				

EPA, eicosapentaenoic acid; DHA, docosahexaenoic acid; NSP, non-starch polysaccharides
 WHO/FAO. Diet, Nutrition and the Prevention of Chronic Diseases. WHO Technical Report Series 916. 2003.
<http://www.fao.org/WAIRDOCS/WHO/AC911E/AC911E00.HTM>

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Resumen de la relación entre factores dietéticos y de estilo de vida y diabetes tipo 2 (WCRF, 1997)

Evidence	Decreased risk	No relationship	Increased risk
Convincing	Voluntary weight loss in overweight and obese people Physical activity	-	Overweight and obesity** Abdominal obesity*** Physical inactivity Maternal diabetes†
Probable	NSPs*	Importancia en prevención y tratamiento	-
Possible	<i>n</i> -3 Fatty acids Low glycaemic index foods Exclusive breastfeeding‡	-	Total fat intake Trans fatty acids
Insufficient	Vitamin E Chromium Magnesium Moderate alcohol	-	Excess alcohol

* NSP = Non-starch polysaccharide.

** Overweight: BMI $\geq 25 \text{ kg/m}^2$, obesity: BMI $\geq 30 \text{ kg/m}^2$.

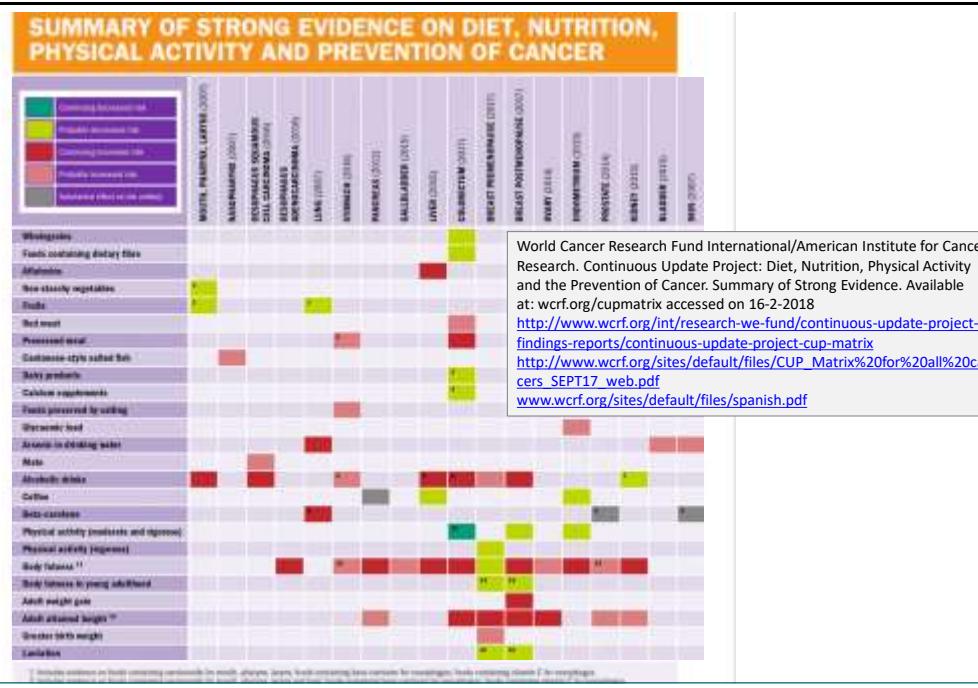
*** Waist circumference: men = 102 cm, women = 88 cm.

† This includes gestational diabetes.

‡ As a global public health recommendation, infants should be exclusively breastfed for the first six months of life to achieve optimal growth, development and health.

(Steyn y col., 2004)

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¿Cómo se llega a estas conclusiones, cuando se dispone de TANTA información y TAN contradictoria?



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Práctica Basada en la Evidencia

Hay que *cribar / filtrar* con criterios de calidad y científicos la información que se publica para hacerla accesible al profesional que tiene que tomar decisiones.

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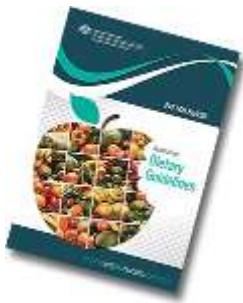
Evidencia científica

Información contrastada para la toma de decisiones

- Práctica clínica y salud pública
- Etiquetado nutricional: Aprobación de alegaciones nutricionales/salud
 - Medicina basada en la evidencia (MBE)
 - Nutrición basada en la evidencia (NuBE)
 - Práctica dietética basada en la evidencia (DNABE)

Cómo se prepara, cómo se jerarquiza, cómo interpreta, cómo se usa y aplica

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EAT FOR HEALTH - Australian Dietary Guidelines *Providing the scientific evidence for healthier Australian diets, 2013*

The guideline is designed to provide information to assist decision-making and **is based on the best available evidence at the time of development of this publication.**

<https://www.eatforhealth.gov.au/guidelines>



**2015 Dietary Guidelines for Americans
SYSTEMATIC REVIEW OF THE SCIENTIFIC EVIDENCE**

<http://health.gov/dietaryguidelines/2015-scientific-report/>

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**Revista Española de
Obesidad**

Vol. 9 • Suplemento 1 • Octubre 2011

http://www.naos.aesan.msp.es/naos/ficheros/investigacion/Consenso_SEEDO.pdf

● Recomendaciones nutricionales
basadas en la evidencia para la prevención
y el tratamiento del sobrepeso y
la obesidad en adultos
(Consenso FESNAD-SEEDO)

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Las declaraciones de propiedades saludables de los alimentos solamente pueden autorizarse después de efectuar una evaluación científica del nivel más elevado posible. Reglamento (CE) № 1924/2006 del Parlamento Europeo y del Consejo, de 20 diciembre 2006, relativo a las declaraciones nutricionales y de propiedades saludables en los alimentos.

Position of the American Dietetic Association:
Functional Foods

ADA. 2004;104(5):814-826

Functional food	Bioactive component	Health benefit	Type of evidence	Strength of evidence	Recommended amount or frequency of intake	Regulatory status
Fortified margarines	Plant sterol and stanole esters	Reduce total and LDL cholesterol (43)	Clinical trials	Very strong	1.3 g/d for sterols 1.7 g/d for stanols	Health claim
Pooyum	Soluble fiber	Reduce total and LDL cholesterol (38)	Clinical trials	Very strong	1 g/d	Health claim
Soy	Protein	Reduce total and LDL cholesterol (22-42)	Clinical trials	Very strong	25 g/d	Health claim
Whole oat products	β-glucan	Reduce total and LDL cholesterol (38)	Clinical trials	Very strong	3 g/d	Health claim

*Foods that have a Food and Drug Administration-approved health claim or intervention claim, such as protein, and generally are reported by the American Dietetic Association as having well-designed published clinical trials. For example, the low-fat claim applies to more than 40 clinical trials, whereas there are only a few clinical trials for cholesterol and coronary heart disease risk reduction.

^{a,b,c}Reprinted with permission and adapted from the American Council on Science and Health. From: *Health* (UK). J Sept. 2002;10(3):573-579.

ADA—American Dietetic Association.
FDA—Food and Drug Administration.
HFA—Health claims for functional foods.
LDL—low-density lipoprotein.
SFA—saturated fat.

2009: <http://www.eatright.org/ada/files/FunctionalFnp.pdf>

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Health Claims de avena, betaglucano y colesterol

FEEU (FDA, 21-enero-1997)

(US FDA final rule for federal labelling: health claims: oats and coronary heart disease. Fed Regist 1997;62:3584-681).
<http://www.cfsan.fda.gov/~lrd/fr970331.html>

"Una dieta alta en fibra soluble de avena integral y baja en grasa saturada y colesterol puede reducir el riesgo coronario"

- Consumo de 4 raciones diarias (0,75 g/ración: 3 g/día) → reduciría un 5% los niveles de colesterol → riesgo coronario

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ADA's Evidence Analysis Library (EAL)

Use of systematically reviewed scientific evidence in making food and nutrition practice decisions by integrating best available evidence with professional expertise and client values to improve outcomes.

Consuming diets high in total fiber (17-30 g/d) and soluble fiber (7-13 g/d) as part of a diet low in SFA and cholesterol can further ↓ TC by 2%-3%

and LDL cholesterol up to 7%

(van Horn y col., J Am Diet Assoc 2008;108:287-331)

Grade I: Good

The evidence consists of results from studies of strong design for answering the question addressed. The results are both clinically important and consistent. The results are free of serious doubts. Studies with negative results have sufficiently large sample sizes to have adequate statistical power.

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¿Cómo valorar/evaluar la evidencia científica?

- Desde finales de 1990s, cualquier procedimiento realizado en Medicina, ya sea preventivo, diagnóstico, terapéutico, pronóstico o rehabilitador, tiene que estar definido por su nivel de evidencia científica (**Medicina basada en la evidencia** o basada en las pruebas).

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Medicina basada en la evidencia (MBE)

«La aplicación consciente, explícita y juiciosa de la **mejor evidencia clínica** disponible para tomar decisiones sobre el cuidado de los pacientes y cuya práctica integra la experiencia del clínico con la mejor evidencia externa disponible procedente de una investigación sistemática»

Prof. David Sackett, 1996



Expresión utilizada por primera vez en **1991** por **Gordon Guyatt**.
En **1992** se constituyó el primer grupo de trabajo en MBE en Canadá



Sackett DL, WM Rosenberg, JA Gray, RB Haynes and WS Richardson 1996. Evidence based medicine: what it is and what it isn't. BMJ; 312:71-72. Disponible en URL: <http://bmj.bmjjournals.com/cgi/content/full/312/7023/71>

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Nutrición basada en la evidencia (NuBE)

- Aplicación sistemática de métodos científicamente rigurosos para evaluar la efectividad de las intervenciones sanitarias, tanto terapéuticas como preventivas, a nivel individual, lo que permitiría juzgar su pertinencia y decidir su aplicabilidad teniendo en cuenta las circunstancias y preferencias de los pacientes en las decisiones clínicas.
- Y, de forma implícita, también estos principios son aplicables a nivel poblacional, lo que se ha dado en llamar **Atención o Política Sanitaria Basada en la Evidencia**, mediante los que debemos valorar la tecnología, la cartera de servicios y los modelos de gestión más efectivos y eficientes, y sus resultados, y así por ejemplo las políticas alimentarias.

(Doreste y Serra, 2005)

http://www.respyn.uanl.mx/vi/2/ensayos/NuBE_Indexado.htm

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Práctica dietética basada en la evidencia (DNABE)

(desde 2009) (ICDA Evidence-based Dietetic Practice Working Group = 16 asociaciones de dietistas en el mundo)

“La práctica dietética basada en la evidencia consiste en la búsqueda sistemática de evidencia científica y la evaluación de la validez, aplicabilidad e importancia de dicha evidencia para que, combinada después con la experiencia clínica del dietista-nutricionista, las opiniones y las circunstancias y valores específicos del cliente o la comunidad, sirva de guía en la toma de decisiones en el ámbito de la dietética”

Además, la práctica dietética basada en la evidencia:

- se utiliza para tomar decisiones en todas las áreas de la práctica dietética con la finalidad de mejorar la salud de clientes (y pacientes), comunidades y poblaciones.
- establece claramente la fuente de la evidencia que sustenta las recomendaciones prácticas. Para ser pertinente y efectiva, la práctica dietética basada en la evidencia debe integrar el conocimiento de otras disciplinas.
- está fundamentada en los principios éticos y códigos de buena práctica. Esto incluye la necesidad de reflexionar acerca de cómo las perspectivas o sesgos personales pueden influenciar la interpretación de la evidencia científica.

