

Primera Reunión Técnica Nacional
“Centros y Servicios de Información de Medicamentos – Gestión del Conocimiento”

OTROS METABUSCADORES

14 y 15 de noviembre del 2011



CenadIM

Centro Nacional de Documentación e Información de Medicamentos

Q.F. María Emilia Ledezma Carbajal

Pirámide de Haynes

METABUSCADORES



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Jon Brassey y el Dr. Chris Precio


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1997

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Jon Brassey
Director, TRIP Database Ltd
Bristol, United Kingdom | Hospital & Health Care

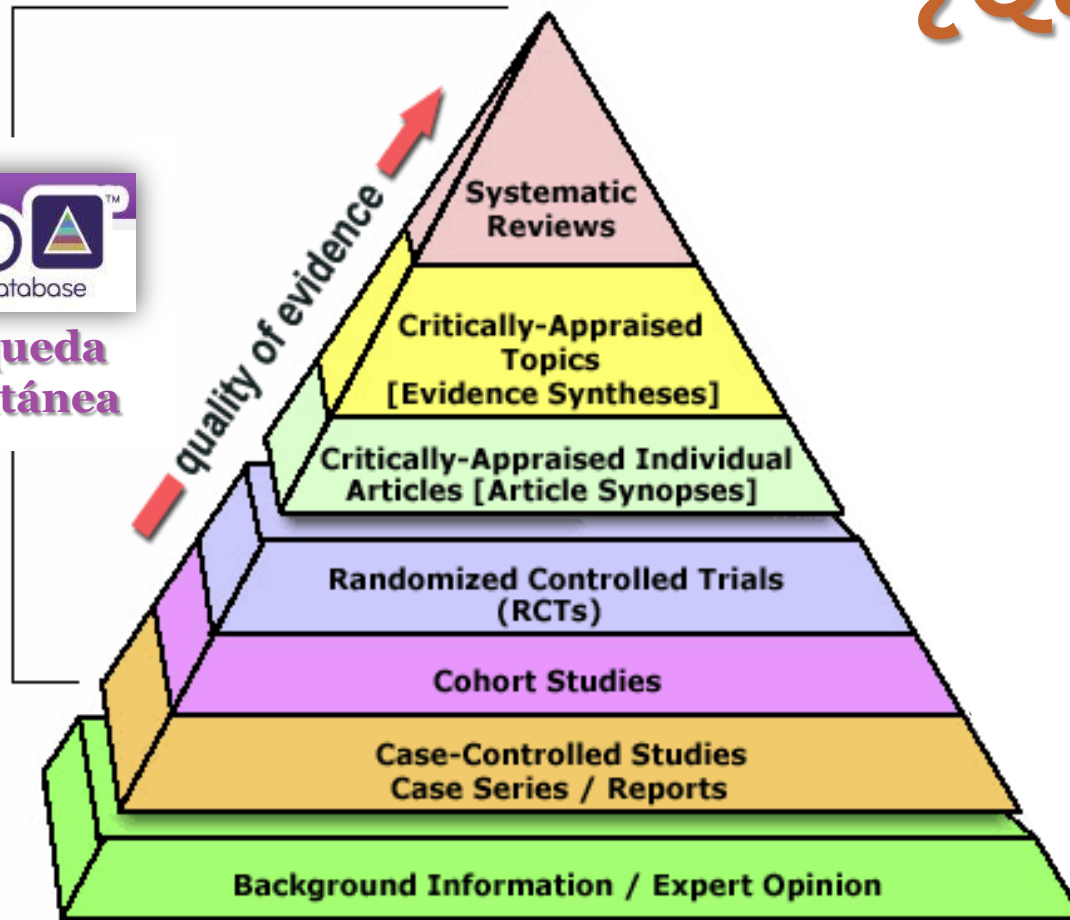
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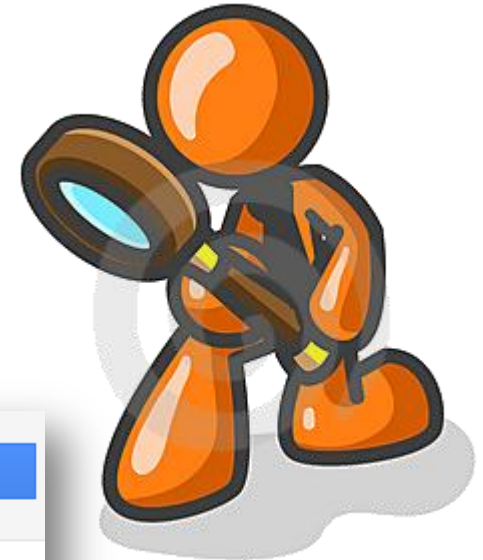


