"Medicina e Saúde: Perspectivas actuais e futuras"

José M. D. Poças
Médico Internista e Infecciologista
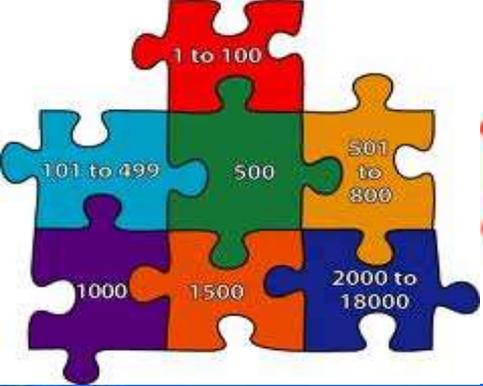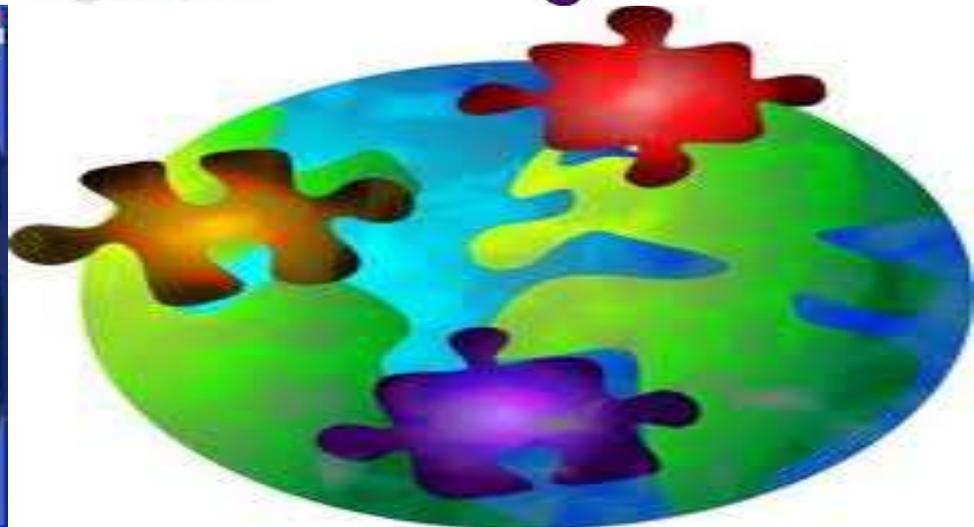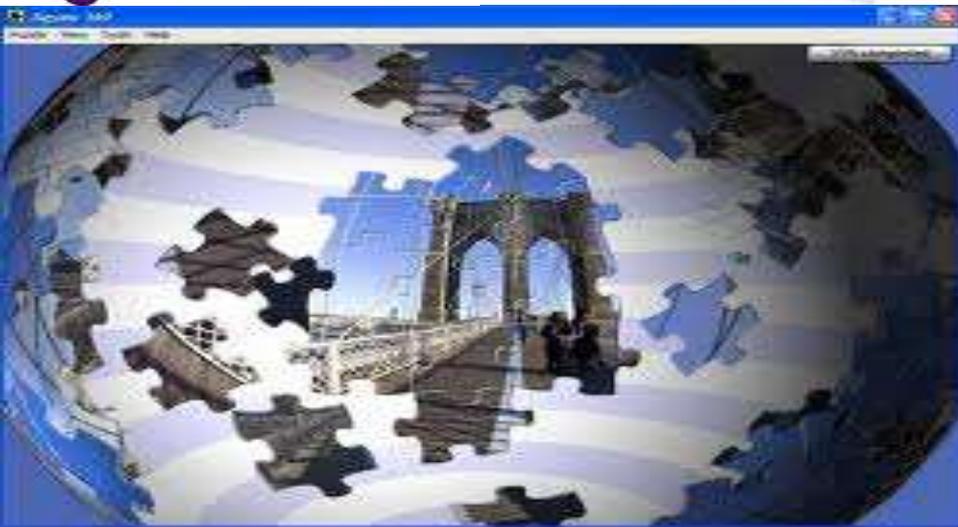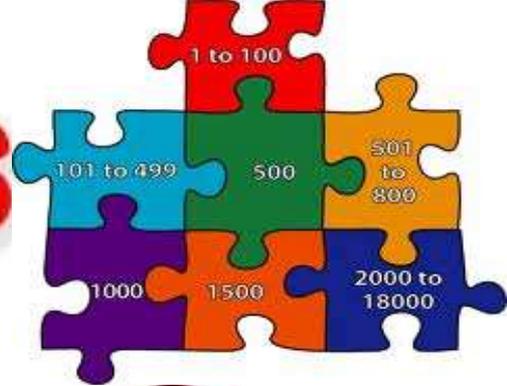
PRIMEIROS PENSAMENOS

- *" ... Time will say nothing but I told you so,
Time only knows the price we have to pay,
If I could tell you I would let you know ... " **(sic.)*** (Wystan Auden, Poeta, 1966)*

- *" ... o Homem só vê verdadeiramente aquilo que ele já conhece ...
saber não é suficiente; devemos aplicar os conhecimentos ...
revelar vontade de fazer não é bastante; o que temos mesmo é de
executar ... " **(sic.)*** (Johann W. Goethe)*

TÓPICOS da CONFERÊNCIA

INTRODUÇÃO
PROBLEMAS de ONTEM, de HOJE e de AMANHÃ
Os PRINCIPAIS PROBLEMAS ACTUAIS
Os PRINCIPAIS PROBLEMAS do FUTURO
E ENTÃO em PORTUGAL?
As RESPOSTAS da INOVAÇÃO CIENTÍFICA
PROBLEMAS DECORRENTES da TECNOLOGIA
CONCLUSÕES
MENSAGENS FINAIS

Can DNA Stop Time?
UNLOCKING THE SECRETS OF **LONGEVITY GENES**

Know Your DNA
Inexpensive **gene readers** will soon
unlock the secrets in your
personal double helix

'A História depende da nossa forma de pensar o futuro'

Como se escreve a História, hoje em dia? Num mundo desprovido de horizonte utópico, a historiografia é hoje em dia dominada pela noção de memória, explica o historiador Enzo Traverso numa obra sobre as violências do século XX.

REVISTA POLITIS
PARIS

UTOPIA

Enzo Traverso
1914-1945
La guerre civile européenne

Enzo Traverso
Understanding the Nazi Genocide
Marxism after Auschwitz

INTRODUÇÃO

Escrito na pedra

E cada um acredita, facilmente, no que teme e no que deseja.

Jean de La Fontaine, escritor francês (1621-1695)

FABLES CHOISIES.

MISES EN VERS

Par M. de la Fontaine.

A PARIS,

Chez CLAUDE BARBIN, au Palais sur le Perron de la sainte Chapelle.

M. DC. LXVIII.

AVEC PRIVILEGE DV ROY.

Marc Chagall
Fables of La Fontaine

THE ORIGINAL
FABLES
OF LA FONTAINE

RENDERED INTO ENGLISH PROSE
BY
FREDK. COLIN TILNEY

WITH COLOURED ILLUSTRATIONS
BY THE AUTHOR

LONDON: J. M. DENT & SONS LIMITED
NEW YORK: E. P. DUTTON & COMPANY

MAIS DOIS PENSAMENTOS

- *" … o custo da inacção é evidente e inaceitável. Através de um investimento sério e determinado numa política sectorial adequadamente direccionada às principais causas de morbi-mortalidade preveníveis, poder-se-á registar um significativo impacto positivo na vida das diversas populações à escala planetária … " (SIC.)* (Lee Jong-Wook, Secretário Geral da OMS, 2005)

- *" … calcula-se que cerca de 70 a 80 % das causas Cardio/Cerebro-Vasculares ou Diabetes, bem como 30 a 40 % das causas de Cancro , 40 % das Doenças Crónicas Respiratórias, e cerca de 50 % das restantes causas de morte por causas secundárias a outras Doenças Crónicas, sejam preveníveis com a adopção de hábitos de vida mais saudáveis e vigilância periódica da saúde … " (sic.)* (N. Unwin, and G. Alberti, 2006, Dele O. Abegunde, et al, 2007)

Annals of Tropical Medicine & Parasitology, Vol. 100, Nos. 5 and 6, 455–464 (2006)

CENTENNIAL REVIEW

Chronic non-communicable diseases

N. UNWIN* and K. G. M. M. ALBERTI*

*School of Population and Health Sciences, University of Newcastle upon Tyne, William Leech Building, Framlington Place, Newcastle upon Tyne NE2 4HH, U.K.
*Department of Endocrinology and Metabolic Medicine, Imperial College London, St Mary's Hospital, Mint Wing, Praed Street, London W2 1NY, U.K.