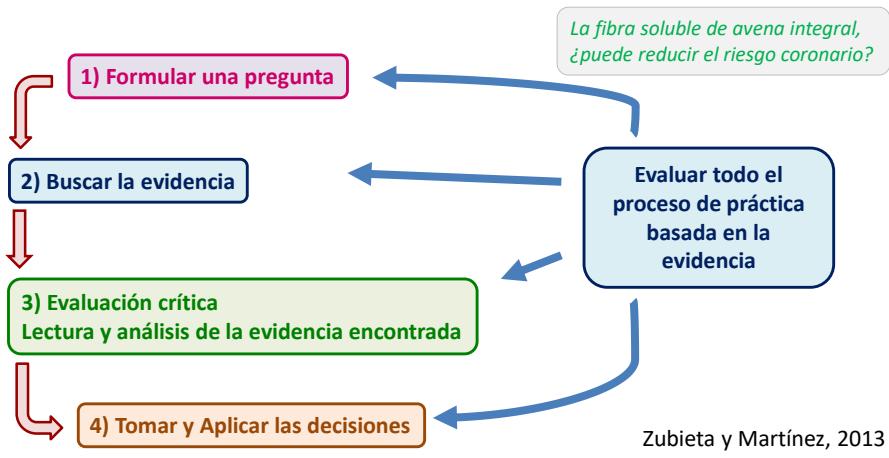
Approved by the ICDA Board of Directors, November 13, 2010

<http://www.grep-aedn.es/newsletter/diciembre2010.htm>

<http://www.internationaldietetics.org/Downloads/ICDA-Report-Evidence-based-Dietetics-Practice-2010.aspx>

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Etapas de la práctica basada en la evidencia



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¡Demasiada información!
“paradoja de la información”

Estado de la cuestión, state of the art

- 1) Revisiones bibliográficas narrativas o clásicas
- 2) Revisiones sistemáticas
- 3) Meta-análisis
- 4) *Umbrella reviews: evidence synthesis with overviews of reviews and meta-epidemiologic studies*

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1) Revisiones bibliográficas narrativas o clásicas

(Traditional, Narrative Review, non-systematic or selective reviews)

2 o más publicaciones

No hay pregunta ni "material y métodos"

Usan métodos «informales», no explícitos y a menudo personales y subjetivos

No se especifica el proceso seguido para buscar, reunir y evaluar la información para sacar conclusiones y, sin esta información, no será posible repetir y verificar los resultados y las conclusiones de la revisión.

*J Am Coll Nutr. 1995 Apr;14(2):124-36.
The health effects of vitamin C supplementation: a review.
Baranick A, Lantzelli L.
Hoffman-La Roche Inc., Paramus, New Jersey 07652, USA.
E-mail to: [REDACTED]
J Am Coll Nutr 1995 Aug;14(4):380.
J Am Coll Nutr 1995 Jun;14(3):258.*

Abstract
A comprehensive review of the literature indicates that populations with long-term consumption of higher than RDA levels of vitamin C (\geq or = 60 mg/day) from foods and/or supplements have reduced risks of cancer at several sites, cardiovascular disease, and cataracts. The safety of higher than RDA intakes of vitamin C is confirmed in eight placebo-controlled, double-blind studies and six non-placebo clinical trials in which up to 10,000 mg of vitamin C was consumed daily for up to 3 years. There are no clinical data which suggest that vitamin C's enhancement of non-heme iron absorption in individuals with low iron status could be a critical factor in the possible increased risk of heterozygous hemochromatosis-related cardiovascular disease. In fact, the cumulative data do not confirm that iron status is related to risk of cardiovascular disease. Moreover, higher than RDA intakes of Vitamin C have been associated with several indices of lowered cardiovascular disease risk including increases in HDL, and decreases in LDL oxidation, blood pressure and cardiovascular mortality.



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2) Revisiones sistemáticas

(Overview, Systematic literature review)

Trabajo de investigación realizado por expertos que revisa la evidencia científica sobre una pregunta claramente definida.

Usa métodos sistemáticos y explícitos para identificar, seleccionar y evaluar críticamente estudios relevantes y extraer y analizar datos de interés para obtener conclusiones consistentes (The Centre for Review and Dissemination).

Mulrow, C. D. (1994). Systematic reviews: Rationale for systematic reviews. BMJ, 309(6954):597-599.



Primera revisión sistemática en el área de nutrición fue publicada en 1953:

Stewart CP & Guthrie P (editors) (1953)

Lind's Treatise on Scurvy.

A bicentenary volume containing reprint on the first Edition of a Treatise of the Scurvy, by James Lind, M. D. with additional notes, p. 314. Edinburgh: Edinburgh University Press.

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2) Revisiones sistemáticas

(Overview, Systematic literature review)

Pasos básicos en la realización de revisiones sistemáticas

1. Especificación del problema. Definición de las preguntas.
2. Especificación de los criterios de inclusión y exclusión de los estudios
3. Formulación del plan de búsqueda de la literatura.
4. Selección y recuperación de los estudios que cumplen los criterios.
5. Valoración crítica de la calidad científica de los mismos.
6. Combinación de los resultados.
7. Formulación de las conclusiones y recomendaciones.

http://www.respyn.uanl.mx/vi/2/ensayos/NuBE_Indexado.htm

Guerra y col., 2003 - <http://www.fisterra.com/>

Systematic Literature Review



Fig. 1. Systematic literature review process
Tiago Silva da Silva (ICT/UNIFESP)

Steps of systematic review:



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2) Revisiones sistemáticas

(Overview, Systematic literature review)

Pretenden ser:

- a) **Rigurosas**, en cuanto a los estudios incluidos (con criterios de calidad, etc.)
- b) **Informativas**, enfocadas hacia **problemas reales**, tratando de contestar una **pregunta claramente delimitada o específica**, e idealmente analizando y presentando los datos de la forma que mejor **ayude a la toma de decisiones**
- c) **Exhaustivas**: su objetivo es identificar y utilizar la **mayor cantidad posible de información pertinente**, sin introducir sesgos (de publicación, de selección, etc.)
- d) **Explícitas**, ya que todos los métodos utilizados en la revisión deben describirse con suficiente detalle.

(Gisbert y Bonfill, 2004)

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Importante: “explicit methods section”

REVIEW ARTICLE

A Systematic Review of the Evidence Supporting a Causal Link Between Dietary Factors and Coronary Heart Disease

Andrew Mente, PhD; Lawrence de Koning, MSc; Hurry S. Shunum, PhD; Senia S. Anand, MD, FRCR, FRCPC

Background: Although a wealth of literature links dietary factors and coronary heart disease (CHD), the strength of the evidence supporting valid associations has not been evaluated systematically in a single investigation.

Methods: We conducted a systematic search of MEDLINE for prospective cohort studies or randomized trials investigating dietary exposures in relation to CHD. We used the Bradford Hill guidelines to derive a causation score based on 4 criteria (strength, consistency, temporality, and coherence) for each dietary exposure in cohort studies and examined for consistency with the findings of randomized trials.

Results: Strong evidence supports valid associations (4 criteria satisfied) of protective factors, including intake of vegetables, nuts, and “Mediterranean” and high-quality dietary patterns with CHD, and associations of harmful factors, including intake of trans-fatty acids and foods with a high glycemic index or load. Among studies of higher methodologic

quality, there was also strong evidence for monounsaturated fatty acids and “prudent” and “western” dietary patterns. Moderate evidence (3 criteria) of associations exists for intake of fish, marine-ω-3 fatty acids, folate, whole grains, dietary vitamins E and C, beta carotene, alcohol, fruit, and fiber. Insufficient evidence (<2 criteria) of association is present for intake of supplementary vitamin E and ascorbic acid (vitamin C); saturated and polyunsaturated fatty acids; total fat; α-linolenic acid; meat; eggs; and milk. Among the dietary exposures with strong evidence of causation from cohort studies, only a Mediterranean dietary pattern is related to CHD in randomized trials.

Conclusions: The evidence supports a valid association of a limited number of dietary factors and dietary patterns with CHD. Future evaluation of dietary patterns, including their nutrient and food components, in cohort studies and randomized trials is recommended.

Arch Intern Med. 2009;169(7):659-669

SEARCH STRATEGY AND STUDY SELECTION

We searched the MEDLINE database for prospective cohort studies and RCTs from 1950 through June 2007. The bibliographies of retrieved articles were scanned for additional cohort studies and RCTs. Two of us (A.M. and L.D.J.) independently assessed study eligibility. Excluded studies and reasons for exclusion were listed, and disagreement was resolved by discussion and consensus. We included original English-language articles pertaining to the effect of diet on the following primary outcomes: coronary or atherosclerotic heart disease and fatal or non-fatal myocardial infarction. These articles were also evaluated for the following secondary outcomes: angina pectoris, sudden death, cardiovascular disease, and total mortality. Relative risks (RRs) of outcomes are presented with their 95% confidence intervals (CIs) after adjusting for potential confounders. We only considered studies that followed up subjects for at least 1 year. Cohort studies had to include estimates of dietary intake using conventional dietary assessment tools (eg, food frequency questionnaire, food records, or 24-hour diet recall). Clinical trials had to be randomized and compare dietary exposure with a control diet or a placebo. Cross-over trials were excluded if plasma biomarkers or adrenocortical measures were not evaluated because carryover outcomes occurring after a crossover would be difficult to interpret.

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Cochrane Collaboration



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Summerbell y Moore, 2007

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<http://uk.cochrane.org/news/what-are-systematic-reviews>

<https://ccrcg.cochrane.org/animated-storyboard-what-are-systematic-reviews>

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Uso de la evidencia en políticas sanitarias internacionales: la colaboración entre Cochrane y la OMS

<http://es.cochrane.org/es/news/uso-de-la-evidencia-en-pol%C3%ADticas-sanitarias-internacionales-la-colaboraci%C3%B3n-entre-cochrane-y-la>

2. The collaboration has contributed to the WHO Programme of Work in the following (below): eLENA:

Category of Work: Non-communicable diseases
Programme Area: Nutrition
Outcome target/s: Reduced nutritional risk factors
Short description of the collaboration:



WHO e-Library of Evidence for Nutrition Actions (eLENA) is an online library of evidence-informed guidance for nutrition interventions. It is a single point of reference for the latest nutrition guidelines and related information including supporting materials such as scientific evidence, background materials and commentaries from invited experts.

Cochrane has contributed to the development of eLENA since its launch in 2011. Thanks to an agreement with the publishers of the Cochrane Library, John Wiley & Sons, Ltd, eLENA users have access to the full reviews and are also able to track the use of reviews in guideline development processes⁴.



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Uso de la evidencia en políticas sanitarias internacionales: la colaboración entre Cochrane y la OMS

<http://es.cochrane.org/es/news/uso-de-la-evidencia-en-pol%C3%ADticas-sanitarias-internacionales-la-colaboraci%C3%B3n-entre-cochrane-y-la>



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Al contribuirse se obtienen los recursos de las evidencias, que facilitan el uso específico de OMS para cada estrategia generada por Cochrane.

- Actualizado el año pasado, se han utilizado 410 revisiones del MUS en las estrategias de salud pública para documentar 144 directrices de la OMS y otras recomendaciones basadas en las evidencias más altas entre 2009 y junio de 2013.
- La estrategia del Comité de Normas de Documentos organiza regularmente talleres y reuniones sobre otras temáticas importantes para la elaboración de directrices. De 2012 a 2014, 20 de Jun. 30 reuniones y 1000 participantes a representantes de la Ejecución, Comisión y Poder Ejecutivo.
- Los contribuyentes a la Línea de Recomendaciones Esenciales (LRE) y a la revisión de la LRE también utilizan evidencias Cochrane. Hay más de 1000 artículos revisados que, en 2011, ya de cuatro 15 editores utilizaban evidencias Cochrane. En 2013 se publicaron 7400 artículos.
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Selenium supplements for the prevention of cardiovascular disease

Alvarez K, Hartley L, Dey C, Flatters R, Clarke A, Stranges S
Published Online: January 31, 2013

This summary was produced using rigorous methods by the impartial and independent Cochrane Collaboration

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Cranberries for preventing urinary tract infections

Jepson RG, Williams G, Craig JC | Published Online: October 17, 2013

Cranberries (usually as cranberry juice) have been used to prevent urinary tract infections (UTIs). Cranberries contain a substance that can prevent bacteria from sticking on the walls of the bladder. This may help prevent bladder and other UTIs. This review identified 24 studies (4473 participants) comparing cranberry products with control or alternative treatments. There was a small trend towards fewer UTIs in people taking cranberry product compared to placebo or no treatment but this was not a significant finding. Many people in the studies stopped drinking the juice, suggesting it may not be an acceptable intervention. Cranberry juice does not appear to have a significant benefit in preventing UTIs and may be unacceptable to consume in the long term. Cranberry products (such as tablets or capsules) were also ineffective (although have the same effect as taking antibiotics), possibly due to lack of potency of the active ingredient.