Yale University Library. <http://guides.library.yale.edu/content.php?pid=9786&sid=73113>



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**diabetes obesity
metformin**

diabetes obesity metformin

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All Secondary Evidence 186

Evidence Based Synopses 49

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More 6

News 63

MEDICAL IMAGES 0

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1. [Cost-utility analysis of intensive blood glucose control with metformin versus usual care in overweight type 2 diabetes mellitus patients in Beijing, PR China](#)

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CLINICAL TRIALS [ClinicalTrials.gov](#)

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diabetes obesity metformin

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1. Cost-utility analysis of intensive blood glucose control with metformin versus usual care in overweight type 2 diabetes mellitus patients in Beijing, PR China

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American Family Physician

A peer-reviewed journal of the American Academy of Family Physicians

November 1, 2007 Table of Contents

FPIN's Clinical Inquiries

Metformin Therapy and Diabetes Prevention in Adolescents Who Are Obese

JOSÉ E. RODRÍGUEZ, MD, and BARBARA SHEARER, MLS, Florida State University College of Medicine, Tallahassee, Florida

Clinical Commentary by DAVID C. SLAWSON, MD, University of Virginia Health System, Charlottesville, Virginia

Am Fam Physician. 2007;76(9):1357-1358.

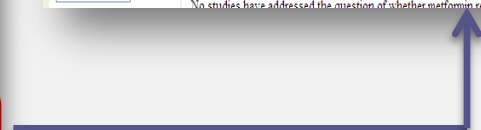
Clinical Question

Does metformin (Glucophage) therapy reduce the likelihood of obese adolescents developing diabetes?

Evidence-Based Answer

No studies have addressed the question of whether metformin reduces development of diabetes among

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End year (inclusive):

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EVIDENCE 465

All Secondary Evidence	111
Evidence Based Synopses	31
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Other	0
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1. Cost-utility analysis of intensive blood glucose control with metformin versus usual care in overweight type 2 diabetes mellitus patients in Beijing, PR China

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Systematic Reviews

- Agency for Healthcare Research and Quality - Evidence-based Practice
- AHRQ - Outcome and Effectiveness
- AHRQ - Preventive Services
- AHRQ - Technology Assessment
- All Wales Medicines Strategy Group
- ASERNIP-S
- Cochrane Database of Systematic Reviews
- DARE
- DARE.
- EPPI Centre
- FFPRHC
- Health Technology Assessment (HTA) Database
- Health Technology Assessment (HTA) Database.
- Institute for Clinical Systems Improvement
- Medical Services Advisory Committee
- National Collaborating Centre for Environmental Health
- National Institute for Health and Clinical Excellence - Interventional Procedures
- National Institute for Health and Clinical Excellence - Medical technologies
- National Institute for Health and Clinical Excellence - Technology Appraisals
- National Institute of Clinical Studies (Australia)
- NHS Economic Evaluation Database.
- NHS EED..
- NHS Quality Improvement Scotland
- NIH Consensus Statements

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There are 4 elements and not all are compulsory. However, we recommend at least 3 boxes are completed.