Received 12 June 2006, Accepted 13 June 2006

Chronic non-communicable diseases (NCD) account for almost 60% of global mortality, and 80% of deaths from NCD occur in low- and middle-income countries. One quarter of these deaths — almost 9 million in 2005 — are in men and women aged <60 years. Taken together, NCD represent globally the single largest cause of mortality in people of working age, and their incidences in younger adults are substantially higher in the poor countries of the world than in the rich. The major causes of NCD-attributable mortality are cardiovascular disease (30% of total global mortality), cancers (13%), chronic respiratory disease (7%) and diabetes (2%). These conditions share a small number of behavioural risk factors, which include a diet high in saturated fat and low in fresh fruit and vegetables, physical inactivity, tobacco smoking, and alcohol excess. In low- and middle-income countries such risk factors tend to be concentrated in urban areas and their prevalences are increasing as a result of rapid urbanization and the increasing globalisation of the food, tobacco and alcohol industries.

Because NCD have a major impact on men and women of working age and their elderly dependents, they result in lost income, lost opportunities for investment, and overall lower levels of economic development. Reductions in the incidences of many NCD and their complications are, however, already possible. Up to 80% of all cases of cardiovascular disease or type-2 diabetes and 40% of all cases of cancer, for example, are probably preventable based on current knowledge. In addition, highly cost-effective measures exist for the prevention of some of the complications of established cardiovascular disease and diabetes. Achieving these gains will require a broad range of integrated, population-based interventions as well as measures focused on the individuals at high risk. At present, the international-assistance community provides scant resources for the control of NCD in poor countries, partly, at least, because NCD continue to be wrongly perceived as predominantly diseases of the better off. As urbanization continues apace and populations age, investment in the prevention and control of NCD in low-and middle-income countries can no longer be ignored.

Chronic Diseases 1

The burden and costs of chronic diseases in low-income and middle-income countries

Dele O Abegunde, Colin D Mathers, Taghreed Adam, Monica Ortegon, Kathleen Strong

This paper estimates the disease burden and loss of economic output associated with chronic diseases—mainly cardiovascular diseases, cancer, chronic respiratory diseases, and diabetes—in 23 selected countries which account for around 80% of the total burden of chronic disease mortality in developing countries. In these 23 selected low-income and middle-income countries, chronic diseases were responsible for 50% of the total disease burden in 2005. For 15 of the selected countries where death registration data are available, the estimated age-standardised death rates for chronic diseases in 2005 were 54% higher for men and 86% higher for women than those for men and women in high-income countries. If nothing is done to reduce the risk of chronic diseases, an estimated US$84 billion of economic production will be lost from heart disease, stroke, and diabetes alone in these 23 countries between 2006 and 2015. Achievement of a global goal for chronic disease prevention and control—an additional 2% yearly reduction in chronic disease death rates over the next 10 years—would avert 24 million deaths in these countries, and would save an estimated $8 billion, which is almost 10% of the projected loss in national income over the next 10 years.

O PLANETA e a SOCIEDADE estão DOENTES ...

70%

of commercial species of fish have already been fished beyond their reproductive capacity.

Almost

20%

of coral reefs in the world have disappeared, and 56% are threatened.

Tropical forests provide a habitat for

50%

of the world's known species. Half the Amazon forest could disappear by **2050**.

13 million

hectares of natural forest disappear in the world every year. That is the equivalent of the surface area of Greece.

PLUS DE

100 MILLIONS

DE PERSONNES VIVENT EN DESSOUS DU **NIVEAU DE LA MER** OU SUR DES TERRES SITUÉES À MOINS D'UN MÈTRE AU-DESSUS.

Courrier *internacional*

JAPÃO
DEPOIS DO SISMO
O PERIGO ATÓMICO

PRIMAVERA
ÁRABE
POSTA À PROVA
NA LÍBIA

BOAT PEOPLE

THE OFTEN-OVERLOOKED
FACTS ABOUT
BOAT PEOPLE

Come and work

GO BACK HOME

Distance no object

Some of the world's more important current migration routes

Sources: National Public Radio; The Economist
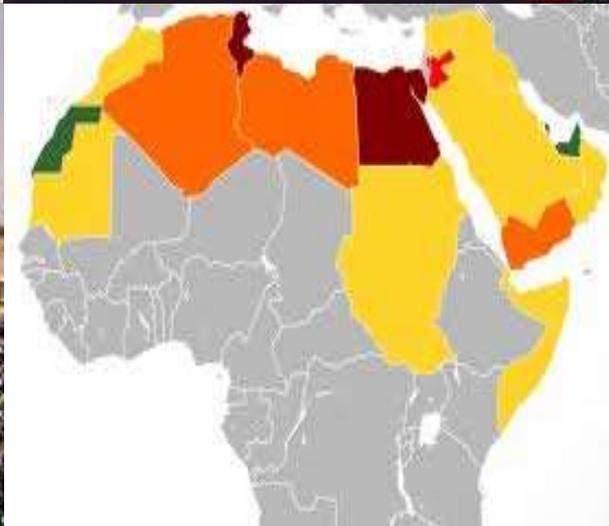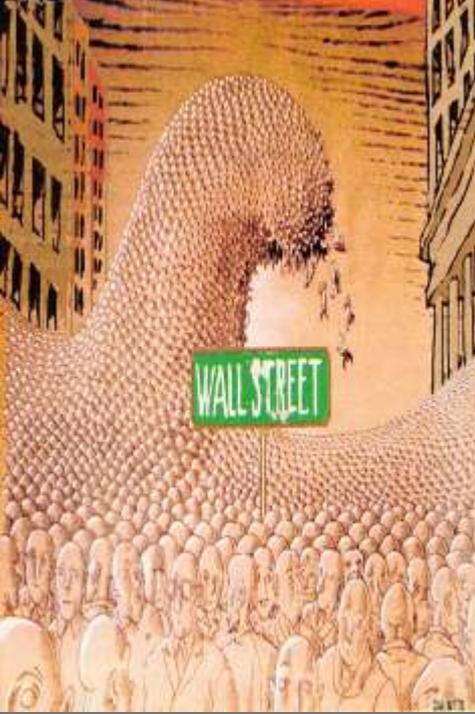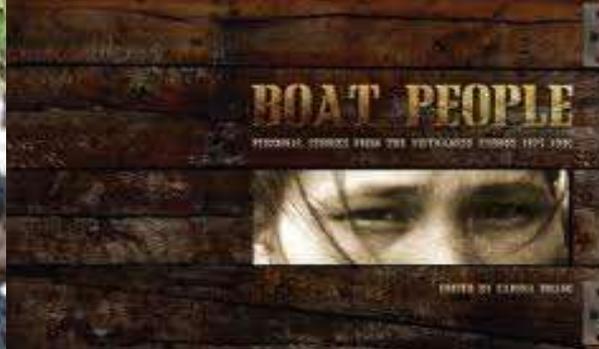
Displaced by violence or disaster

Haiti
500,000 in refugee camps

Libya
440,000 fled to Tunisia, Egypt and European contries bordering the Meditterranean

Iraq
1.5 million refugees, partly in Syria, Jordan and Lebanon.

Afghanistan
319,000 in the country alone

Sudan (Darfur)
4.9 million, mainly in country itself

Colombia
3 million refugees. Partly fled to Ecuador and Venezuela

Ivory Coast
Some 700,000, mainly in country itself

Uganda
Nearly 1 million refugees

Myanmar
1.5 million in the country itself. 340,000 to Malaysia, Thailand and Bangladesh