- Abstract (click to read)

Background:
Cranberries have been used widely for several decades for the prevention and treatment of urinary tract infections (UTIs). This is the third update of our review first published in 1998 and updated in 2004 and 2008.

Objectives:
To assess the effectiveness of cranberry products in preventing UTIs in susceptible populations.

This summary was produced using rigorous methods by the **independent and independent** Cochrane Collaboration.

Find the research
Get full text in The Cochrane Library for this Review (PDF):
Cranberries for preventing urinary tract infections

Primary Review Group: Renal Group



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Perlas Cochrane: Evidencia práctica para situaciones de la vida real

Breves resúmenes de revisiones sistemáticas orientadas a problemas atendidos en el ámbito de la atención primaria.

- Abstract (click to read)

- PEARLS - Practical Evidence About Real Life Situations (click to read)

Clinical question:
How effective are cranberry products in preventing urinary tract infections (UTIs) in susceptible populations?

Bottom line:
There was some evidence that cranberries (juice or capsules) may decrease the number of symptomatic UTIs over a 12-month period, particularly for women with recurrent UTIs (NNT=7). The evidence for elderly men and women was less clear, and there was evidence cranberry products were not effective in people who needed either intermittent or indwelling catheters. *NNT = number needed to treat to benefit 1 individual.

Response:
Caveat:
Many people in the trials stopped drinking the juice, suggesting it may not be a popular intervention. It is not clear how long cranberry juice needs to be taken to be effective or what the required dose might be.

Context:
No definite mechanism of action has been established for cranberries in the prevention or treatment of UTI. However, the main suggestion is cranberries prevent bacteria, particularly *Escherichia coli*, from adhering to uroepithelial cells lining the bladder. Without adhesion, *E. coli* cannot infect the mucosal surface of the urinary tract.

Cochrane Systematic Review:
Jepson RG and Craig JC. Cranberries for preventing urinary tract infections. Cochrane Reviews 2006, Issue 1. Article No. CD001321.
DOI: 10.1002/14651858.CD001321.pub4. This review contains 18 trials involving 1649 participants.

Authored by:
Brian R McAvay
Cochrane Primary Health Care Field

<http://www.cochraneprimarycare.org/pearls-2012-254-296>
<http://summaries.cochrane.org/CD001321/cranberries-for-preventing-urinary-tract-infections>

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3) Meta-análisis

Tipo de revisión sistemática que usa métodos estadísticos para obtener un estimado único (un riesgo relativo, una diferencia de riesgos) a partir de los resultados de los estudios independientes.

- Procedimiento estadístico que integra los resultados de estudios independientes pero con un diseño y objetivos similares.
- Proceso de recopilación y combinación de información de diversos estudios relacionados con el propósito de llegar a una conclusión.

Se consigue:

- Una mayor potencia estadística para detectar diferencias.
- Una estimación más precisa del efecto global.
- Una técnica claramente superior y más objetiva que la revisión bibliográfica tradicional.

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3) Meta-análisis

1904

Karl Pearson (matemático británico)

1976

Glass GV (Psicólogo)

Introduce el término de Meta-análisis

1970-1980

Se redescubre para la medicina (ECV, cáncer)

Tom Chalmers (EEUU)

Richard Peto (RU)

Iam Chalmers (RU)

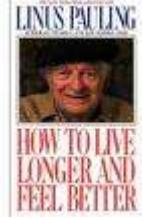
1990

Meta-análisis: Término incluido como “Medical subject heading” en el sistema de indexación de MedLine de la National Library of Medicine
Cochrane Centre

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Linus Pauling (1901-1994)

1954. Premio Nobel de Química
1962. Premio Nobel de la Paz



How to live longer and feel better
Vitamina C, resfriado común y gripe
Cáncer y vitamina C, 1970.



Portuguese edition of Vitamin C and the Common Cold, a book that was translated into nine different languages.

Linus Pauling; José María Ladero Quesada (trad.)
Madrid : AC, 1980; ISBN: 84-7288-202-0

Propone que el consumo de
1.000 mg/día de vitamina C reduce la
incidencia del resfriado común en un 45%.
(IDR: 60 mg/día)

USA: production of vitamin C increased from 8.9 million pounds in 1969 to 11.7 in 1971 (increase of 39% in two years and an annual growth rate of 18%) (CMR 1972a), in contrast to the annual growth rate of about 6% in the 1960s (CMR 1972c). About 5.6 million pounds of vitamin C were also imported to the US between January and November 1971, up about 160% from the amount imported during the same period in 1970 (CMR 1972b).

<http://paulingblog.wordpress.com/2010/11/09/vitamin-c-the-common-cold-and-controversy/>
<http://www.ltdk.helsinki.fi/users/hemila/reviews/pauling/>

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Proc. Am. Acad. Sci. USA
Vol. 68, No. 11, pp. 3024-3030, December 1971

The Significance of the Evidence about Ascorbic Acid and the Common Cold

(vitamin C/ascorbic acid/statistical significance)
L20108 PAULING

Department of Chemistry, Stanford Research Institute, Menlo Park, California 94036

Translated by Goro Terada, Japan, 6, 1971

ABSTRACT (This) has been reported that ascorbic acid has value in preventing protein against the common cold and in reducing the manifestations of the cold disease. This popular belief has, however, not been generally shared by physicians, nutritionists, scientists, and other bodies.

I was prompted to re-examine the evidence behind this belief and to perform, as I did, the published research reported thereof as well, in relation to the common cold. On the basis of this study and of some general arguments about the nature of the common cold, I have drawn conclusions and the results thereof by carrying the conclusions from the human body of substances that are normally present in the body and are required for health and the prevention of disease. These substances are, in other words, those in the human nutrients, chemicals, the breakdown products and secretions of the body. In my book, "Vitamin C and the Common Cold" (1), the evidence will be presented in this book with reference to existing literature, physicians and authorities on nutrition. Many references concerning my conclusions were made.

ascorbate can in most cases kill, and those in the most potent form were administered at apparently therapeutic (optimal), i.e., highest. The studies were double-blind, with one of the subjects receiving the placebo and the other receiving the ascorbate and which received the placebo. But knowledge being keys to some other persons and all, the information had been released.

This paper is intended to examine by analyzing the published reports to the following. Then the major characteristics of placebo and at a dose greater than 100 mg per day, and the results of the placebo and the active drug, i.e., ascorbate, in preventing the common cold, and with the outcome reported in only those under ordinary living conditions, both as often different from that of a patient as among the literature on the prevention of the common cold. A second point with a placebo with the subjects not knowing which group they are in, is essential because of the well-known "placebo effect" of even inactive substances.

The statistical methods used in this analysis are the conventional ones, for the most part Student's *t*-test or the calculation of χ^2 and that of the probability. Probability is the chance of obtaining a result as extreme as, or more extreme, than that actually observed if the null hypothesis or a large difference would be obtained by chance alone. If the two groups taken at random from a healthy population, if the differences between them are not statistically significant, then the null hypothesis is accepted. If the differences are statistically significant, then the null hypothesis is rejected. That is, they choose to give (from their) either more than (or less than) because we are interested that the placebo (usually costs more) has a greater effect. In this case, the difference of opinion is between those people who think that ascorbic acid is better than a placebo and those who say that it is a better placebo. My conclusion is that the evidence does not indicate that a greater preventive effect of the placebo than of ascorbic acid in most individuals used is reported to

Pauling (1971) carried out a meta-analysis of 4 placebo-controlled trials, which was one of the very first meta-analyses in medicine.

of four groups of subjects selected at random from one population. The subjects in every group were administered the active substance: ascorbic acid and vitamin C in various

amounts, in amounts were 200 mg per day, and placebo, which was taken regularly over a period of time, beginning before colds were learned. The outcome will be considered to be the disease.

<http://www.ncbi.nlm.nih.gov/pmc/articles/PMC389499/?page=1>

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1971

Segundo metaanálisis

Linus Pauling, Ph.D.

The American Journal of Clinical Nutrition

For a number of years I have been interested in the possibility that the state of health of many people could be significantly improved by the ingestion in the optimum amounts of certain substances normally present in the human body, including the vitamins. This interest developed from the work that my associates and I have done on molecular diseases, especially the hemoglobinopathies (1). I decided in 1953 that it would be worthwhile to make a study of the extent to which mental diseases could be described as molecular diseases. Work along these lines was carried out in our laboratory in the California Institute of Technology from 1954 to 1964, and was continued in the University of California, San Diego, and (since 1969) in Stanford University. In the course of this period I formulated some ideas about orthomolecular medicine, defined as the preservation of good health and the treatment of disease by varying the concentrations in the human body of substances that are normally present in the body and are required for health (2-4). I also became aware of arguments indicating that the optimum rate of intake of ascorbic acid may be far greater than the recommended daily allowance of this vitamin, which is approximately 50 mg/day.

For example, as recently as November 1970, Dr. Philip L. White (10), Secretary of the Council on Foods and Nutrition of the American Medical Association, stated that "Unfortunately, it is still a widespread belief that extra ascorbic acid can not only prevent colds but also lessen the severity and duration of colds and other respiratory infections. Even when consumed at the first sign of a sniffle, large doses of the vitamin are useless." Also, many statements contradicting my conclusions were made by physicians, experts in nutrition, and health officials within a few weeks after the publication of my book. For example, Dr. Charles C. Edwards, United States Food and Drug Commissioner, was reported in the press on December 29, 1970 as having said that the use of ascorbic acid was ridiculous, and that there was no scientific evidence and never had been any meaningful studies indicating that vitamin C is capable of preventing or curing colds. The Editors of *The Medical Letter* published an article in which nearly all my statements were contradicted; for example, it was stated that there had been no controlled trials of the effectiveness of vitamin C, in comparison with a placebo, against upper respiratory infections over a long period and including many hu-

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ANÁLISIS DE 30 TRABAJOS

El falso mito de la vitamina C para el resfriado

- Este micronutriente acorta los catarros y alivia los síntomas, pero no los previene
- En el caso de maratonianos o esquiadores se reduce un 50% el número de resfriados

Investigación online (2017/2018) de la UCI

ISABEL F. LANTIGUA

Madrid.- Durante más de 60 años el papel de la vitamina C para prevenir los resfriados comunes ha sido objeto de controversia. Una última revisión de más de 30 ensayos en los que han intervenido 11.350 participantes carga la cuestión al afirmar que "hay claras evidencias que muestran que no tiene sentido tomar suplementos de vitamina C todo el año, ya que no reduce la incidencia de los catarros".

Aunque esta sustancia es muy popular entre la gente para no "pollar" un resfriado "hence comprado durante varias décadas que no es la panacea", afirma Ham Haméla, del departamento de Salud Pública en la Universidad de Helsinki (Finlandia) y uno de los autores de esta revisión que se publica en el último número de *"The Cochrane Library"*, una publicación que revisa las evidencias científicas sobre un tema.

Los 30 ensayos analizados muestran que la utilización de suplementos de vitamina C todos los días —una práctica bastante extendida— para prevenir los catarros "no está justificada, puesto que este efecto no se ha comprobado en la población general". Sin embargo, Haméla explica que "si tenemos un ejercicio intenso o una infección viral, acorta la duración de los resfriados y alivia sus síntomas".