Population:

This is the type of patient you're interested in e.g. diabetics. However, for this trial version add the disease name e.g. diabetes not diabetics!

osteoporosis

Search

Intervention:

This is any intervention e.g. treatment, diagnostic test

bisphosphonates

Comparison:

This is a comparison intervention; are you comparing your intervention with another treatment or test?

calcium

Outcome:

What outcome are you interested in e.g. reduced mortality, fewer exacerbations

fractures

(title:OSTEOPOROSIS)(title:BISPHOSP

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EVIDENCE

All Secondary Evidence	9
Evidence Based Synopses	0
Systematic Reviews	2
Guidelines	0
Aus. & NZ	0
Canada	0
UK	0
USA	0
Other	0
Clinical Q&A	3
Core primary research	0
Extended primary research	1
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Patient Decision Aids	0
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1. The efficacy of bisphosphonates in the prevention of vertebral, hip, and nonvertebral-nonhip fractures in osteoporosis: a network meta-analysis
DARE. 2011
CPD/CME
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2. In a man with a previous hip fracture, should he be on osteoporosis treatment e.g. bisphosphonates and calcium/vitamin D? Or does he need specialist referral and a DEXA scan?
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SUMSearch

CAMBIOS



SUMSearch 2



- 1) Más rápida
- 2) Las estrategias de búsqueda son validadas
- 3) Las Guías de Práctica Clínica de PubMed y del National Guidelines Clearinghouse; y Revisiones Sistemáticas de DARE (Database of Abstracts of Reviews of Effects) y del PubMed se han fusionado en una sola lista ordenada por año de publicación.

¿Qué busca?

Busca simultáneamente:

- Estudios originales
- Revisiones sistemáticas y
- Guías de práctica clínica en varios sitios web.

Siempre se encuentra actualizado.



SUMSearch Fusiona y ordena resultados de **PubMed**, **Dare**, y **NGC**



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**diabetes obesity
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Focus: Intervention Diagnostic

Age: Adult Pediatric

Human only: English only: Require abstracts:

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[Heart Groups](#) (posted)

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NEJM - Cases with CME

1. [A 56-Year-Old Man with Hypophosphatemia](#) (posted 2011-10-27)
2. [A 55-Year-Old Man with Oligometastatic Lung Cancer](#) (posted 2011-10-13)
3. [A 62-Year-Old Woman with Renal Failure](#) (posted 2011-09-29)
4. [A 66-Year-Old Woman with Cardiac and Renal Failure](#) (posted 2011-09-22)
5. [A 74-Year-Old Man with Pemphigus Vulgaris and Lung Nodules](#) (posted 2011-09-15)
6. [A 17-Year-Old Boy with Abdominal Pain and Weight Loss](#) (posted

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“Filtrar”

Original studies

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60 possible original studies PubMed found after 4 searches. The first 50 citations are:

1. Effectiveness of primary care-relevant treatments for obesity in adults: a systematic evidence review for the U.S. preventive services task force.

Ann Intern Med 2011;155:7. PMID: [21969342](#) , doi: [10.1059/0003-4819-155-7-201110040-00006](#). [Cite](#)

Conclusion: Behaviorally based treatments are safe and effective for weight loss and maintenance.
Primary Funding Source: Agency for Healthcare Research and Quality.

2. Current perspectives of insulin resistance and polycystic ovary syndrome.

Diabet Med 2011;:. PMID: [21950959](#) , doi: [10.1111/j.1464-5491.2011.03460.x](#). [Cite](#)

Conclusion: Insulin resistance is linked to polycystic ovary syndrome. Further study of lifestyle and pharmacologic interventions that reduce insulin resistance, such as metformin, are needed to demonstrate that they are effective in reducing the risk of diabetes, endometrial abnormalities and cardiovascular disease events in women with polycystic ovary syndrome.


3. Metformin should be considered in the treatment of gestational diabetes: a prospective randomised study.

BJOG 2011;118:7. PMID: [21083860](#) , doi: [10.1111/j.1471-0528.2010.02763.x](#). [Cite](#)

Conclusion: Metformin seems to be suitable for the prevention of fetal macrosomy, especially in lean or moderately overweight women developing GDM in late gestation. Women with considerable obesity, high fasting blood glucose and an early need for pharmacological treatment may be more suitable for insulin therapy.