Central African Republic
200,000, more than half in Chad and Cameroon

© RNW - Tamar Visscher Source: UNHCR   070411

Conflict and Emerging Infectious Diseases

Michelle Gayer,* Dominique Legros,* Pierre Formenty,* and Maire A. Connolly*

Legend:
- ● Ebola, Marburg, and CCHF
- ● Influenza (H5N1)
- ○ Lassa fever
- ○ Monkeypox
- ● Nipah and Hendra
- ○ Rift Valley fever
- ○ SARS-CoV
- ○ Yellow fever
- ● Poliovirus
- ★ Plague
- ★ Tularemia
- ▽ Malaria
- ▼ Trypanosomiasis
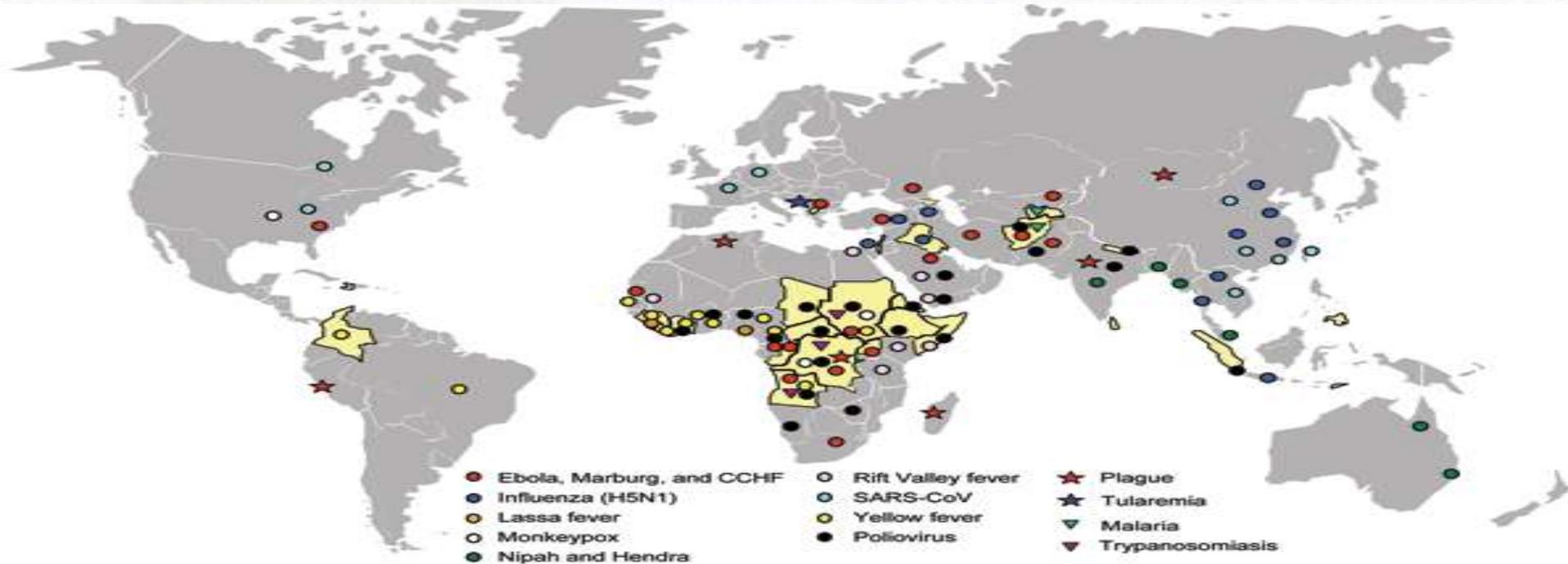
Figure. Geographic distribution of recent emerging or reemerging infectious disease outbreaks and countries affected by conflict, 1990–2006. Countries in yellow were affected by conflict during this period (source: Office for the Coordination of Humanitarian Affairs, World Health Organization, www.reliefweb.int/ocha_ol/onlinehp. html). Symbols indicate outbreaks of emerging or reemerging infectious diseases during this period (source: Epidemic and Pandemic Alert and Response, World Health Organization, www. who.int/csr/en). Circles indicate diseases of viral origin, stars indicate diseases of bacterial origin, and triangles indicate diseases of parasitic origin. CCHF, Crimean-Congo hemorrhagic fever; SARS-CoV, severe acute respiratory syndrome coronavirus.

QUATORZE AGENTS DE BIOTERRORISME OU DE GUERRE BIOLOGIQUE

The virus hierarchy

LEVEL 4: eg: **Lassa fever, Marburg, Ebola Zaire viruses**
No known cure. Highly infectious. Fast acting. High mortality rates.

LEVEL 3: eg: **Yellow fever, dengue haemorrhagic fever virus**
Easily contracted. Potentially fatal, and serious when not fatal.

LEVEL 2: eg: **HIV (Aids)**
Difficult to contract. Easily destroyed. Serious, possibly fatal.

LEVEL 1: eg: **cold viruses**
Relatively innocuous – not fatal.

VIRUSES are infectious packages of genetic material (DNA or RNA) in a protein coat. There is debate about whether they should be considered alive: their only activity is to take over living cells of other organisms, without which they cannot survive. Despite these limitations, viruses plague us: flu, colds, glandular fever, polio, some eye infections, possibly some cancers – all are viruses.

Lassa fever virus

Level 4

Yellow fever virus

Level 3

Human immuno-deficiency virus (HIV)

Level 2

Colony of coronaviruses (responsible for colds)

Level 1

YANKS
SINK
MARINERS

DAILY NEWS

ANTHRAX
NATION

FORMER
SUPER
POWER.
ANYTHING
HELPS

% POPULATION
LIVING ON LESS
THAN $2 PER DAY
- 75 - 92
- 50 - 75
- 28 - 50
- 15 - 28
- 2 - 12
- No Data

www.theodora.com/maps

Population Living Below
National Poverty Line

>60%
50-60%
40-50%
30-40%
20-30%
10-20%
<10%
No data

Source: CIA World Factbook 2008

Population With Sustainable Access to Improved Sanitation (Percent)
2008
(Go to Table or Notes and Sources below)

Population Undernourished (Percent of Total Population)
2005-2007
(Go to Table or Notes and Sources below)
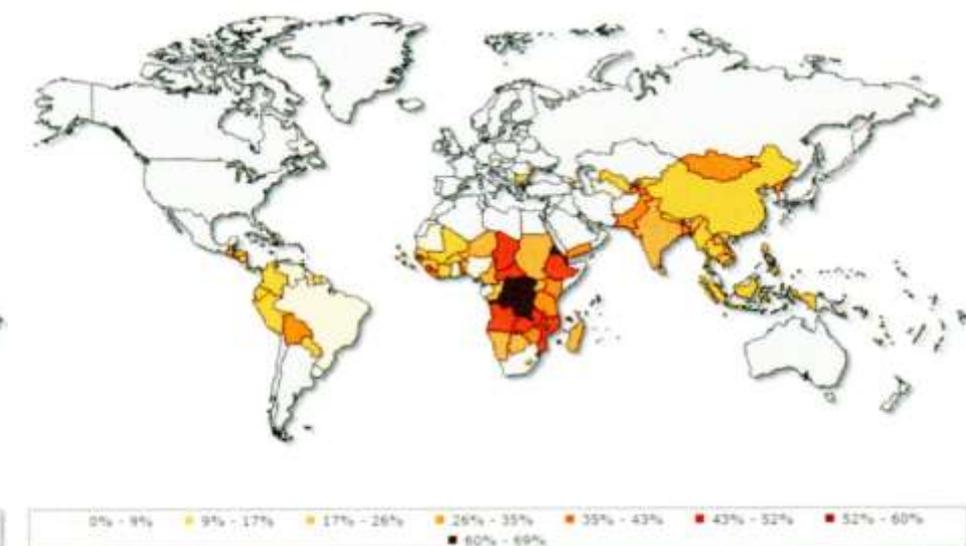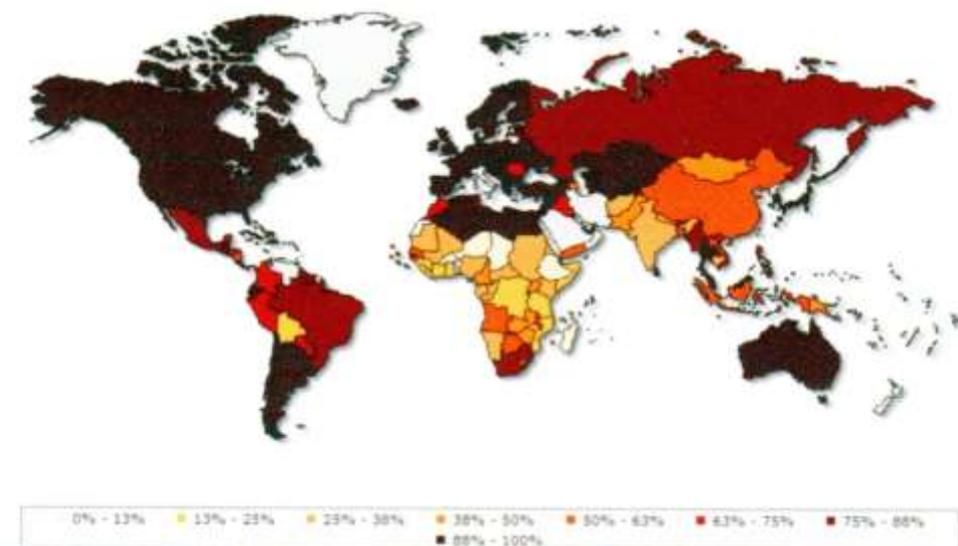
0% - 13%   13% - 25%   25% - 38%   38% - 50%   50% - 63%   63% - 75%   75% - 88%
88% - 100%

0% - 9%   9% - 17%   17% - 26%   26% - 35%   35% - 43%   43% - 52%   52% - 60%
60% - 69%