(Foto: José María Pérez)

Ángeles Carbajal Azcona. Dpto de Nutrición y Ciencia de los Alimentos. Fac. Farmacia. Universidad Complutense de Madrid. <https://www.ucm.es/nutricioncarbajal>

**Vitamin C for preventing and treating the common cold
(Review)**

Hemill H, Chalker E, Douglas B

2010



The COCHRANE
COLLABORATION®

This is a report of a Cochrane review, prepared and maintained by The Cochrane Collaboration and published in The Cochrane Library.
2010, Issue 1
<http://www.thecochranelibrary.com>

Ángeles Carbajal Azcona. Dpto de Nutrición y Ciencia de los Alimentos. Fac. Farmacia. Universidad Complutense de Madrid. <https://www.ucm.es/nutricioncarbajal>

AUTHORS' CONCLUSIONS

Implications for practice

The lack of effect of prophylactic vitamin C supplementation on the incidence of the common cold in the general population throws doubt on the usefulness of this practice. In special circumstances, where people are engaged in extreme physical exertion or exposed to significant cold stress, or both, vitamin C supplementation may have a beneficial effect, but caution should be exercised in generalising this finding. The prophylaxis trials found a reduction in common cold duration of 8% in adults and 13% in children. The practical relevance of these findings is open. In our opinion, this level of benefit does not justify long-term prophylaxis in its own right. So far, therapeutic supplementation has not been shown to be effective. Nevertheless, given the consistent effect of vitamin C on duration and severity in the regular supplementation studies, and the low cost and safety, it may be worthwhile for common cold patients to test on an individual basis whether therapeutic vitamin C is beneficial for them.

Implications for research

It does not seem worthwhile to carry out further regular supplementation trials in the general population. However, the findings in marathon runners, skiers and soldiers operating in subarctic conditions warrant further research.

None of the therapeutic trials carried out so far have examined the effect of vitamin C on children, even though the regular supplementation trials have found substantially greater effect on cold duration in children than in adults. In view of the greater incidence of respiratory infections in children, such therapeutic trials are warranted.

The findings in the Anderson 1974 study on the greater benefit of 8 g than 4 g dose on the day of onset of respiratory symptoms suggest that doses in further therapeutic trials with adults should be at least 8 g/day.

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Revisado, 2012

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Hans Hemila^{1,*}, Elizabeth Chalker²
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Authors' conclusions

The failure of vitamin C supplementation to reduce the incidence of colds in the general population indicates that routine vitamin C supplementation is not justified, yet vitamin C may be useful for people exposed to brief periods of severe physical exercise. Regular supplementation trials have shown that vitamin C reduces the duration of colds, but this was not replicated in the few therapeutic trials that have been carried out. Nevertheless, given the consistent effect of vitamin C on the duration and severity of colds in the regular supplementation studies, and the low cost and safety, it may be worthwhile for common cold patients to test on an individual basis whether therapeutic vitamin C is beneficial for them. Further therapeutic RCTs are warranted.

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BMJ RESEARCH

Sofi et al., BMJ 2008;337:a1344
Sofi F et al. Am J Clin Nutr 2010;92:1189-1196
Sofi et al., Public Health Nutr. 2014 Dec;17(12):2769-82

Adherence to Mediterranean diet and health status: meta-analysis

Conclusiones: una mayor adherencia a la Dieta Mediterránea se asocia con una mejora de la salud, con reducciones significativas de:

- Mortalidad total (9%),
- Mortalidad cardiovascular (9%),
- Incidencia y mortalidad de cáncer (6%), y
- Incidencia de Parkinson y Alzheimer (13%).

Resultados clínicamente relevantes para la salud pública. Sería recomendable fomentar la Dieta Mediterránea para prevención primaria y secundaria de las principales enfermedades crónicas.

Salud pública

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Características de la dieta Mediterránea tradicional (Mediterranean diet score)

1. Alta relación AGM/AGS
2. Consumo moderado de alcohol (vino)
3. Alto de leguminosas
4. Alto de cereales (integrales y pan)
5. Alto de frutas
6. Alto de verduras y hortalizas
7. Bajo de carnes y derivados
8. Moderado de leches y derivados
9. Alto consumo de pescados

Óptimo = score de 9

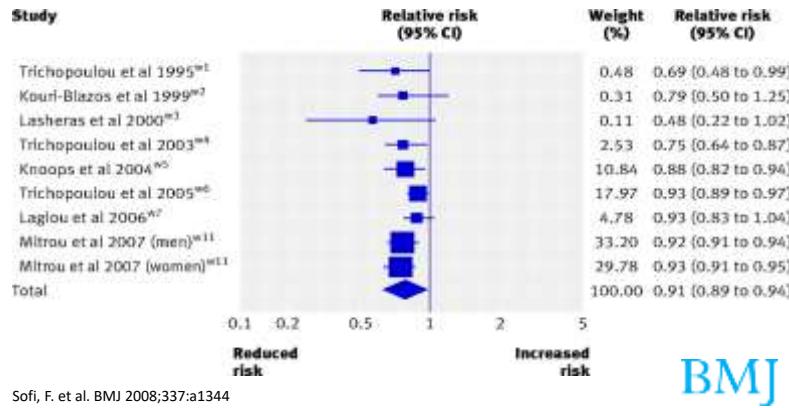
(Trichopoulou y col., 2000)



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Fig 2 Risk of all cause mortality associated with two point increase in adherence score for Mediterranean diet

Squares represent effect size; extended lines show 95% confidence intervals; diamond represents total effect size



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4) Umbrella reviews: evidence synthesis with overviews of reviews and meta-epidemiologic studies

Coffee, Caffeine, and Health Outcomes: An Umbrella Review.

Grosso G^{1,2}, Godas J^{1,3}, Galvao F³, Giovannucci E^{4,5,6}

Abstract

To evaluate the associations between coffee and caffeine consumption and various health outcomes, we performed an umbrella review of the evidence from meta-analyses of observational studies and randomized controlled trials (RCTs). Of the 59 unique outcomes examined in the selected 112 meta-analyses of observational studies, coffee was associated with a probable decreased risk of breast, colorectal, colon, endometrial, and prostate cancers; cardiovascular disease and mortality; Parkinson's disease; and type-2 diabetes. Of the 14 unique outcomes examined in the 20 selected meta-analyses of observational studies, caffeine was associated with a probable decreased risk of Parkinson's disease and type-2 diabetes and an increased risk of pregnancy loss. Of the 12 unique acute outcomes examined in the selected 9 meta-analyses of RCTs, coffee was associated with a rise in serum lipids, but this result was affected by significant heterogeneity, and caffeine was associated with a rise in blood pressure. Given the spectrum of conditions studied and the robustness of many of the results, these findings indicate that coffee can be part of a healthful diet.

KEYWORDS: caffeine; cancer; cardiovascular disease; coffee; diabetes; neurodegenerative disease

PUBMED ID: 28826374 DOI: 10.1146/annurev-nutr-071016-064941

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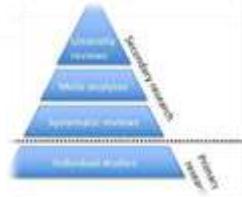
Statistics in practice

Ten simple rules for conducting umbrella reviews FREE

Paolo Fusar-Poli^{1, 2, 3}, Joaquim Radua^{1, 4, 5}

Author affiliations +

<https://ebmh.bmjjournals.org/content/21/3/95>



Abstract

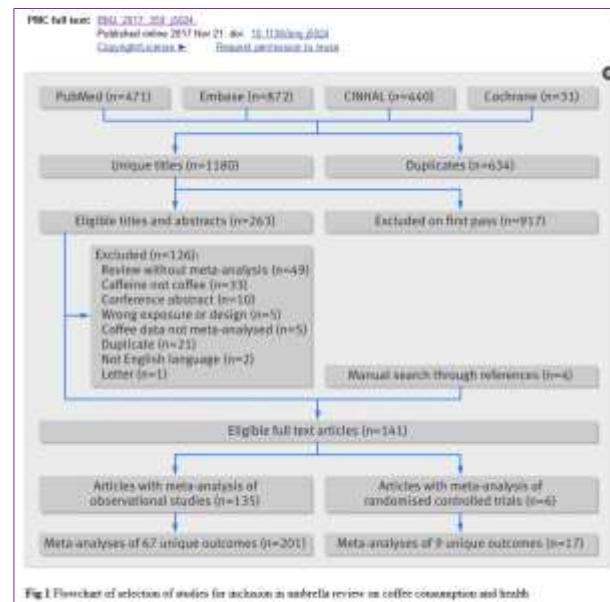
Objective Evidence syntheses such as systematic reviews and meta-analyses provide a rigorous and transparent knowledge base for translating clinical research into decisions, and thus they represent the basic unit of knowledge in medicine. Umbrella reviews are reviews of previously published systematic reviews or meta-analyses. Therefore, they represent one of the highest levels of evidence synthesis currently available, and are becoming increasingly influential in biomedical literature. However, practical guidance on how to conduct umbrella reviews is relatively limited.

Methods We present a critical educational review of published umbrella reviews, focusing on the essential practical steps required to produce robust umbrella reviews in the medical field.

Results The current manuscript discusses 10 key points to consider for conducting robust umbrella reviews. The points are: ensure that the umbrella review is really needed, prespecify the protocol, clearly define the variables of interest, estimate a common effect size, report the heterogeneity and potential biases, perform a stratification of the evidence, conduct sensitivity analyses, report transparent results, use appropriate software and acknowledge the limitations. We illustrate these points through recent examples from umbrella reviews and suggest specific practical recommendations.

Conclusions The current manuscript provides a practical guidance for conducting umbrella reviews in medical areas. Researchers, clinicians and policy makers might use the key points illustrated here to inform the planning, conduction and reporting of umbrella reviews in medicine.

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<https://www.ncbi.nlm.nih.gov/pubmed/28826374>

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<https://www.ncbi.nlm.nih.gov/pubmed/28826374>

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The screenshot shows the homepage of the NMCD (Nutrition, Metabolism & Cardiovascular Diseases) journal. At the top, there's a navigation bar with links for 'Articles & issues', 'Collections', 'For Authors', 'Journal Info', 'Subscribe', 'Society Information', and 'More Periodicals'. Below the navigation is a search bar with dropdowns for 'All Content' and 'Search'.

The main content area displays an article abstract:

Mediterranean diet and multiple health outcomes: An umbrella review of meta-analyses of observational studies and randomized trials

By M. Díaz, G. Padias, A. Casas, F. Sofi

PainX Metrics

DOI: <https://doi.org/10.1016/j.nmcd.2016.11.055>

To the right of the abstract is a sidebar titled 'Article Tools' containing links for PDF download, email, add to reading list, export citation, create citation alert, and cite by in Scopus. There's also a link to 'Order Reprints'.

Below the abstract, a URL is provided: [http://www.nmcd-journal.com/article/S0939-4753\(16\)30247-2/fulltext](http://www.nmcd-journal.com/article/S0939-4753(16)30247-2/fulltext)

At the bottom of the page, a footer note reads: 'Ángeles Carbajal Azcona. Dpto de Nutrición y Ciencia de los Alimentos. Fac. Farmacia. Universidad Complutense de Madrid. <https://www.ucm.es/nutricioncarbajal>'

¿Cuándo hacer un meta-análisis?

- 1) **Acumular evidencia** sobre una situación ya documentada en distintos trabajos que individualmente no tienen suficiente peso estadístico para poder tomar decisiones sólidamente fundadas.
- 2) **Aclarar resultados** de trabajos contradictorios.
- 3) **Resumir resultados** cuando hay información muy abundante.
- 4) Para **justificar la necesidad** de realizar ensayos de mayor tamaño que los realizados hasta ese momento o, por el contrario, para justificar que no es necesario gastar más tiempo y recursos en estudios de mayor tamaño.

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AUTHORS' CONCLUSIONS**Implications for practice**

The lack of effect of prophylactic vitamin C supplementation on the incidence of the common cold in the general population throws doubt on the usefulness of this practice. In special circumstances, where people are engaged in extreme physical exertion or exposed to significant cold stress, or both, vitamin C supplementation may have a beneficial effect, but caution should be exercised in generalising this finding.