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4. Metformin extended release treatment of adolescent obesity: a 48-week randomized, double-blind, placebo-controlled trial with 48-week follow-up.

Arch Pediatr Adolesc Med 2010;164:2 PMID: 20124139 doi: 10.1001/archpediatrics.2009.264. Cite

Conclusion: Metformin XR caused a small but statistically significant decrease in BMI when added to a lifestyle intervention program.

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Display Settings: Abstract Send to

Arch Pediatr Adolesc Med. 2010 Feb;164(2):116-23.

Metformin extended release treatment of adolescent obesity: a 48-week randomized, double-blind, placebo-controlled trial with 48-week follow-up.

Wilson DM, Abrams SH, Aye T, Lee PD, Lenders C, Lustig RH, Osoanian SV, Feldman HA, Glaser Pediatric Research Network Obesity Study Group.

Collaborators (33)

Division of Pediatric Endocrinology and Diabetes, Stanford University and the Lucile Packard Children's Hospital at Stanford, G-311 Medical Center, Stanford, CA 94305-5205, USA. d.wilson@stanford.edu

Abstract

BACKGROUND: Metformin has been proffered as a therapy for adolescent obesity, although long-term controlled studies have not been reported.

OBJECTIVE: To test the hypothesis that 48 weeks of daily metformin hydrochloride extended release (XR) therapy will reduce body mass index (BMI) in obese adolescents, as compared with placebo.

DESIGN: Multicenter, randomized, double-blind, placebo-controlled clinical trial.

SETTING: The 6 centers of the Glaser Pediatric Research Network from October 2003 to August 2007.

PMID (PubMed Unique Identifier)

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Vol. 164 No. 2, February 2010

Article

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Metformin Extended Release Treatment of Adolescent Obesity

A 48-Week Randomized, Double-Blind, Placebo-Controlled Trial With 48-Week Follow-up

Glaser Pediatric Research Network Obesity Study Group

Arch Pediatr Adolesc Med. 2010;164(2):116-123.

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Metformin Extended Release Treatment of Adolescent Obesity

A 48-Week Randomized, Double-Blind, Placebo-Controlled Trial With 48-Week Follow-up

Glaser Pediatric Research Network Obesity Study Group

Background: Metformin has been proffered as a therapy for adolescent obesity, although long-term controlled studies have not been reported.

Objective: To test the hypothesis that 48 weeks of daily metformin hydrochloride extended release (XR) therapy will reduce body mass index (BMI) in obese adolescents, as compared with placebo.

Design: Multicenter, randomized, double-blind, placebo-controlled clinical trial.

Setting: The 6 centers of the Glaser Pediatric Research Network from October 2003 to August 2007.

Participants: Obese (BMI \geq 95th percentile) adolescents (aged 13-18 years) were randomly assigned to the intervention (n=39) or placebo groups.

Intervention: Following a 1-month run-in period, sub-

domized 1:1 to 48 weeks' treatment with metformin hydrochloride XR, 2000 mg once daily, or an identical placebo. Subjects were monitored for an additional 48 weeks.

Main Outcome Measure: Change in BMI, adjusted for site, sex, race, ethnicity, and age and metformin vs placebo.

Results: After 48 weeks, mean (SE) adjusted BMI increased 0.2 (0.5) in the placebo group and decreased 0.9 (0.5) in the metformin XR group ($P=.03$). This difference persisted for 12 to 24 weeks after cessation of treatment. No significant effects of metformin on body composition, abdominal fat, or insulin indices were observed.

Conclusion: Metformin XR caused a small but statistically significant decrease in BMI when added to a lifestyle intervention program.

Trial Registration: clinicaltrials.gov Identifiers: NCT00209482 and NCT00120146.

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	Year	Database	Record type	Title
<input type="checkbox"/>	2011	DARE	Systematic review	Effectiveness of primary care-relevant treatments for obesity in adults: a systematic evidence review for the US preventive services task force [Preview]

[of Effects](#)

The systematic reviews are a list merged from [PubMed](#) (using the [systematic reviews subset strategy](#)) and [Database of Abstracts of Reviews of Effects \(DARE\)](#) in approximate order of date. Duplicates may occur if a guideline is published both in PubMed and DARE.