Guerra de audiências

OS NOVOS HERÓIS

PESO PESADO

www.sic.sapo.pt
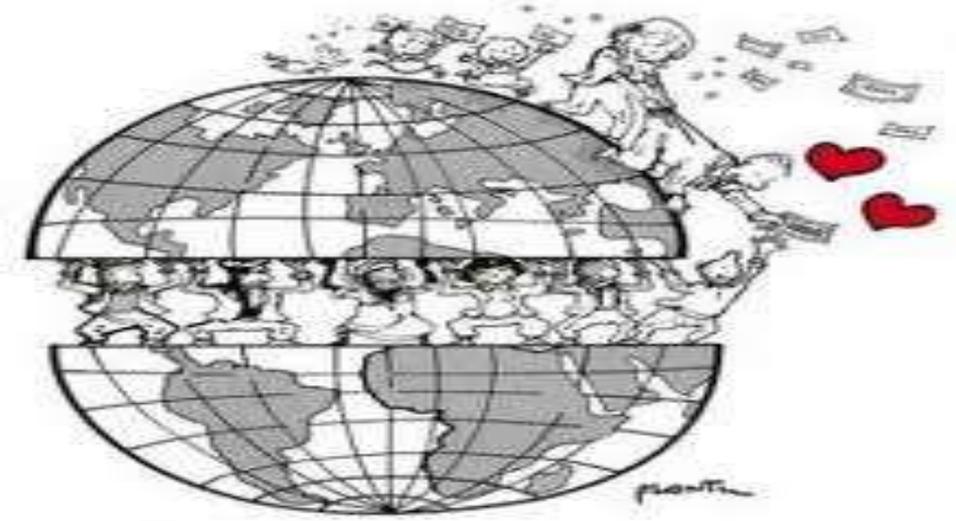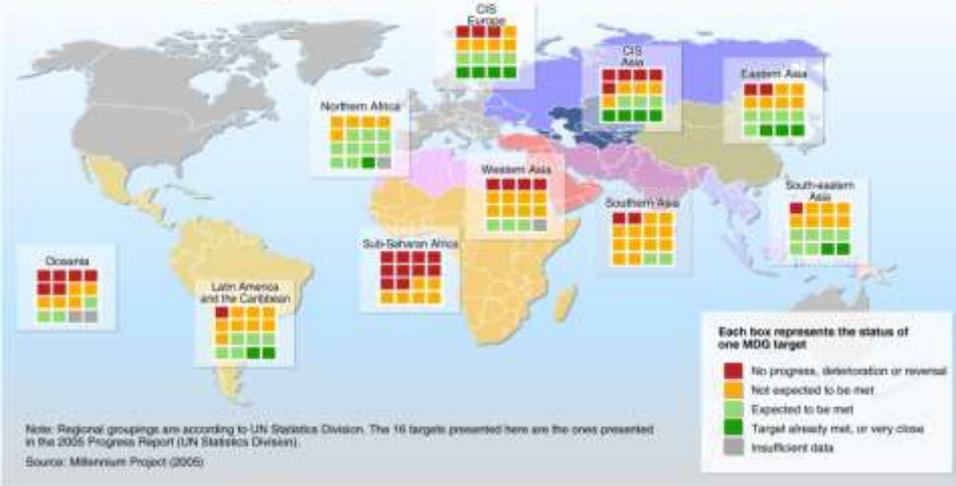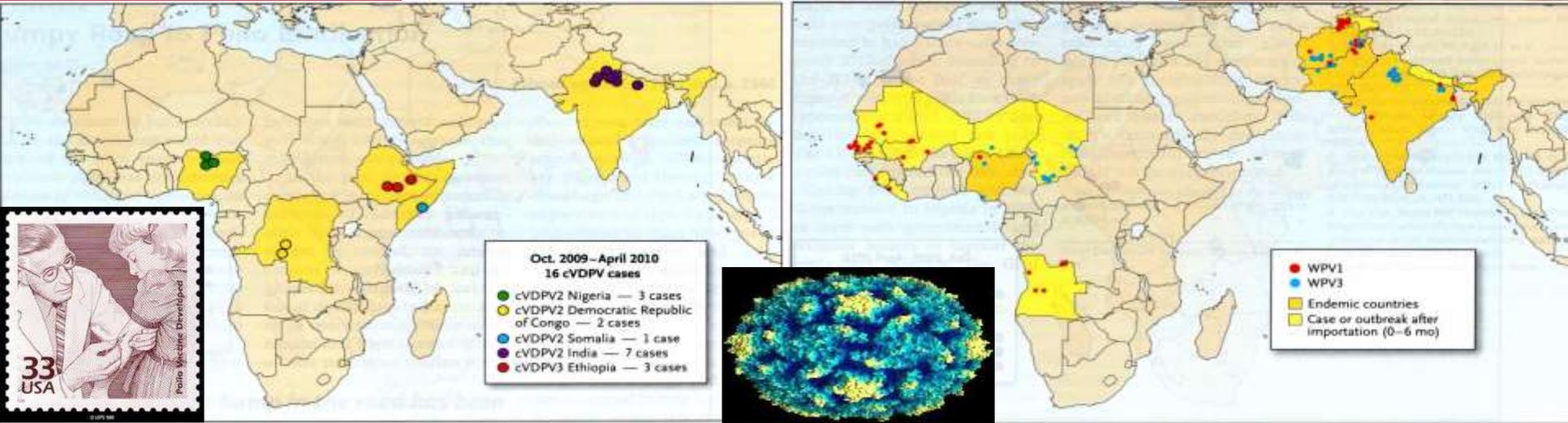
Figure 2. Current Outbreaks of Circulating Vaccine-Derived Poliomyelitis.
The abbreviation cVDPV denotes circulating vaccine-derived poliovirus, and cVDPV2 and cVDPV3 denote types 1 and 3, respectively. Data are from the WHO Global Poliomyelitis Eradication Initiative.

Figure 1. Global Distribution of Wild-Type Poliomyelitis Cases, January 1, 2010–June 1, 2010.
WPV1 and WPV3 denote cases of wild-type poliovirus types 1 and 3, respectively. The total number of cases is 254. Data are from the WHO Global Poliomyelitis Eradication Initiative.

WORLD DRUG REPORT 2010

UNODC
United Nations Office on Drugs and Crime

Main problem drugs as reflected in treatment demand, by region, from the late 1990s to 2008 (or latest year available)

North America
Late 1990s 42%
2008 31%

Europe
Late 1990s 72%
2008 57%

Asia
Late 1990s 73%
2008 62%

Africa
Late 1990s 61%
2008* 63%

South America
Late 1990s 65%
2008 49%

Oceania
Late 1990s 66%
2008 47%

Legend:
- Opiates
- Cannabis
- Cocaine-type
- Amphetamine type
- Others
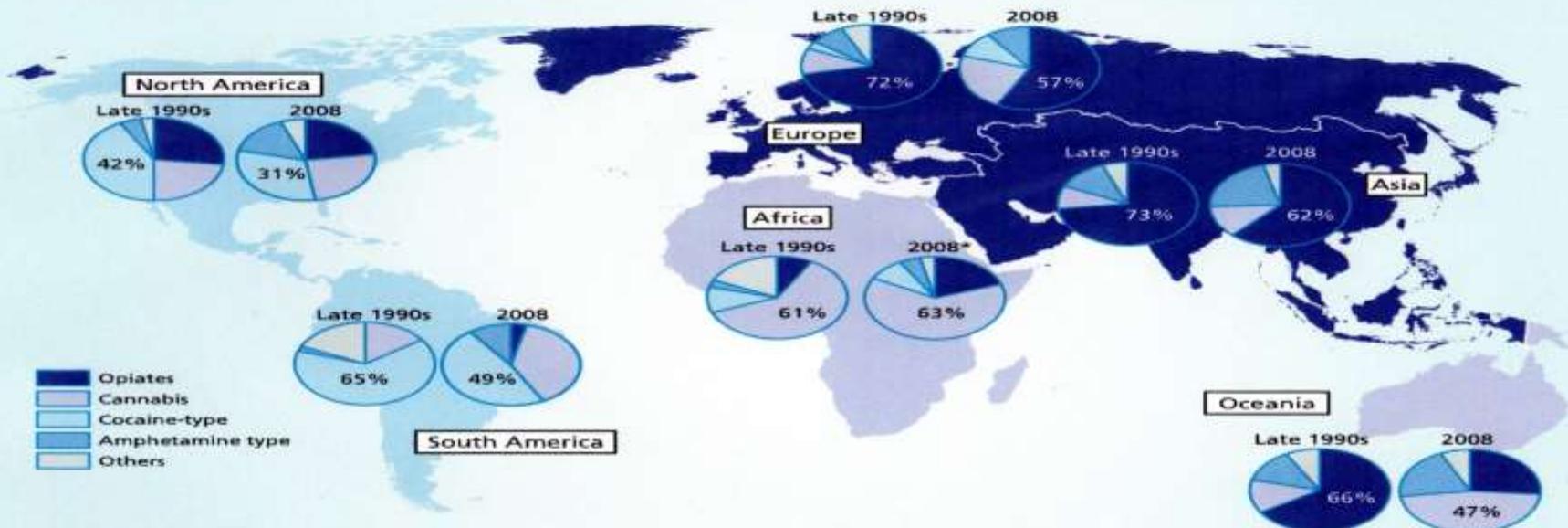
Source: UNODC, Annual Reports Questionnaire Data/DELTA and National Government Reports

Notes: Percentages are unweighted means of treatment demand from reporting countries. Number of countries reporting data for 2008: Europe (45); Africa (26); North America (3); South America (24); Asia (42); Oceania (2). Data generally account for primary drug use. Polydrug use may increase totals beyond 100%. * Treatment data dating back more than 10 years were removed from the 2008 estimates and therefore caution should be taken comparing the data from 2008 with previous years. The boundaries and names shown and the designations used on this map do not imply official endorsement or acceptance by the United Nations.