The prophylaxis trials found a reduction in common cold duration of 8% in adults and 13% in children. The practical relevance of these findings is open. In our opinion, the evidence does not support the use of prophylaxis in its own right. So far, therapeutic supplementation has been shown to be effective in marathon runners. Nevertheless, given the consistent effect of vitamin C on the incidence of colds in children, it would be reasonable to consider therapeutic supplementation on an individual basis whether therapeutic vitamin C supplementation is justified.

AUTHORS' CONCLUSIONS

- **Implications for practice**
- **Policy implications**
- **Implications for research**

Implications for research

It does not seem worthwhile to carry out further regular supplementation trials in the general population. However, the findings in marathon runners, skiers and soldiers operating in subarctic conditions warrant further research.

None of the therapeutic trials carried out so far have examined the effect of vitamin C on children, even though the regular supplementation trials have found substantially greater effect on cold duration in children than in adults. In view of the greater incidence of respiratory infections in children, such therapeutic trials are warranted.

The findings in the Anderson 1974 study on the greater benefit of 8 g than 4 g dose on the day of onset of respiratory symptoms suggest that doses in further therapeutic trials with adults should be at least 8 g/day.

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Sesgos de publicación/selección/citación

- Resultados "positivos"
- Resultados significativos
- Muestras grandes
- Grupos de investigación de prestigio

- Rápida publicación
- Publicación múltiple
- Mayor citación
- Aumenta la probabilidad de que aparezcan en la bibliografía

BMJ 2005;331:433-434

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Why science needs to publish negative results
<https://www.elsevier.com/authors-update/story/innovation-in-publishing/why-science-needs-to-publish-negative-results>

Journal
of Negative & No Positive Results

<http://revistas.proeditio.com/jonpr/index>



Figure 1: The most common approach taken by journals, in which only those experiments yielding positive results end up as publication material.

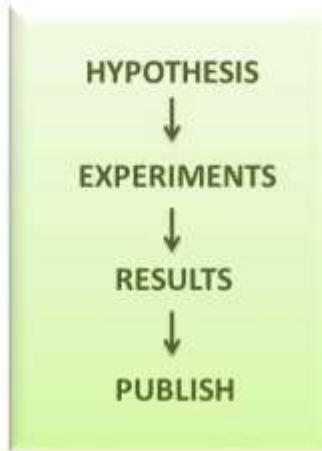


Figure 2: A more neutral approach, in which all results are published, as long as they are generated by well-carried out experiments based on sound hypotheses.

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Sesgo del idioma

- Revistas de lengua inglesa son las de mayor prestigio y difusión. Los trabajos escritos en inglés aparecen con mayor frecuencia en las bases de datos.

Sesgos de publicación

- Financiación por parte de la industria: puede condicionar que sólo se publiquen los resultados favorables.
- Sesgo del propio autor en la selección de los trabajos (subjetividad) (es importante que en el meta-análisis queden bien definidos los criterios de selección).

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*Descubrimiento de los nutrientes
"The golden age of Nutrition"
(Carpenter, 2003)*

El “milagro” de las vitaminas

“descubrimiento tan importante como el de las vacunas o el saneamiento de las aguas” (Trichopoulou y col., 2000)

1881. N Lunin (1844-1920). Basilea (Suiza)

“A la dieta sintética (mezcla artificial de componentes purificados de leche) + agua le falta alguna sustancia desconocida sin la cual no puede llevarse a cabo la vida”.

Dieta sintética

1897. Christian Eijkman (1858-1930).

Holandés que trabajaba en la isla de Java (Indonesia).

Provoca el Beri-beri en pollos alimentándolos con una dieta que sólo contenía arroz pulido.

Las aves curaban cuando se les daba arroz integral o la cascarilla del arroz.

Enfermedad carencial

Sesgo del idioma

1905. Cornelius Adrianus Pekelharing (1848-1922). Utrecht (Holanda).

Dieta sintética + agua + **leche**.

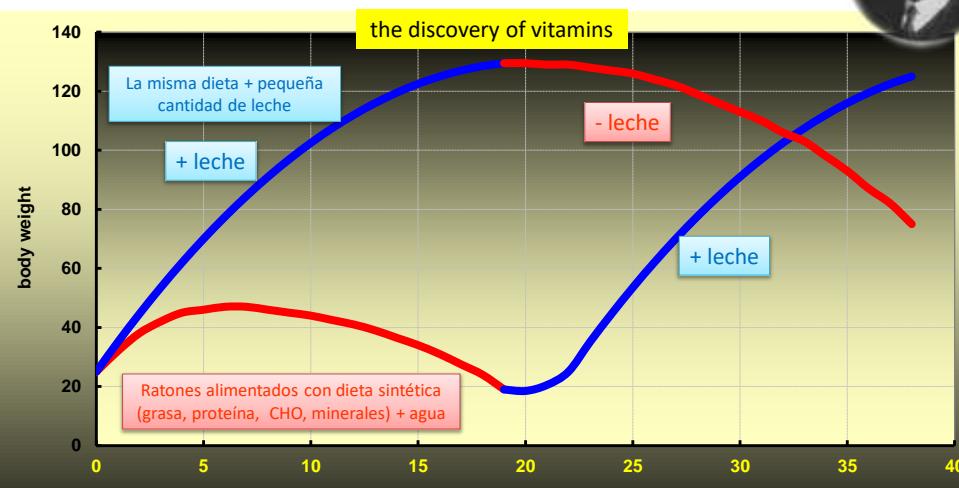
“En la leche hay una sustancia que, incluso en pequeñas cantidades, es necesaria para la vida y sin la cual el animal pierde la capacidad de usar los otros componentes de la dieta”. **Trabajo escrito en holandés.**



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1906-1912. Frederick Gowland Hopkins (1861-1947). Bioquímico inglés.

Establece la existencia de las vitaminas.



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1906-1912. Frederick Gowland Hopkins (1861-1947). Bioquímico inglés.
Establece la existencia de las vitaminas.



“En los alimentos naturales, como en este caso la leche, existen sustancias que en pequeña cantidad son necesarias para la nutrición de los animales:

“factores accesorios de la alimentación”.

(“Feeding experiments illustrating the importance of accessory factors in normal diets”. J Physiol 1912;44:425-460).

Concepto de “enfermedad carencial”.

Alimentos: portadores de “reguladores metabólicos” = **vitaminas**.

Premio Nobel de Medicina (1929) junto con el holandés Eijkman.

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The Nobel Prize in Physiology or Medicine 1929
Christiaan Eijkman, Sir Frederick Hopkins



The Nobel Prize in Physiology or Medicine 1929

Nobel Prize Award Ceremony

Christiaan Eijkman

Sir Frederick Hopkins



Biographical

Nobel Lecture

Banquet Speech

Documentary

Nobel Lecture

Nobel Lecture, December 11, 1929

http://nobelprize.org/nobel_prizes/medicine/laureates/1929/hopkins-lecture.html

The Earlier History of Vitamin Research

When the present century began, animal nutrition was being viewed too exclusively from the standpoint of energy requirements. The fundamental pioneer work of Rubner and its later extension to human subjects in the remarkable enterprise of Atwater, Benedict, Rosa, and others in the United States could not fail to produce a deep impression upon the thought of the time. The quantitative character of the data obtained and the attractive circumstance that such data appeared to supply a real measure of nutritional needs, independent of, and apparently superior to, considerations based upon chemical details, induced a feeling that knowledge concerning these needs had become highly adequate and was approximating even to finality. As a matter of fact, however, these calorimetric studies, invaluable in themselves, were then

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**Sir Frederick Hopkins. The Nobel Prize in Physiology or Medicine 1929
The Earlier History of Vitamin Research. Nobel Lecture, December 11, 1929**



Yet the pregnant suggestions arising from the observations just discussed did ultimately, though not for many years after the latest of them were published, awaken (as we may suppose) the interest of an investigator distinguished in many fields who was led to repeat and extend them. **I allude to the late Professor Pekelharing, whose own observations (published in 1905) unhappily again remained unknown to the majority till very recently. It is indeed astonishing that the results of such significant work as his, though published in the Dutch language alone, should not have become rapidly broadcast. I cannot refrain from referring to the circumstance that the paper was not abstracted or mentioned in Maly's *Jahresbericht für Thierchemie*, so adequate, and in general so complete, in its dealings with current literature.** Many of us were accustomed to rely upon it for references to work published in journals that we could not consult, or in a language that we could not read. Though other work by Pekelharing was duly recorded at this time, no mention was made of the extraordinarily interesting paper in question. My own experiments began soon after the paper was published, and as a proportion of my own work was very similar to that of Pekelharing, I shall never cease to regret that, in common with so many others, I was then completely ignorant of the latter. After speaking of experiments carried out on lines similar to some of those done in Bunge's laboratory, and indicating that they pointed to the existence of some unknown essential, Pekelharing goes on to say: "Till now my efforts, constantly repeated during the last few years, to separate this substance and get to know more about it, have not led to a satisfactory result, so I shall not say any more about them. My intention is only to point out that there is a still unknown substance in milk which even in very small quantities is of paramount importance to nutrition. If this substance is absent, the organism loses the power properly to assimilate the well-known principal parts of food, the appetite is lost and with apparent abundance the animals die of want. Undoubtedly this substance occurs not only in milk but in all sorts of foodstuffs both of vegetable and animal origin." Here we have a clear statement of the vitamin doctrine already a quarter of a century old. It is noteworthy that Pekelharing records prolonged endeavours towards the isolation of a vitamin.

http://nobelprize.org/nobel_prizes/medicine/laureates/1929/hopkins-lecture.html

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Conflictos de intereses

"No investigators have any affiliations or financial involvement (e.g., employment, consultancies, honoraria, stock options, expert testimony, grants or patents received or pending, or royalties) that conflict with material presented in this report."

Los autores declaran que no existe ningún compromiso o vínculo con la entidad financiadora que pueda ser entendido como un conflicto de intereses.

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Estandarización, normas y control de revisiones sistemáticas y meta-análisis. Control de calidad

- MOOSE (Meta-analysis of Observational Studies in Epidemiology)
JAMA 2000;283:2008–12.
- QUOROM (Quality of Reporting of Meta-Analyses)
Lancet 1999;354:1896–900.
- Cochrane Handbook, <http://www.cochrane-handbook.org>
- AHRQ Methods Guide for Effectiveness and Comparative Effectiveness Reviews
- Institutes of Medicine Standards for Systematic Reviews
- The PRISMA Statement (Preferred Reporting Items for Systematic Reviews and Meta-Analyses)
<http://www.prisma-statement.org/>

Reporting of systematic reviews of micronutrients and health: a critical appraisal1–4
Mei Chung, Ethan M Balk, Stanley Ip, Gowri Raman, Winifred W Yu, Thomas A Trikalinos, Alice H Lichtenstein, Elizabeth A Yetley, and Joseph Lau
Am J Clin Nutr 2009;89:1–15.

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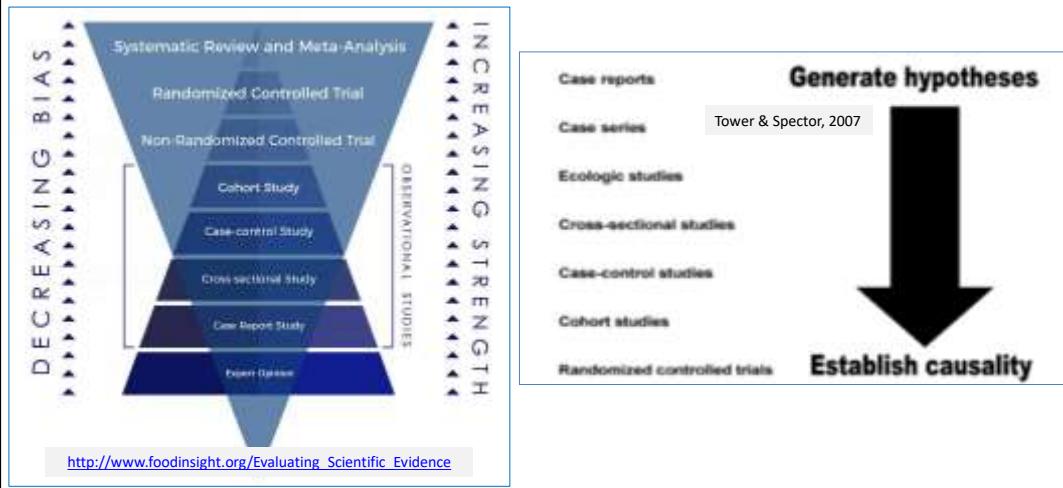
TODA LA INFORMACIÓN CIENTÍFICA ¿“PESA” LO MISMO?