Systematic evidence
ID: [21969342](#) [DARE](#)

Centre for Reviews and Dissemination

Effectiveness of primary care-relevant treatments for obesity in adults: a systematic evidence review for the US preventive services task force
LeBlanc ES, O'Connor E, Whitlock EP, Patnode CD, Kapka T

CRD summary
This review concluded that behavioural interventions were safe and effective for weight loss and maintenance. The authors' conclusions reflect the evidence presented, but this evidence had limitations, such as poor reporting, high rates of withdrawal, and variability between studies, that should be borne in mind when interpreting the conclusions.

Authors' objectives
To assess the effectiveness and harms of weight loss interventions, for overweight and obese adults, conducted in settings relevant to primary care.

Searching
The authors used published systematic reviews and guidelines to identify studies published up to 2005. They searched MEDLINE, PsycINFO and the Cochrane Central Register of Controlled Trials (CENTRAL) for articles from 2005 to September 2010. Search terms were included in the full report (LeBlanc, et al. 2011, see 'Other Publications of Related Interest'). The search was supplemented by consulting relevant systematic reviews, experts, and reference lists. It was restricted to English-language studies.

Expert model. Diabetes
Screening Committee.
Form Metab Res. 2010
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[DARE summary](#) if
ring? Curr Pharm Des.

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Guidelines

19 guidelines from NGC found.

1 possible guidelines found from PubMed ([View at PubMed](#))

Merged list:

1. **American Association of Clinical Endocrinologists medical guidelines for clinical practice for developing a diabetes mellitus comprehensive care plan**
American Association of Clinical Endocrinologists - Medical Specialty Society. 2011. NGC: [008577](#)
2. **Standards of medical care in diabetes. VII. Diabetes care in specific populations**

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The screenshot shows the National Guideline Clearinghouse interface. At the top, there is a navigation bar with links for Help, RSS, Subscribe to weekly e-mail, Site map, Contact us, and For web developers. Below this is a search bar with a 'Search' button and links for Search Tips, Advanced Search, and About Search. The main content area is titled 'Guideline Summary' and includes options for Print, Download as PDF (Adobe Reader), Word, HTML, XML, and Export to Citation Manager. The 'Guideline Title' is 'American Association of Clinical Endocrinologists medical guidelines for clinical practice for developing a diabetes mellitus comprehensive care plan.' The 'Bibliographic Source(s)' section lists the authors and publication details: 'Handelsman Y, Mechanick JI, Blonde L, Grunberger G, Bloomgarden ZT, Bray GA, Dagogo-Jack S, Davidson JA, Einhorn D, Ganda O, Garber AJ, Hirsch IB, Horton ES, Ismail-Beigi F, Jellinger PS, Jones KL, Jovanovic L, Lebovitz H, Levy P, Moghissi ES, Orzeck EA, Vinik AI, Wyne KL, AACE Task Force for Developing Diabetes Comprehensive Care Plan. American Association of Clinical Endocrinologists medical guidelines for clinical practice for developing a diabetes mellitus comprehensive care plan. Endocr Pract 2011 Mar-Apr;17 (Suppl 2):1-53. [375 references] PubMed'. The 'Guideline Status' section states 'This is the current release of the guideline.'

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Min Hae Park, MSC¹, Sanjay Kinra, MD, PHD¹, Kirsten J. Ward, PHD², Billy White, MBBS² and Russell M. Viner, MBBS, PHD²

Author Affiliations

Corresponding author: Russell M. Viner, r.viner@ich.ucl.ac.uk

Abstract

OBJECTIVE To summarize the efficacy of metformin in reducing BMI and cardiometabolic risk in obese children and adolescents without diabetes.

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Metformin for Obesity in Children and Adolescents: A Systematic Review

MIN HAE PARK, MSC¹
SANJAY KINRA, MD, PHD¹
KIRSTEN J. WARD, PHD²

BILLY WHITE, MBBS³
RUSSELL M. VINER, MBBS, PHD³

OBJECTIVE — To summarize the efficacy of metformin in reducing BMI and cardiometabolic risk in obese children and adolescents without diabetes.