STRESS & DRUG ABUSE

The world of drugs

Number of Disability-Adjusted Life Years (96s)

- Tobacco
- Alcohol
- Illicit drugs

| | AFRO | AMRO | EMRO | EURO | SEARO | WPRO |

Total users
Numbers of people

190m Cannabis
50m Amphetamines
23m Ecstasy
21m Opiates
20m Cocaine

Main problem drugs
For those being treated

Key
- Opiates
- Cannabis
- Cocaine type
- Amphetamines
- Others
- No data

Crack	75.6
Inhalants	72.6
Heroin	69.4
Sedatives*	63.7
Hallucinogens	61.5
Stimulants*	59.1
Tranquilizers*	58.8
Cocaine (Not Including Crack)	57.5
Pain Relievers*	56.6
Marijuana	42.4
Alcohol	25.7

0%   20%   40%   60%   80%

Cannabis use
Life-time prevalence, Western Europe, %

14 (1995) 17 (1995) 22 (1995) 17 (1995)

Proportion of users in the world
Amongst people age 15-64 worldwide

4,343m
All people aged 15-64 in 2007

250m Number who have used drugs once in past year
38m Problem drug users
21m Number who inject drugs

Opium poppy cultivation
Hectares worldwide

200,000
150,000
100,000
50,000
0

Laos / Burma / Afghanistan

2004 2005 2006 2007 2008

Cocaine use
Annual prevalence in population aged 15-59, England, %

1.2 (1995) 2.0 2.4 2.6 2.3

Coca bush cultivation
Hectares worldwide

150,000
100,000
50,000
0

Bolivia / Peru / Colombia

2004 2005 2006 2007 2008

SOURCE: WORLD DRUG REPORT, 2009

COURRIER internacional

A CRISE DO ESTADO SOCIAL

O fim do Estado social

GDP

HAUSSE DU PRIX DU PÉTROLE...

EUROPE

CREDIT RATING

Courrier internacional

PESADELO

A SUSTENTABILIDADE do SISTEMA

Managing for Sustainability

Health & Safety, Equity, Training, IR
Our People
Environment
Waste, Water, Biodiversity
Resources
Maintenance, Assets, Design Guidelines
Community
Membership of Associations, Good Neighbours, Stakeholders
Financials
Expenditure & Income, Budgets

Managing for Sustainability

Impacts by brands / products   Influences on brands / products

Environmental
Consumers
Market forces
Key opinion formers
Social
Greenhouse

Sustainability, Energy & Environment
Frequently asked questions … and some answers

Sustainability
Social quality of life
Environmental, Natural Environment
Economic
Human Health

Building sustainability

Strategic sustainability
Helping top teams develop strategies built on sustainability principles

Practical sustainability
Helping to find and implement practical sustainability solutions throughout the organisation

Cultural sustainability
Helping to build the culture & capabilities to make sustainability stick

CFSP Possible workflow.ppt   3

Climate
• Emissions of greenhouse gases

Ozone layer
• Emissions that damage the ozone layer

Resources
• Consumption of fossil fuels
• Proportion of renewable energy sources
• Energy intensity

True costs
• Level to which operation costs are covered
• External costs of damage

Price
• Specific price of defined transport services

Safety
• Victims of transport accidents
• Crime
• Safe outdoor play
• Safe routes to school

Environment

Economy

Society

Habitats and landscape
• Unfragmented areas

Clean air
• Pollution in settlement areas $(O_3/NO_2/PM10)$
• NO_x emissions
• VOC emissions

Noise
• Pollution at place of residence
• Pollution of protected and recreational area

Living/land use
• Land used for transport
• Proportion of transport in the settlement area

Individuality
• Opportunity to choose mode of transport
• Degree of regulation

Participation
• Subjective satisfaction with opportunities to participate

Solidarity
• Public Service Obligations of transport per capita
• Accessibility of regional centre by public transport
• Accessibility of amenities for daily needs by public transport
• Accessibility of leisure facilities for young people
• Consideration of the needs of social groups

What
Sustainability Value Web

Quality of Life
Family & Community
Health
Education
Food Production
Standard of Living
Biodiversity
Atmosphere / Air
Water
Life Support

Your Organization

Economy & Infrastructure
Housing & Cities
Transportation
Energy
Resources & Trade
Population
Land

The Three Spheres of Sustainability

Social-Environmental
Environmental Justice
Natural Resources Stewardship
Locally & Globally

Environmental
Natural Resource Use
Environmental Managemet
Pollution Prevention
(air, water, land, waste)

Environmental-Economic
Energy Efficiency
Subsidies / Incentives for use of Natural Resources

Social
Standard of Living
Education
Community
Equal Opportunity

Sustainability

Economic
Profit
Cost Savings
Economic Growth
Research & Development

Economic-Social
Business Ethics
Fair Trade
Worker's Rights

Adopted from the 2002
University of Michigan
Sustainability Assessment

CONSTATAÇÕES da REALIDADE

HEALTH SYSTEMS FINANCING
The path to universal coverage

- " … existiu uma previsão supostamente realista segundo a qual no ano 2.000 nenhum país, ou sequer qualquer dos seus cidadãos, viesse a ter um nível de saúde inferior ao aceitável … a Saúde Pública é a arte de saber aplicar a ciência num determinado contexto político para reduzir as desigualdades na prestação dos cuidados de saúde enquanto se assegura simultaneamente a melhor saúde possível a um maior número de pessoas … a proporção do PNB alocado à Saúde deixou de crescer, e tem mesmo vindo a diminuir à escala planetária … constatando-se que do total do dispêndio financeiro em investigação de novos fármacos, somente 10% corresponde às doenças que são responsáveis por 90% da mortalidade em todo o Mundo … existindo cerca de 2 biliões de pessoas sem qualquer acesso aos medicamentos considerados essenciais … calcula-se ainda que o desperdício orce entre 20 a 40 % do total daquilo que se gasta com a Saúde … somente 20 % das pessoas têm uma cobertura social adequada, sendo calculado que para tal seriam apenas necessário cerca de 60 USD / pessoa / ano, em contraste com os actuais 20 USD, sendo certo que cerca de 50 % desta despesa é suportada directamente pelo próprio cidadão … " *(WHO Reports, 1998, 2004 e 2010)*

Fig. 1. **Three dimensions to consider when moving towards universal coverage**

Reduce cost sharing and fees

Include other services

Direct costs: proportion of the costs covered

Extend to non-covered

Current pooled funds

Services: which services are covered?

Population: who is covered?

The access framework

1 Rational use

3 Sustainable financing

ACCESS

2 Affordable prices

4 Reliable health and supply systems
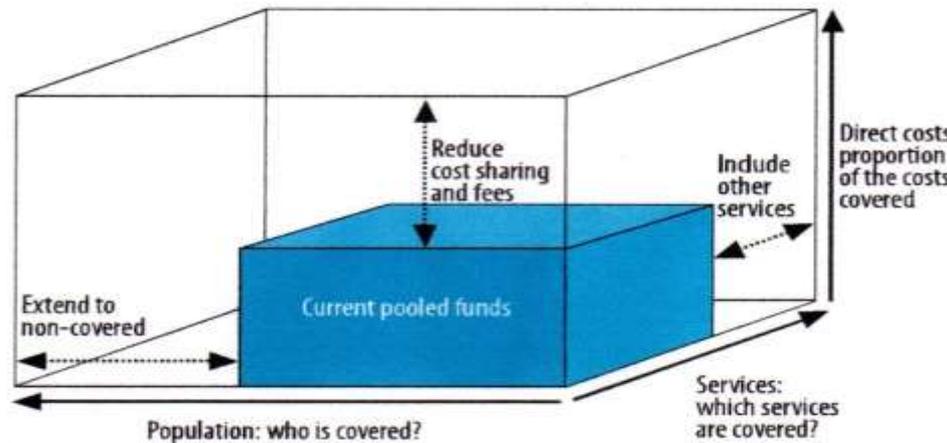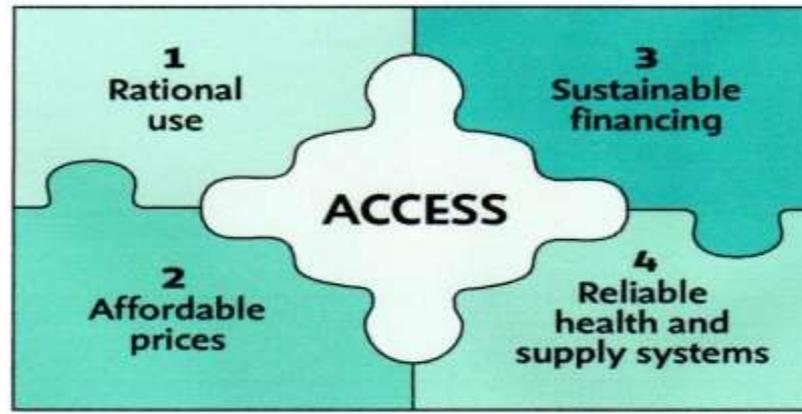
Source: WHO, 2000

Building of the global movement for health equity: from Santiago to Rio and beyond

Michael Marmot, Jessica Allen, Ruth Bell, Peter Goldblatt

Health inequalities are present throughout the world, both within and between countries. The Commission on Social Determinants of Health drew attention to dramatic social gradients in health within most countries and made proposals for action. These inequalities are not inevitable. The purpose of this article is to report on activity that has taken place worldwide after the report by the Commission on Social Determinants of Health. First, we summarise the global situation. Second, we summarise an interim report of the emerging findings from an independent review of social determinants and the health divide, which was commissioned by the WHO European region. The world conference on social determinants of health will be held in Rio de Janeiro, Brazil, in October, 2011. This summit provides an opportunity to galvanise support, prioritise action, and respond to the call by the Commission on Social Determinants of Health for social justice as a route to a fair distribution of health.