NO



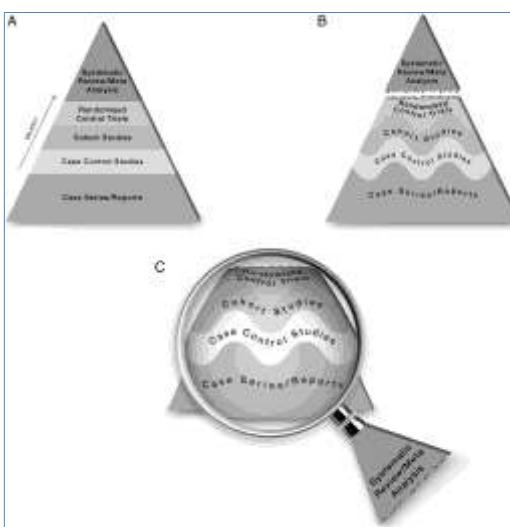
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Jerarquía de la Evidencia Científica



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The proposed new evidence-based medicine pyramid.



M Hassan Murad et al. Evid Based Med 2016;21:125-127

<https://ebm.bmjjournals.org/content/21/4/125>

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Niveles de Evidencia Científica y Grados de recomendación

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Importancia de tomar decisiones

Convinciente: Evidencia de una relación causal concluyente. **Información suficiente para realizar recomendaciones dietéticas (RD) en la población en general.** Poca o ninguna evidencia de lo contrario. La asociación debe ser biológicamente plausible.

Probable: Evidencia lo suficientemente fuerte ($RR/OR > 2$ o < 0.5 , estadísticamente significativo) para concluir que puede haber una relación causal. **En función de esta información también pueden realizarse RD.**

Possible: Puede existir una relación causal, pero la evidencia **no es lo suficientemente fuerte como para establecer RD.**

Insuficiente: Evidencia sugerente pero tan escasa o contradictoria que **no permite llegar a ninguna conclusión.**

WCRF, 2007; <http://eprints.ucl.ac.uk/4841/1/4841.pdf>

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Adapted from the World Health Organization, Diet, Nutrition and the Prevention of Chronic Diseases: report of a joint WHO/FAO expert consultation (WHO, FAO, 2003).

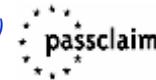
Grading	Evidence
Convinciente	Evidencia de estudios epidemiológicos que muestren <u>asociaciones consistentes</u> entre exposición y enfermedad, con poca o ninguna evidencia de lo contrario. Se basa en resultados de un <u>número importante</u> de estudios: <ul style="list-style-type: none">•Observacionales prospectivos y•Ensayaos controlados aleatorizados, realizados con muestras grandes y adecuada duración. La asociación debe ser biológicamente plausible
Probable	Evidencia basada en estudios epidemiológicos que muestren asociaciones <u>bastante consistentes</u> entre exposición y enfermedad. Limitaciones por: evidencia de lo contrario, insuficiente duración del estudio, insuficientes estudios, muestras inadecuadas, seguimiento incompleto, etc. La asociación debe ser biológicamente plausible
Possible	Evidencia basada principalmente en resultados de: <ul style="list-style-type: none">•Estudios caso-control y•estudios transversales. Puede haber también resultados de algunos: <ul style="list-style-type: none">•Ensayaos controlados aleatorizados•Ensayaos controlados no aleatorizados•Estudios observacionales Son necesarios más ensayos que apoyen la asociación. La asociación debe ser biológicamente plausible
Insuficiente	Pocos estudios que sugieran la asociación, insuficientes para establecer dicha asociación. Limitada información o ninguna procedente de ensayos controlados aleatorizados. Son necesarios más estudios mejor diseñados

(WCRF, 1997; WHO, 2003. World Cancer Research Fund. American Institute for Cancer Research. Food, Nutrition and Prevention of Cancer: a Global Perspective. 1997. <http://www.wcrf.org/>. WHO/FAO. 2003. Diet, Nutrition and the Prevention of Chronic Diseases. Technical Report Series 916. <http://www.who.int/dietphysicalactivity/publications/trs916/en/>)

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Etiquetado nutricional de los alimentos

PASSCLAIM ("Process of the Assessment of Scientific Support for Claims on Foods")



Framework for strength of evidence

- **CONVINCING:** A Significant scientific agreement can be trusted to guide practice
- **PROBABLE:** B Good to moderate level of scientific agreement can be trusted to guide practice in most situations
- **POSSIBLE:** C Low level of scientific agreement some support for the recommendations but care should be taken in its application
- **INSUFFICIENT:** D Very low level of scientific agreement evidence is weak and any recommendation must be applied with caution

WCRF (2007); USFDA (2003); WHO (2004); CODEX (2007)

Professor D. P. Richardson, DPR Nutrition Ltd, UK

<http://www.eski.hu/new3/konyvtar/bookshop/EFSA%20Conference%20on%20Nutrition%20and%20Health%20Claims.pdf>

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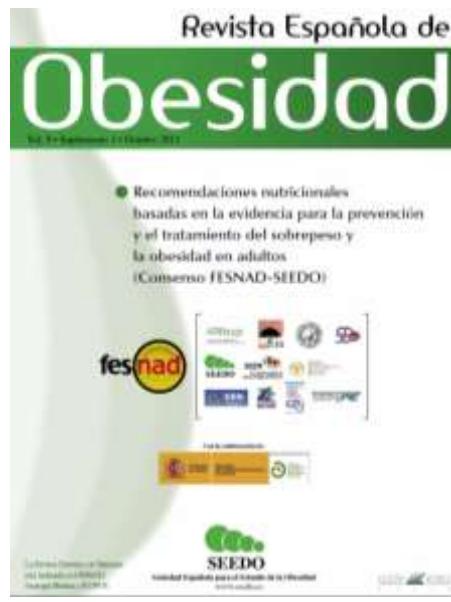
Evidence Grading Summary

Types of Research: Evidence Hierarchies	
Agency for Healthcare Research and Quality (AHRQ)	
Level I	Meta-Analysis (Combination of data from many studies)
Level II	Experimental Designs (Randomized Control Trials)
Level III	Well designed Quasi Experimental Designs (Not randomized or no control group)
Level IV	Well designed Non-Experimental Designs (Descriptive-can include qualitative)
Level V	Case reports/clinical expertise

Strength of Evidence	
United States Preventive Services Task Force (USPSTF) Grading	
A	Strongly recommended; Good evidence
B	Recommended; At least fair evidence
C	No recommendation; Balance of benefits and harms too close to justify a recommendation
D	Recommend against; Fair evidence is ineffective or harm outweighs the benefit
I	Insufficient evidence; Evidence is lacking or of poor quality, benefit and harms cannot be determined

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Ejemplo



http://www.naos.aesan.msp.es/naos/ficheros/investigacion/Consenso_SEDO.pdf

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LA DIETA EN LA PREVENCIÓN DE LA OBESIDAD

1. Equilibrio energético y peso corporal

1.1. Densidad energética

Evidencia

1. Los patrones alimentarios de alta densidad energética pueden conducir a un incremento de peso en adultos (nivel de evidencia 1+).

Recomendaciones

1. El aumento de peso puede prevenirse mediante dietas que contengan alimentos con baja densidad energética (recomendación de grado A).

1.2. Equilibrio energético y ambiente obesogénico

Evidencia

2. La ausencia de supermercados con disponibilidad de frutas y hortalizas o su ubicación a grandes distancias, sobre todo en núcleos humanos con niveles socioeconómicos desfavorecidos, son factores condicionantes de un mayor IMC medio poblacional (nivel de evidencia 1+).

Recomendaciones

2. Deben arbitrarse estrategias que hagan posible la disponibilidad alimentaria y el acceso a alimentos saludables, en especial a frutas y hortalizas, para crear ambientes favorables

1.4. Equilibrio energético: tamaño de las raciones

Evidencia

4. El ofrecimiento de raciones de mayor tamaño condiciona un aumento en la ingesta energética de los individuos (nivel de evidencia 2++).

Recomendaciones

4. La utilización de raciones de menor tamaño limita la ingesta energética (recomendación de grado B).

1.5. Equilibrio energético: desayuno

Evidencia

5. Son controvertidas e inconsistentes las investigaciones que estudian la relación entre la omisión del desayuno en adultos y el riesgo de sobrepeso y obesidad.

1.6. Equilibrio energético: aperitivos

Evidencia

6. Son controvertidas e inconsistentes las investigaciones que sugieren que el consumo de aperitivos está asociado con el incremento de peso.

1.7. Equilibrio energético: frecuencia

Evidencia

7. Las investigaciones que estudian la relación entre la

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MÉTODO SIGN: Niveles de evidencia

(Scottish Intercollegiate Guidelines Network)

Tabla 1. NIVELES DE EVIDENCIA^{III}

1	1++	Metaanálisis de alta calidad, revisiones sistemáticas de ECA, o ECA con un riesgo muy bajo de sesgo
	1+	Metaanálisis bien realizados, revisiones sistemáticas de ECA, o ECA con bajo riesgo de sesgo
	1-	Metaanálisis, revisiones sistemáticas de ECA o ECA con alto riesgo de sesgo
2	2++	Revisiones sistemáticas de alta calidad de estudios caso-control o de cohortes
	2+	Estudios caso-control o de cohortes de alta calidad con un riesgo muy bajo de confusión o sesgo, y una alta probabilidad de que la relación sea causal
	2-	Estudios caso-control o de cohortes bien realizados: con un riesgo bajo de confusión o sesgo, y una probabilidad moderada de que la relación sea causal
3		Estudios no analíticos (p. ej.: casos clínicos o series de casos)
4		Opinión de experto/s

ECA: ensayo controlado aleatorizado

Tabla 2. GRADOS DE RECOMENDACIÓN^{III}

A	Como mínimo un metaanálisis, revisión sistemática o ECA con una clasificación de 1++ y directamente aplicable a la población diana; o una revisión sistemática o ECA con un cuerpo de evidencia consistente principalmente en estudios puntuados como 1+, directamente aplicable a la población diana, y que demuestre una consistencia global en sus resultados
	Un cuerpo de evidencias que incluya estudios puntuados como 2++, directamente aplicables a la población diana y que demuestre una consistencia global en sus resultados; o evidencias extrapoladas de estudios puntuados como 1++ o 1+
B	Un cuerpo de evidencias que incluya estudios puntuados como 2+, directamente aplicables a la población diana y que demuestre una consistencia global en sus resultados; o evidencias extrapoladas de estudios puntuados como 2++
	Un cuerpo de evidencias que incluya estudios puntuados como 2-, directamente aplicables a la población diana y que demuestre una consistencia global en sus resultados; o evidencias extrapoladas de estudios puntuados como 2+
C	Evidencias de nivel 3 o 4; o evidencias extrapoladas de estudios puntuados como 2+
	ECA: ensayo controlado aleatorizado Los estudios clasificados como 1- y 2- no deben usarse en el proceso de elaboración de recomendaciones, por su alto potencial de sesgo

ECA: ensayo controlado aleatorizado

Los estudios clasificados como 1- y 2- no deben usarse en el proceso de elaboración de recomendaciones, por su alto potencial de sesgo

http://www.naos.aesan.msp.es/naos/ficheros/investigacion/Consenso_SEEDO.pdf

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MÉTODO SIGN: Niveles de evidencia

(Scottish Intercollegiate Guidelines Network)

- SIGN evalúa la calidad de la evidencia científica en función de:

- Diseño del estudio:

- Ensayos clínicos , revisiones sistemáticas o metanálisis (1)
- Opinión de expertos (4)

- Riesgo de sesgo:

++	Se han cumplido todos o la mayoría de los criterios de calidad metodológica. En los puntos en que no se han cumplido, se considera muy poco probable que dicho incumplimiento pueda afectar a las conclusiones del estudio o revisión.
+	Se han cumplido algunos de los criterios de calidad metodológica. Se considera poco probable que los criterios que no se han cumplido o que no se describen de forma adecuada puedan afectar a las conclusiones.
-	Se han cumplido sólo unos pocos criterios de calidad metodológica, o ninguno de ellos. Se considera probable o muy probable que esto afecte a las conclusiones.