RESEARCH DESIGN AND METHODS — We performed a systematic review and meta-analysis of randomized controlled trials (RCTs). Double-blind RCTs of ≥6 months duration in obese subjects age ≤19 years without diabetes were included. Our primary outcomes of interest include changes in BMI and measures of insulin sensitivity.

RESULTS — Five trials met inclusion criteria (n = 320 individuals). Compared with placebo, metformin reduced BMI by 1.42 kg/m² (95% CI 0.83–2.02) and homeostasis model assessment insulin of resistance (HOMA-IR) score by 2.01 (95% CI 0.75–3.26).

CONCLUSIONS — Metformin appears to be moderately efficacious in reducing BMI and insulin resistance in hyperinsulinemic obese children and adolescents in the short term. Larger, longer-

A random-effects model was selected. Sensitivity analyses were performed using fixed-effects models and by dose of metformin (1,000 vs. 2,000 mg), age of participants (12–19 vs. <12 years), co-intervention (metformin vs. metformin + co-intervention), baseline BMI (mean ≥35 vs. <35 kg/m²), and by excluding one study reporting greater treatment effects than the other studies (4).

RESULTS — Five studies published between 2001 and 2008 met the inclusion criteria (4–8). This included one crossover trial (5). Three studies took place in the U.S. (6–8) and one each in Australia (5) and

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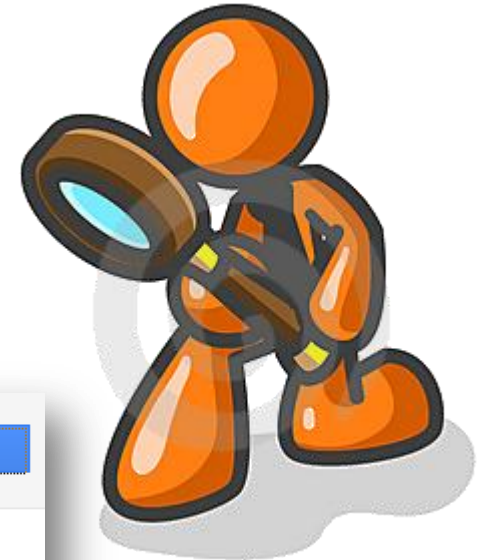
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Silvio E. Inzucchi, M.D., David G. Maggs, M.D., Cheryl Veronika Walton, B.A., and Gerald I. Shulman, M.D., Ph.D. | N Engl J Med 1998; 338:867-873 | March 26, 1998

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BACKGROUND

Combination therapy is logical for patients with type II diabetes mellitus, because it allows for the use of single-drug therapy. We studied the efficacy and physiologic effects of metformin and troglitazone alone and in combination in patients with type II diabetes.

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METHODS

EFFICACY AND METABOLIC EFFECTS OF METFORMIN AND TROGLITAZONE IN TYPE II DIABETES MELLITUS

EFFICACY AND METABOLIC EFFECTS OF METFORMIN AND TROGLITAZONE IN TYPE II DIABETES MELLITUS

SILVIO E. INZUCCHI, M.D., DAVID G. MAGGS, M.D., GERALYN R. SPOLLETT, A.P.R.N., STEPHANIE L. PAGE, R.N., FRANCES S. RIFE, R.N., VERONIKA WALTON, B.A., AND GERALD I. SHULMAN, M.D., PH.D.

ABSTRACT

Background Combination therapy is logical for patients with non-insulin-dependent (type II) diabetes mellitus, because they often have poor responses to single-drug therapy. We studied the efficacy and physiologic effects of metformin and troglitazone alone and in combination in patients with type II diabetes.

Methods We randomly assigned 29 patients to receive either metformin or troglitazone for three months, after which they were given both drugs for another three months. Plasma glucose concentrations during fasting and postprandially and glycosylated hemoglobin values were measured periodically during both treatments. Endogenous glucose production and peripheral glucose disposal were measured at base line and after three and six months.