"Social Injustice is killing on a grand scale"

"A toxic combination...of poor social policies and programmes, unfair economic arrangements, and bad politics...is responsible for the fact that a majority of people in the world do not enjoy the good health that is biologically possible."

"When you cannot measure it, when you cannot express it in numbers, you have scarcely . . . advanced to the stage of Science, whatever the matter may be."

William Thomson, Lord Kelvin, 1824–1907

The Economic Burden of Malaria

John Luke Gallup* and Jeffrey D. Sachs.

Center for International Development, Harvard University, Cambridge, Massachusetts

KILLS 3000 CHILDREN EVERYDAY
IN AFRICA. PLEASE HELP.
HTTP://MALARIANOMORE.ORG

THE GLOBAL MALARIA BUSINESS PLAN (*Executive Summary*)

OF THE

ROLL BACK MALARIA PARTNERSHIP

MAY 5, 2008

Abstract

Malaria and poverty are intimately connected. Controlling for factors such as tropical location, colonial history, and geographical isolation, countries with intensive malaria had income levels in 1995 of only 33% that of countries without malaria, whether or not the countries were in Africa. The high levels of malaria in poor countries are not mainly a consequence of poverty. Malaria is geographically specific. The ecological conditions that support the more efficient malaria mosquito vectors primarily determine the distribution and intensity of the disease. Intensive efforts to eliminate malaria in the most severely affected tropical countries have been largely ineffective. Countries that have eliminated malaria in the past half century have all been either subtropical or islands. These countries' economic growth in the 5 years after eliminating malaria has usually been substantially higher than growth in the neighboring countries. Cross-country regressions for the 1965–1990 period confirm the relationship between malaria and economic growth. Taking into account initial poverty, economic policy, tropical location, and life expectancy, among other factors, countries with intensive malaria grew 1.3% less per person per year, and a 10% reduction in malaria was associated with 0.3% higher growth. Controlling for many other tropical diseases does not change the correlation of malaria with economic growth, and these diseases are not themselves significantly negatively correlated with economic growth. A second independent measure of malaria has a slightly higher correlation with economic growth in the 1980–1996 period. We speculate about the mechanisms that could cause malaria to have such a large impact on the economy, such as foreign investment and economic networks within the country.

PRELIMINARY DRAFT FOR DISCUSSION

Figure 1. Ten-year Comparison of Differences Between Actual and Expected Disease-Specific NIH Funding Relative to US Burden of Disease in DALYs. A comparison of differences between actual and expected funding values as predicted by DALYs burden alone in 1996 (light blue) and 2006 (navy). Negative values reflect actual funding dollars less than expected and positive values represent actual funding dollars more than expected.
doi:10.1371/journal.pone.0016837.g001
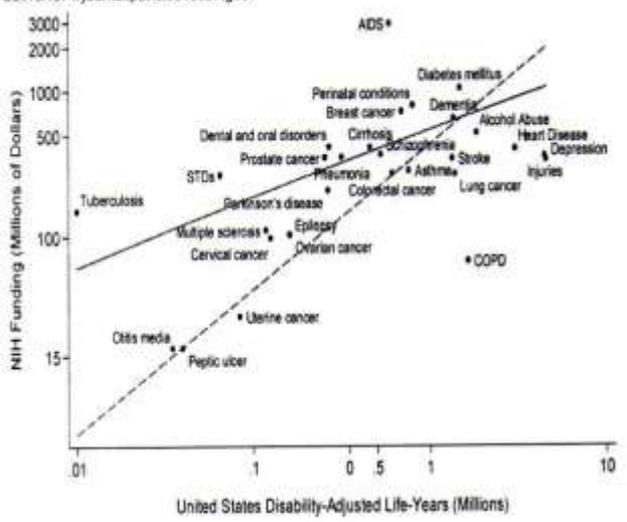
Figure 2. NIH Funding in 2006 and US Disease Burden in DALYs in 2004 for 29 Common Medical Conditions. The solid line represents the results of a traditional multivariable analysis, showing the relationship between US disease-specific DALYs burden and actual 2006 NIH funding dollars. The dashed line projects NIH funding levels in a similar multivariable model that requires that a disease with no burden receives no funding (constrained model). Though the models produce similar results, several diseases that would be considered overfunded in one model are considered underfunded in the other. For example, cervical cancer appears to be overfunded relative to the dashed line, while it is underfunded relative to the solid one.
doi:10.1371/journal.pone.0016837.g002

NIH Disease Funding Levels and Burden of Disease

Leslie A. Gillum[1], Christopher Gouveia[1], E. Ray Dorsey[3], Mark Pletcher[2], Colin D. Mathers[4], Charles E. McCulloch[2], S. Claiborne Johnston[1,2]*

1 Department of Neurology, University of California San Francisco, San Francisco, California, United States of America, 2 Department of Epidemiology and Biostatistics, University of California San Francisco, San Francisco, California, United States of America, 3 Department of Neurology, Johns Hopkins University Medical Center, Baltimore, Maryland, United States of America, 4 Department of Health Statistics and Informatics, World Health Organization, Geneva, Switzerland

Abstract

Background: An analysis of NIH funding in 1996 found that the strongest predictor of funding, disability-adjusted life-years (DALYs), explained only 39% of the variance in funding. In 1998, Congress requested that the Institute of Medicine (IOM) evaluate priority-setting criteria for NIH funding; the IOM recommended greater consideration of disease burden. We examined whether the association between current burden and funding has changed since that time.

Methods: We analyzed public data on 2006 NIH funding for 29 common conditions. Measures of US disease burden in 2004 were obtained from the World Health Organization's Global Burden of Disease study and national databases. We assessed the relationship between disease burden and NIH funding dollars in univariate and multivariable log-linear models that evaluated all measures of disease burden. Sensitivity analyses examined associations with future US burden, current and future measures of world disease burden, and a newly standardized NIH accounting method.

Results: In univariate and multivariable analyses, disease-specific NIH funding levels increased with burden of disease measured in DALYs (p = 0.001), which accounted for 33% of funding level variation. No other factor predicted funding in multivariable models. Conditions receiving the most funding greater than expected based on disease burden were AIDS ($2474 M), diabetes mellitus ($390 M), and perinatal conditions ($297 M). Depression ($719 M), injuries ($691 M), and chronic obstructive pulmonary disease ($613 M) were the most underfunded. Results were similar using estimates of future US burden, current and future world disease burden, and alternate NIH accounting methods.

Conclusions: Current levels of NIH disease-specific research funding correlate modestly with US disease burden, and correlation has not improved in the last decade.

Farmacoeconomia e TARV

Table 2. Survival Gains in Resulting From Medical/Surgical Advances in Various Diseases

Condition	Treatment	Per-person survival gains (months)
Non-small-cell lung cancer	Chemotherapy	7
Node and breast cancer	Adjuvant chemotherapy	29
Coronary artery disease	Bypass surgery	50
Relapsed non-Hodgkin's lymphoma	Marrow transplant	92
Prophylaxis in persons with HIV/AIDS	Opportunistic infection prophylaxis: trimethoprim/sulfamethoxazole, azithromycin	3
HIV/AIDS	Antiretroviral therapy	160

Cost-Effectiveness Ratios for HIV Care and Screening for Other Medical Conditions

Intervention/Population	Agent	Cost/QALY*	Reference
HIV			
PCP/toxo prophylaxis	TMP/SMX	$ 2800	4
Antiretroviral therapy	EFV/ZDV/3TC	$11,700	5
Resistance testing	————	$20,200	6
Inpatient screening	————	$38,600	2
MAC prophylaxis	Azithromycin	$43,300	4
HIV screen: high-risk patients every 5 yrs	————	$50,000	8
Other Conditions			
Breast cancer, women age > 50 yrs	Annual mammogram	$57,700	9
Colon cancer, age 50-85 yrs	Colonoscopy	$57,500	9
Diabetes, age > 25 yrs	FBS	$70,000	9

EXPERIMENTAL DRUGS ON TRIAL

A controversial lawsuit challenges the FDA's system of controlling access to experimental drugs and, some say, the scientific basis of drug approval

By Beryl Lieff Benderly
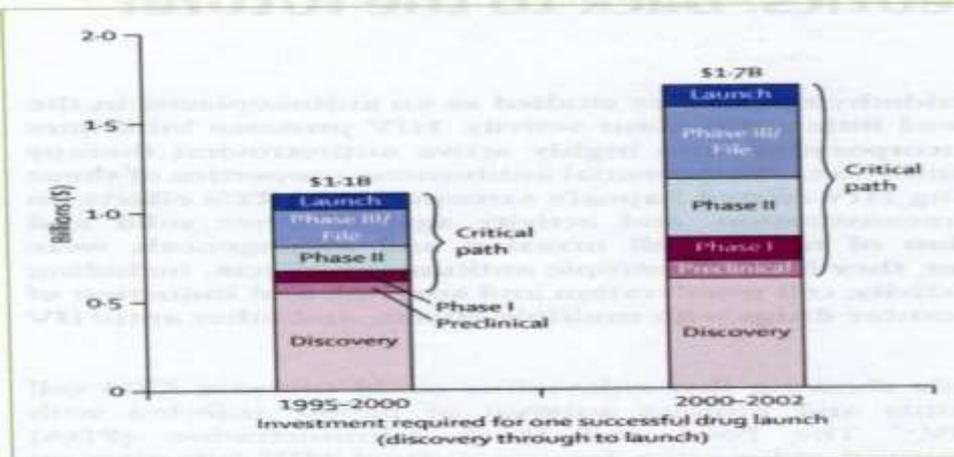
Figure 2: Growing cost of successful drug development
The cost of successful drug development increased 55% from 1995–2000 to 2000–02. Most of the rising costs are attributable to clinical trials. Reproduced with permission from reference 2.