Scottish Intercollegiate Guidelines Network. A guideline developers' handbook (Publication no 50). Edinburgh: SIGN: 2001 [actualizado Noviembre 2011]. Disponible en: <http://www.sign.ac.uk/pdf/sign50.pdf>

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<https://www.ucm.es/innovadieta/nube>
<https://www.ucm.es/innovadieta/documentos-consenso>

En las BBDD de Innovadieta busca la evidencia sobre los efectos del chocolate negro en la salud

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BBDD de Evidencia científica Algunos ejemplos

The screenshot shows the homepage of the Innovadieta database. At the top, there's a navigation bar with links to 'Estadística web', 'Documentos', 'Bibliografía', 'Valores Nutricionales', 'Nutrición en massa-masa', and 'ABC de la Nutrición'. Below the navigation bar, there's a search bar with the placeholder 'Ángela... Busca en el inventario general... Nutrición y Dietética basadas en la evidencia (NuBe)'. The main content area has a heading 'Nutrición y Dietética basadas en la evidencia (NuBe)'. Underneath this, there's a section titled 'Sistemas de evidencia basados en la evidencia' with a link to 'Nutrición y Dietética basadas en la evidencia (NuBe)'. A large image of a healthy meal is displayed, along with a snippet of text: 'NICE Standard for Health and Care Excellence (NICE Guidance)' and 'Office of Dietary Supplements, National Institutes of Health, United States National Institutes of Health'. At the bottom, there's a footer with links to 'The All-Complaint Evidence Analysis Library (AllEvidence), of the Office of Health and Disease', 'The Evidence Library', 'The Evidence Library', 'The Evidence Library', and 'The Evidence Library'.

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The screenshot shows the homepage of the RED-Nube website. At the top, there is a navigation bar with links: INICIO, QUÉNES SOMOS, COLABORA, PROYECTOS, RE-CURSOS, and BLOG. Below the navigation bar, there is a main content area featuring the RED-Nube logo and a brief description of the network's mission. A sidebar on the left contains the PEN logo and some text. A callout box in the center provides information about the FEDN and CAEC-FEDN.

<http://www.rednube.net/>

RED de Nutrición
BÁSADA en la EVIDENCIA

La Red de Nutrición Basada en la Evidencia (RED-Nube) es una red de trabajo colaborativo que pretende unir todos los esfuerzos que se están realizando en investigación secundaria (elaboración de guías basadas en pruebas o evidencias y metasíntesis) en el campo de la Nutrición Humana y Dietética, fomentando tanto el acceso libre y equitativo como el acceso más económico y asequible a las principales herramientas y bases de datos especializadas existentes.

Entre los objetivos de RED-Nube se encuentra acercar las mejores pruebas importantes (evidencias) a aquellos profesionales que más las necesitan, los dietistas-nutricionistas que trabajan día a día con los pacientes/clientes. Para ello hemos desarrollado un sistema de evidencias rápidas que permite el acceso a las mejores recomendaciones basadas en pruebas (evidencias) en pocos minutos. La base al inicio de dicho sistema se realiza a través del software de ayuda de tama de decisiones Practice-based Evidence in Nutrition (PEN®) un sistema creado por y para dietistas-nutricionistas de todo el mundo. RED-Nube, a través de la Academia Española de Nutrición y Dietética y del Consejo General de Dietistas-Nutricionistas de España se ha convertido en Partner Iberoamericano con permiso oficial para ofrecer licencias individuales al PEN® a un precio doble y adaptado a la situación económica de este profesional. Más información sobre licencias del PEN® aquí.

Nutrición basada en la evidencia, FEDN | Centro de Análisis de la Evidencia Científica de la Fundación Española de Dietistas-Nutricionistas (CAEC-FEDN)
<http://www.academianutricionydietetica.org/>

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The screenshot shows the 'Publicaciones RED-Nube' section of the website. It features a header with the RED-Nube logo and a sub-header 'Entradas en el blog de RED-Nube'. Below this, there is a link to the blog page. Further down, there is a section for 'Digestor semanal de movimiento en redes sociales' with a link to a paper. The section is divided into years: 2017, 2016, and 2015. Each year has a list of publications with their titles, authors, and links. At the bottom of the section, there is a link to the publications page.

Publicaciones RED-Nube

Entradas en el blog de RED-Nube:
<http://www.rednube.net/blog/>

Digestor semanal de movimiento en redes sociales:
<http://paperkit.e-1400136177/>

Revistas Científicas y otras monografías

2017

- Baladía E, Martínez-Rodríguez R. Revisión científica sobre la alimentación y nutrición en el embarazo. RED-Nube; 2017. Disponible en: http://diamandibularstudinutricional.org/wp-content/uploads/2017/05/Baladia_evidencias_low.pdf / en Researchgate

2016

- Baladía E, Martínez-Rodríguez R. Legumbres y salud: resumen de evidencias rápidas. RED-Nube; 2016. Disponible en: http://www.rednube.net/docs/legumbres_2016.pdf / Presentación del documento en YouTube / en Researchgate
- Baladía E, Martínez-Rodríguez R. Conflictos de interés en nutrición humana y dietética / Conflicts of interest in human nutrition and dietetics. Rev Esp Nutr Hum Diet. 2016; 20(2):77-79. doi: 10.14306/rehyd.20.3.261
- Baladía E, Frutos Pérez-Surio A, Martínez-Rodríguez R. Resumen de recomendaciones nutricionales basadas en la evidencia de la Guía de Práctica Clínica para el manejo de pacientes con enfermedad de Parkinson /Summary of evidence-based nutritional recommendations of the Clinical Practice Guideline for the management of patients with Parkinson's disease. Nutr Hosp. 2016; 33(3):749-760 [factor de impacto = 1,487]. Texto completo. También en Researchgate.
- Estarí M, Aguilar Barreira ES, Martínez Rodríguez R, Baladía E, Duran Agüero S, Camacho S, Bühring K, Herrero López A, Gil González DM. Fases de referencia para publicar Protocolos de Revisões Sistemáticas y Metaanálisis: Declaración PRISMA-P 2015. Rev Esp Nutr Hum Diet. 2016; 20(2):148 - 160. doi: 10.14306/rehyd.20.3.2323

<http://www.rednube.net/publicaciones-red-nube/>

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The screenshot shows the homepage of the PEN website. At the top, there is a navigation bar with links for 'Help', 'Contact Us', 'Log In', 'Connect', 'Username', 'Password', 'Forgot It?', 'Logout', 'Home', 'About PEN', 'Subscribe', and 'eNews Issues'. A yellow box highlights the text 'Dietitians of Canada 2012' and the URL 'http://www.pennutrition.com/index.aspx'. A large green banner at the top says 'Welcome to PEN' and 'Practice-based Evidence in Nutrition [PEN]'. Below the banner, it states: 'PEN is a dynamic knowledge translation subscription service available internationally as individual or group licenses for food, nutrition, and dietetic practice.' There are several call-to-action buttons: 'Become a PEN Author or Reviewer', 'Take our PEN Survey', 'Curious About PEN?', 'Start your free 15-day trial today.', 'PEN eNews is Free! Sign up Today.', and 'Sign Up'. Below these are four main sections with icons: 'What's Being Said About PEN' (speech bubble icon), 'PEN Orientation Tutorial' (play button icon), 'What's New in PEN Global' (globe icon), and 'PEN Terminology' (two 'A' icons). Each section has a brief description and a 'Watch' or 'Explore' button.

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The screenshot shows the homepage of the Evidence Analysis Library (EAL). At the top, there is a navigation bar with links for 'SEARCH', 'Sign In', 'Register', 'Contact Us', and 'Help'. A yellow box highlights the text 'Necesita registro'. Below the navigation bar, there is a green banner with the text 'EVIDENCE ANALYSIS LIBRARY' and a 'Welcome' message. The main content area features a grid layout. On the left, there is a sidebar with a 'EVIDENCE BASED PRACTICE' section containing a green icon of a person, two bullet points, and a copyright notice: 'Copyright 2013 © Academy of Nutrition and Dietetics UK'. The main content area includes sections for 'Topics' (with categories like Diseases & Conditions Topics, Foods Topics, Nutrients Topics, and Nutrition Care Process Topics), 'A-Z Index' (listing categories from A to Z), and 'Topics in Nutrition' (listing categories like About the EAL, ADA Evidence-Based Positions, Adolescent Nutrition, Adult Nutrition, Adult Weight Management (AWM) index, Advance Food Production and Sustainable Agriculture Project, Aging, and Other Adults, Antioxidants, Aspartame index, and Assessment).

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THE ADA NUTRITION EVIDENCE ANALYSIS LIBRARY

<http://www.adaevidencelibrary.com/default.cfm?auth=1>

Grade I: Good—The evidence consists of results from studies of **strong** design for answering the question addressed. The results are both clinically important and consistent with minor exceptions at most. The results are free of serious doubts about generalizability, bias, and flaws in research design. Studies with negative results have sufficiently large sample sizes to have adequate statistical power.

Grade II: Fair—The evidence consists of results from studies of strong design answering the question addressed, but there is uncertainty attached to the conclusion because of inconsistencies among the results from different studies or because of doubts about generalizability, bias, research design flaws, or adequacy of sample size. Alternatively, the evidence consists solely of results from weaker designs for the questions addressed, but the results have been confirmed in separate studies and are consistent with minor exceptions at most.

Grade III: Limited—The evidence consists of results from a limited number of studies of **weak** design for answering the questions addressed. Evidence from studies of strong design is either unavailable because no studies of strong design have been done or because the studies that have been done are inconclusive due to lack of generalizability, bias, design flaws, or inadequate sample sizes.

Grade IV: Expert Opinion Only—The support of the conclusion consists solely of the statement of informed medical commentators based on their clinical experience, unsubstantiated by the results of any research studies.

Grade V: Not Assignable—There is no evidence available that directly supports or refutes the conclusion.

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Conditional versus Imperative Recommendations

Recommendations can be worded as conditional or imperative statements. **Conditional** statements clearly define a specific situation, while **imperative** statements are broadly applicable to the target population without restraints on their pertinence.

More specifically, a **conditional recommendation** can be stated in **if/then** terminology (e.g., If an individual does not eat food sources of omega-3 fatty acids, then 1g of EPA and DHA omega-3 fatty acid supplements may be recommended for secondary prevention).

In contrast, **imperative recommendations** “require,” or “must,” or “should achieve certain goals,” but do not contain conditional text that would limit their applicability to specified circumstances. (e.g., Portion control should be included as part of a comprehensive weight management program. Portion control at meals and snacks results in reduced energy intake and weight loss).

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¿Cuántas comidas al día deben hacerse?

Position of the American Dietetic Association: Weight Management
Journal of the American Dietetic Association - Volume 109/2, Pages 185-364 (February 2009)

Revisión sistemática de todos los estudios y concluyen:

EAL Recommendation "Total caloric intake should be distributed throughout the day, with the consumption of four to five meals/snacks per day including breakfast. Consumption of greater energy intake during the day may be preferable to evening consumption"

Rating: Fair, Imperative.