HYPERGLYCEMIA in patients with non-insulin-dependent (type II) diabetes mellitus is caused by peripheral insulin resistance, which results in decreased insulin-mediated glucose disposal; increased endogenous glucose production, chiefly from the liver; and inadequate pancreatic insulin secretion.¹ Reversal of these defects, either individually or in concert, improves glycemic control. New drugs are now available that affect each of these defects separately, and an understanding of their mechanisms of action is important for their proper use, especially when they are administered in combination.

Until recently in the United States, the only oral drugs available for patients with type II diabetes were sulfonylureas. These increase insulin secretion.²

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JOHN E. NESTLER, M.D., DANIELA J. JAKUBOWICZ, M.D., WILLIAM S. EVANS, M.D., AND RENATO PASQUALI, M.D.

ABSTRACT

Background Obese women with the polycystic ovary syndrome are relatively unresponsive to the induction of ovulation by clomiphene. We hypothesized that reducing insulin secretion by administering metformin would increase the ovulatory response to clomiphene.

Methods We performed oral glucose-tolerance tests before and after the administration of 500 mg of metformin or placebo three times daily for 35 days in 61 obese women with the polycystic ovary syndrome. Women who did not ovulate spontaneously were then given 50 mg of clomiphene daily for five days while continuing to take metformin or placebo. Serum progesterone was measured on days

drome²⁻⁵ and appears to have a pathophysiologic role in the hyperandrogenism of the disorder. Ovarian androgen production and serum free testosterone concentrations decrease in women with polycystic ovary syndrome when insulin secretion is reduced by drugs such as diazoxide,⁶ metformin,⁷⁻¹⁰ and troglitazone.^{11,12} However, whether such therapy improves ovulatory function is not known.

Clomiphene citrate, an antiestrogenic drug, is the primary therapy used for ovulation induction in women with the polycystic ovary syndrome.¹³⁻¹⁵ However, obese women with the syndrome often require multiple courses and high doses of clomiphene, and there is a positive correlation between

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J.P. Després ^{1, 2, 3}

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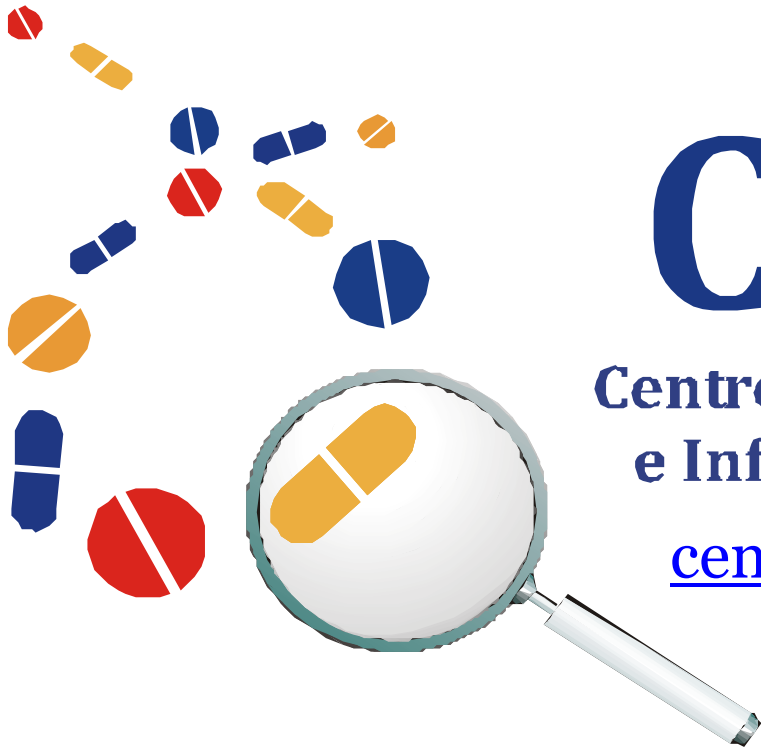


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