RATIONAL USE OF MEDICINES

SUMMARY

- Worldwide, it is estimated that half of all medicines are inappropriately prescribed, dispensed or sold, and that half of all patients fail to take their medicine properly.

- An estimated two-thirds of global antibiotic sales occur without any prescription, and studies in Indonesia, Pakistan and India show that over 70% of patients were prescribed antibiotics. The great majority – up to 90% – of injections are estimated to be unnecessary.

- The inappropriate use of medicines is not only widespread, it is costly and extremely harmful both to the individual and the population as a whole. Adverse drug events rank among the top 10 causes of death in the USA and are estimated to cost that country between US$ 30 and US$ 130 billion each year.

- Growing resistance to antimicrobial medicines is a particularly serious challenge in countries at all economic levels, and results largely from inappropriate prescribing and use. For the treatment of malaria, chloroquine resistance is now established in 81 of the 92 countries in which the disease is endemic.

- Much greater use of evidence-based diagnostic and treatment guidelines by health professionals is needed.

- More effective monitoring and regulation of medicines, and public education and information are important components of a strategy for increased rational use.

- Por cada molécula aprovada para uso clínico, são investigadas 25 novos compostos, de entre aproximadamente 5.000 possíveis alternativas, durante cerca de 10 a 15 anos
- Somente 21,5 % das moléculas que entram em fase I de desenvolvimento chegam a ser comercializadas
- Custos triplicaram em 4 anos (1998-2001)
- Em 2007: 1,3 biliões de USD / molécula
- Em 2008: Investimento total (USA): 65.2 biliões de USD, contabilizando-se cerca de 2.900 compostos em investigação (300 para Doenças Raras, 750 para Doenças Neoplásicas, 277 para Doenças Cerebro/Cardio-Vasculares, e 109 para a SIDA)
- Em 2008: Existia um repositório de cerca de 100.000 moléculas que poderão ser futuramente investigadas, embora apenas 100 dos cerca de 3.000 alvos proteicos conhecidos a partir do Genoma Humano poderão permitir a síntese de novos medicamentos a curto/médio prazo
- Calcula-se que os medicamentos são responsáveis por cerca de 50 a 60% da redução da mortalidade presentemente verificada em todo o mundo, sendo mais acentuada nas Doenças Neoplásicas, Doenças Cerebro/Cardio-Vasculares e SIDA.

MALARIA IS BIG PROBLEM IN AFRICA

MNAZI MMOJA HOSPITAL ZANZIBAR TANZANIA E AFRICA
RECEPTION

FARMACIA
FARMACIA RYFER
DE DESCUENTO

CONSULTORIO MEDICO URGENCIAS 24 HORAS
Dr. José Guirao López

BEWARE OF MALARIA IT KILLS
BEWARE OF HIV/AIDS IT KILLS
MNAZI MMOJA HOSPITAL Z'BAR

Tooth Brush Tree.
salvadora persica
(swahili-mswaki).
The twigs are used as
a toothbrush. Roots
and leaves used for
treatment of malaria.

FEVER TREE
Acacia Xanthophlea (olerai-Masai)
The tree grows in groups in a low lying
swampy areas which is favourable breeding
ground for Malarial Mosquitoes. Early Pioneers
were convinced that the tree caused Malaria
which is not true. Roots used for
treatment of Stomach-ache.

DISCOUNT FARMACY
• GENERICS
• GERIATRICS
• ANTIBIOTICS

30%
GENERICS
GERIATRICS
ANTIBIOTICS

DOCTOR-TRADICIONAL
TRATA-SE TRADICIONAL-
MENTE E SPIRITUALMENTE
TLL 323815483
FEBRE AMARELA - MALUCU
CHUCHA GRANDE - DOR DE DENTE
MULHER OU HOMME NUNCA TEZ FILHO
CRIANÇA QUE TEM PEDRA NA BARRIGA
PALUDISMO - MAKULU
TROMBOSE - ANSIMA
PESSOAS QUE DESMAIA
VIA BERTA - PESSOA QUE AJUDA TODA
HORA - DOR DE BARRIGA
DOR DE BEXIGA - INFAMAÇÃO NA BEXIGA
XIGA - TUBERCULOSE - LUKUKA
IMORROIDE - APENDECITE
ASMA - DIABETA - ESCONDAMI
TO - MICROBIO - FORCA SEXUEL
MESTRUAÇÃO NÃO APARECE
QUE TEM QUALQUER PROBLEMA-
RESOLVE-SE - LAVAGEM DE BARRIGA
HELNA INTERNA E EXTERNA - GOTA
DOR DE COSTELA - DARMATISMO ANDA

Global Malaria Portfolio 2008

RESEARCH | | TRANSLATIONAL | | | DEVELOPMENT | |
Lead Generation | Lead Optimization | Preclinical | Phase I | Phase II | Phase III | Registration

Medicines for Malaria Venture

FIGURE 9-2 Drug development as an iterative process. The figure illustrates how biomarkers may be used to bridge, or translate, early preclinical findings to clinical findings, and how clinical findings may be used to inform and corroborate the basic science. Many of the workshop participants emphasized that the ultimate goal of applying these new technologies in safety science is to create a continual iterative process in which the basic scientific data can help to inform and predict clinical outcomes.
SOURCE: Frueh, 2007.

FIGURE 11: **The R&D Process: Long, Complex, and Costly**
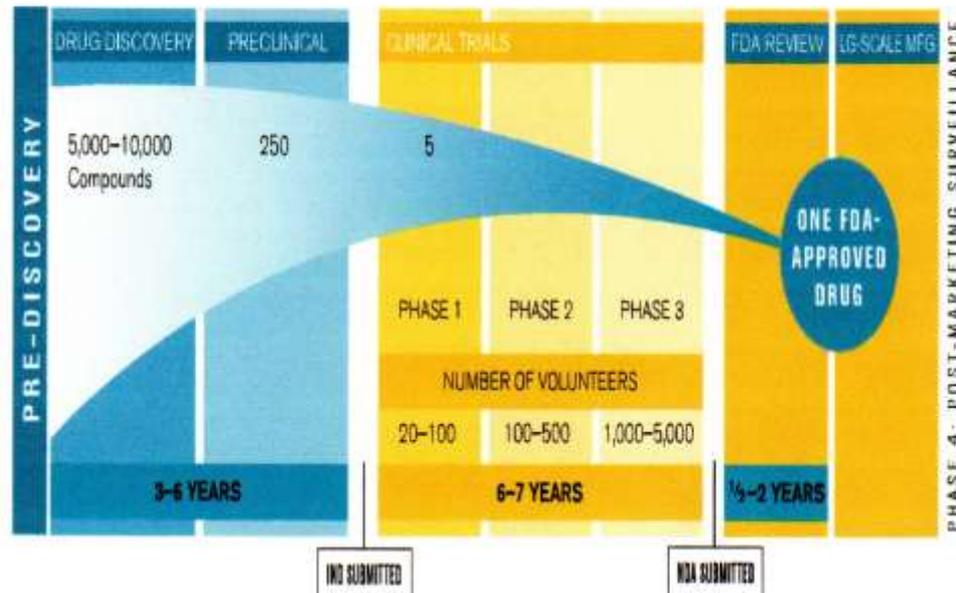
Figure 1: The Drug Discovery, Development, and Review Process