Conclusion. Several studies show that consumption of four to five meals or snacks per day is associated with reduced or no obesity risk, while three or fewer and six or more meals or snacks per day may result in increased risk of obesity, depending on gender. Higher eating frequency is related to lower total daily energy intake and body weights in men, but in women the data is less conclusive. Five studies demonstrate that consumption of greater energy intake in the morning vs. the evening is associated with lower body weights and greater loss of weight. Further research is needed on the distribution of calories consumed at meals and snacks during the day.

Grade II

Recommendation Strength Rationale

- Conclusion statements both given a Grade II
- Consistent findings among a variety of study designs
- Minority Opinions

Consensus reached

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Nutrition Evidence Library (NEL):
Answering food and nutrition questions
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A state-of-the-art method for evaluating scientific evidence to answer a precise question or series of questions developed by a

WELCOME TO NEL

USDA's Nutrition Evidence Library (NEL) specializes in conducting systematic reviews to inform Federal nutrition policy and programs. NEL staff collaborate with stakeholders and leading scientists using state-of-the-art methodology to

WHAT'S NEW

2015 Dietary Guidelines Advisory Committee's Systematic Reviews

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TOPICS

- Digital Media and Technology
- Parental Involvement
- Type of Educator
- Food Environment
- Single/Multi Component
- Methodology
- Acknowledgements

A Series of Systematic Reviews on the Effects of Nutrition Education on Children's and Adolescents' Dietary Intake

Full Report
Appendices A-C
Appendices D-J

Systematic Review Questions

Executive Summary

Consuming a healthy diet consistent with the Dietary Guidelines for Americans, 2010[[1]] can help individuals achieve and maintain a healthy weight, reduce the risk of developing chronic diseases, and promote good health. However, many children are consuming excess calories, while not meeting nutrient needs, and are overweight/obese and/or at increased risk for future health problems.

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Cont.

Systematic Review Questions

The systematic review questions addressed were identified and prioritized to focus the reviews on topics that would enhance current nutrition education practice and programs. A broad range of school- and community-based nutrition education interventions involving children in preschool, kindergarten, and elementary school and adolescents in middle and high school were considered, in order to answer the following systematic review questions:

1. What is the effect of nutrition education delivered via digital media and technology on children's and adolescents' dietary intake related behaviors?
2. What is the effect of nutrition education with parental involvement compared to no parental involvement on children's and adolescents' dietary intake related behaviors?
3. Which type of educator who delivers nutrition education is most effective in changing children's and adolescents' dietary intake related behaviors?
4. What are the effects of combining changes in the food environment and nutrition education compared to active or other strategies alone in children's and adolescents' dietary intake related behaviors?
5. What are the effects of multi-component compared to single component nutrition education interventions on children's and adolescents' dietary intake related behaviors?

Background and Methodology

USDA's Nutrition Evidence Library (NEL) conducted these systematic reviews. The NEL uses a rigorous, transparent, and reproducible methodology to conduct systematic reviews on food- and nutrition-related topics to support Federal nutrition policies and programs using a six-step process:

1. Develop systematic review questions
2. Create and implement literature search and sort plans
3. Develop evidence portfolios
4. Synthesize the bodies of evidence
5. Develop conclusion statements and grade the evidence
6. Describe research recommendations

This NEL systematic review project was planned, organized, and guided by a NEL Systematic Review Management Team composed of Federal nutritionists trained in systematic review methodology. The NEL Systematic review team worked with a Technical Expert Collaborative (TEC), which consisted of eight leading nutrition education experts, whose expertise was needed to address specific issues related to the scope of nutrition education and to a synthesis of the body of evidence.

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e-Library of Evidence for Nutrition Actions (eLENA)

Iodine supplementation during pregnancy

Pregnant women need about 60% more iodine than non-pregnant women. This nutrient is essential for healthy brain development in the fetus and young child.

Most foods are relatively low in iodine content. To ensure that everyone has a sufficient intake of iodine, WHO and UNICEF recommend universal salt iodization as a global strategy. However, in certain countries salt iodization may not be feasible in all regions.

WHO and UNICEF therefore recommend iodine supplementation for pregnant and lactating women in countries where less than 20% of households have access to iodized salt, until the salt iodization programme is scaled up. Countries with a household access to iodized salt between 20 and 90% should make efforts to accelerate salt iodization or assess the feasibility of increasing iodine intake in the form of a supplement or iodine-fortified foods by the most susceptible groups.

<http://www.who.int/elenas/en/index.html>
En español: <http://www.who.int/elenas/es/index.html>

Iodine supplementation during pregnancy
Status: guidelines under development
Publication year: expected 2012

Last update: 21 October 2011 11:31 CEST

Related links

- Publication: Nutrition essentials: a guide for health managers [pdf, 1.9 Mb]
- Publication: Iodine deficiency in 2007: Global progress since 2003 [pdf, 421kb]
- Publication: Prevention and control of iodine deficiency in pregnant and lactating women and in children less than 2 years old: conclusions and recommendations of the technical consultation [pdf, 87kb]
- Publication: Iodized salt during pregnancy: safe use of iodized salt to prevent iodine deficiency in pregnant women [pdf, 475kb]

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e-Library of Evidence for Nutrition Actions (eLENA)

Increasing fruit and vegetable consumption to reduce the risk of noncommunicable diseases

Fruits and vegetables are important components of a healthy diet. Reduced fruit and vegetable consumption is linked to poor health and increased risk of noncommunicable diseases (NCDs). An estimated 6.7 million deaths worldwide were attributable to inadequate fruit and vegetable consumption in 2010.

Including fruits and vegetables as part of the daily diet may reduce the risk of some NCDs including cardiovascular diseases and certain types of cancer. More limited evidence suggests that when consumed as part of a healthy diet, low in fat, sugars and salt/sodium, fruits and vegetables may also help to prevent weight gain and reduce the risk of obesity, an independent risk-factor for NCDs.

Moreover, fruits and vegetables are rich sources of vitamins and minerals, dietary fibre and a host of beneficial non-nutrient substances including plant sterols, flavonoids and other antioxidants and consuming a variety of fruits and vegetables helps to ensure an adequate intake of many of these essential nutrients.

WHO recommendations
As part of a healthy diet low in fat, sugars and sodium, WHO suggests consuming more than 400 grams of fruits and vegetables per day to improve overall health and reduce the risk of certain NCDs.

Category 2 intervention
Systematic review(s) have been conducted but no recent guidelines yet available that have been approved by the WHO Guidelines Review Committee.
More about categories of interventions

Biological, behavioural and contextual rationale
Increasing fruit and vegetable consumption to reduce the risk of noncommunicable diseases

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Evaluar la calidad de la Evidencia

**Grading of Recommendations Assessment, Development,
and Evaluation (año 2000)**



www.gradeworkinggroup.org

Ver también: <http://www.fisterra.com/guias-clinicas/la-evaluacion-calidad-evidencia-graduacion-fuerza-recomendaciones-sistema-grade/>
<http://www.sciencedirect.com/science/article/pii/S0212656714000493>

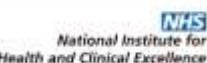
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MÉTODO GRADE



GRADE working group <http://www.gradeworkinggroup.org>

Organizations that have endorsed or that are using GRADE*



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Método GRADE: Representación



Calidad de la evidencia científica		
Alta	⊕⊕⊕⊕	A
Moderada	⊕⊕⊕○	B
Baja	⊕⊕○○	C
Muy baja	⊕○○○	D
Fuerza de la recomendación		
Recomendación fuerte a favor de utilizar una intervención	↑↑	1
Recomendación débil a favor de utilizar una intervención	↑?	2
Recomendación débil en contra de utilizar una intervención	↓?	2
Recomendación fuerte en contra de utilizar una intervención	↓↓	1

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Método GRADE: Niveles de evidencia



Calidad de la evidencia científica	Definición
Alta (⊕⊕⊕⊕)	Estamos muy seguros de que el verdadero efecto se encuentra cerca de la estimación del efecto.
Moderada (⊕⊕⊕)	Estamos moderadamente seguros en la estimación del efecto: el verdadero efecto es probable que esté cerca de la estimación del efecto, pero hay una posibilidad de que sea sustancialmente diferente.
Baja (⊕⊕)	Nuestra confianza en la estimación del efecto es limitada: el verdadero efecto puede ser sustancialmente diferente de la estimación del efecto.
Muy Baja (⊕)	Tenemos muy poca seguridad en el efecto estimado: El verdadero efecto es probable que sea sustancialmente diferente de la estimación del efecto.

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Método GRADE: Grados de recomendación



Tipos de recomendación	Definición
Fuerte	El grupo elaborador confía en que los efectos beneficiosos que se pueden dar como resultado de adherirse/llevar a cabo una recomendación son mayores que los efectos perjudiciales.
Débil (Condicional, Discrecional)	El grupo elaborador concluye , aunque no está completamente seguro, que los potenciales efectos beneficiosos de llevar a cabo una recomendación probablemente son mayores que los potenciales efectos perjudiciales.

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Ejemplo: Evaluation, Treatment, and Prevention of Vitamin D Deficiency:
an Endocrine Society Clinical Practice Guideline

<https://www.ncbi.nlm.nih.gov/pubmed/21646368>

1912 Holick et al.

Guidelines on Vitamin D Deficiency

J Clin Endocrinol

2.0 Recommended dietary intakes of vitamin D for patients at risk for vitamin D deficiency

2.1 We suggest that infants and children aged 0–1 yr require at least 400 IU/d (IU = 25 ng) of vitamin D and children 1 yr and older require at least 600 IU/d to maximize bone health. Whether 400 and 600 IU/d for children aged 0–1 yr and 1–18 yr, respectively, are enough to provide all the potential nonskeletal health benefits associated with vitamin D to maximize bone health and muscle function is not known at this time. However, to raise the blood level of 25(OH)D consistently above 30 ng/ml (75 nmol/liter) may require at least 1000 IU/d of vitamin D (2).

2.2 We suggest that adults aged 19–50 yr require at least 600 IU/d of vitamin D to maximize bone health and muscle function. It is unknown whether 600 IU/d is enough to provide all the potential nonskeletal health benefits associated with vitamin D. However, to raise the

vitamin D₂ or vitamin D₃, or vitamin D₃ once weekly of 25(OH)D above 30 ng/ml therapy of 400–1000 IU/d

3.3 For children aged 1–18 yr, we suggest treatment with vitamin D₃ for at least 1000 IU/d once a week for a blood level of 25(OH)D above 30 ng/ml once weekly therapy of 600–1000 IU/d.

3.4 We suggest that all patients be treated with 50,000 IU/d of vitamin D₃ once a week for 8 wk until the blood level of 25(OH)D above 30 ng/ml weekly therapy of 1500–2000 IU/d (2).

3.5 In obese patients, p

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Evidencia científica

Información contrastada para la toma de decisiones

"¿Quién sabe, preguntó Robert Browning, si el mundo no terminará esta noche?

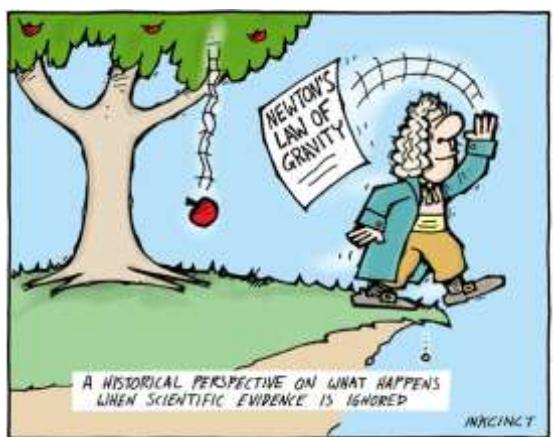
Cierto, pero con la evidencia disponible, la mayoría de nosotros deberíamos prepararnos para salir a trabajar mañana a las 8.30 h."

A.B. Hill

[Schoenbach, 1999]

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Evidencia científica



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