Source: Pharmaceutical Research and Manufacturers of America.
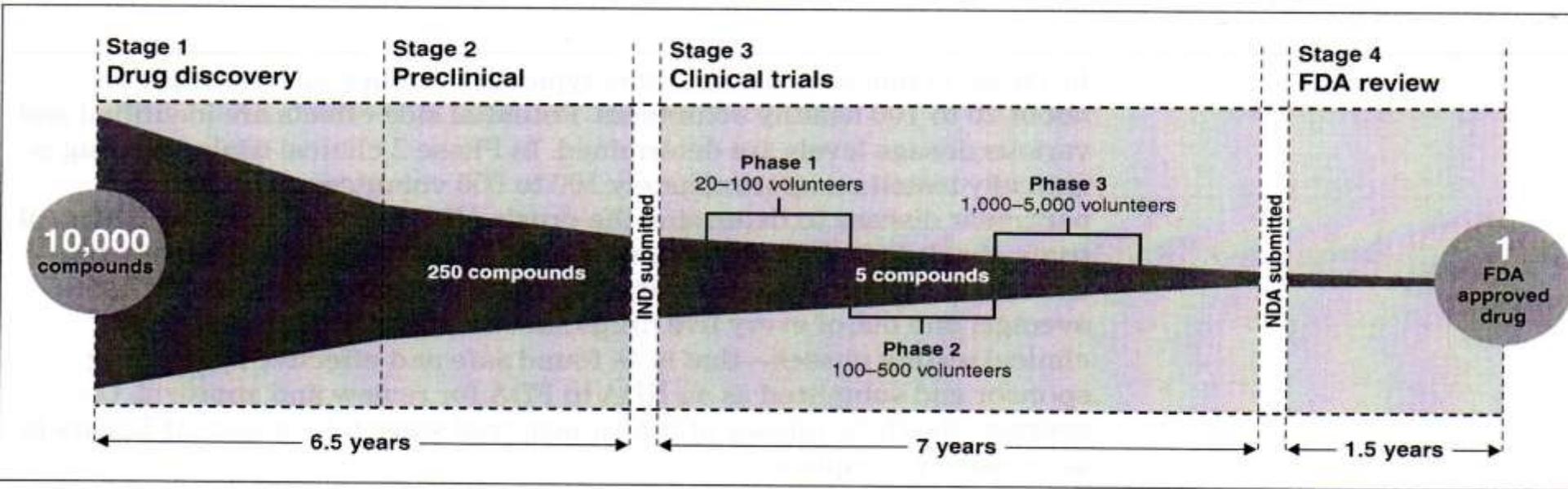
FIGURE 13: **Cost to Develop One New Drug**
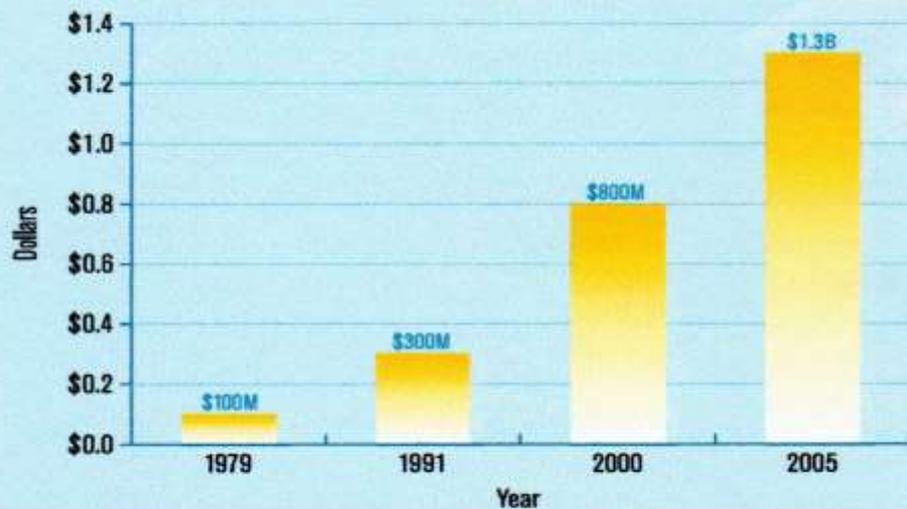
SOURCES: J. A. DiMasi and H. G. Grabowski, "The Cost of Biopharmaceutical R&D: Is Biotech Different?" *Managerial and Decision Economics* 28, no. 4–5 (2007): 469–479; J. A. DiMasi, et al., "The Price of Innovation: New Estimates of Drug Development Costs," *Journal of Health Economics* 22 (2003): 151–185.

FIGURE 9: **Biopharmaceutical Companies' Investment in R&D Remains Strong**
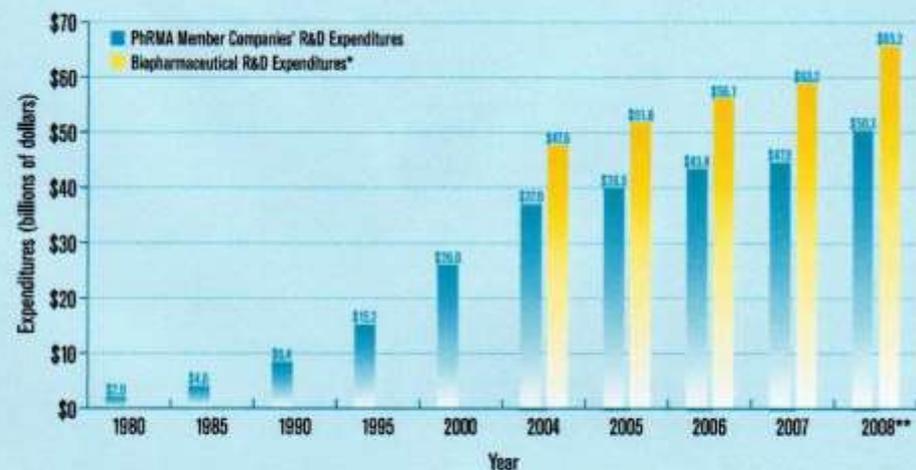
*The "Biopharmaceutical R&D Expenditures" figures include PhRMA research associates and nonmembers; these are not included in "PhRMA Member Companies' R&D Expenditures." PhRMA first reported this data in 2004.

**Estimated.

SOURCES: Burrill & Company, analysis for Pharmaceutical Research and Manufacturers of America, 2005–2009; Pharmaceutical Research and Manufacturers of America, *PhRMA Annual Member Survey* (Washington, DC: PhRMA, 1981–2008).

FIGURE 10: **Number of Compounds in Development, by Region***
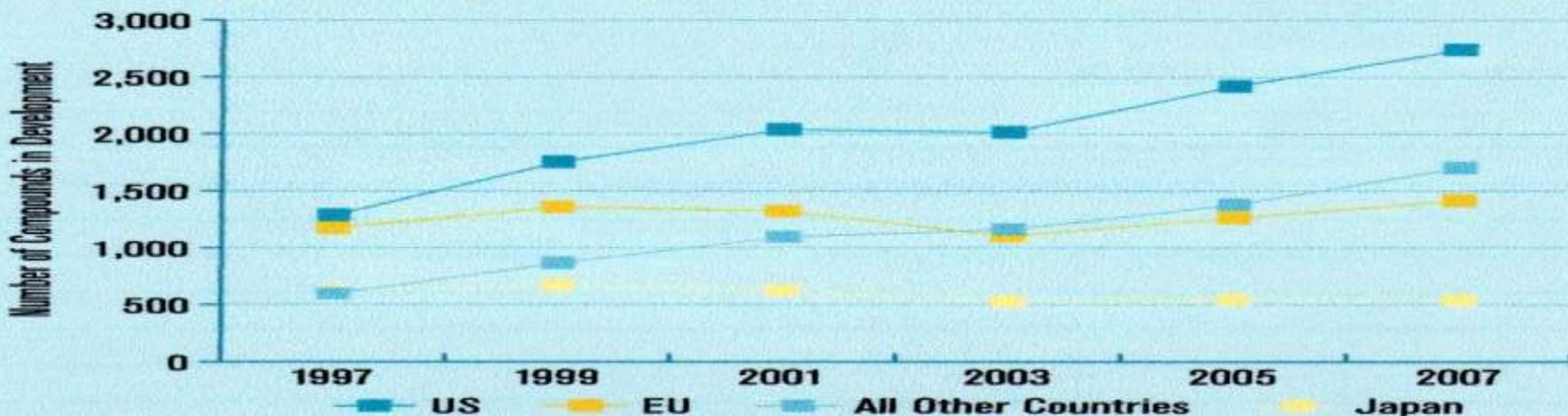
*Note: Reflects the number of compounds in clinical trials or awaiting approval as of June of each year. Compounds in development for multiple regions are counted in each region for which regulatory approval is sought, and multiple indications are counted only once.

SOURCE: Adis R&D Insight Database, Wolters Kluwer Health, Customized Run, December 2007.

Demythologizing the high costs of pharmaceutical research

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E-mail: rnwarbur@uvic.ca
*Corresponding author.

Abstract It is widely claimed that research to discover and develop new pharmaceuticals entails high costs and high risks. High research and development (R&D) costs influence many decisions and policy discussions about how to reduce global health disparities, how much companies can afford to discount prices for lower- and middle-income countries, and how to design innovative incentives to advance research on diseases of the poor. High estimated costs also affect strategies for getting new medicines to the world's poor, such as the advanced market commitment, which built high estimates into its inflated size and prices. This article takes apart the most detailed and authoritative study of R&D costs in order to show how high estimates have been constructed by industry-supported economists, and to show how much lower actual costs may be. Besides serving as an object lesson in the construction of 'facts', this analysis provides reason to believe that R&D costs need not be such an insuperable obstacle to the development of better medicines. The deeper problem is that current incentives reward companies to develop mainly new medicines of little advantage and compete for market share at high prices, rather than to develop clinically superior medicines with public funding so that prices could be much lower and risks to companies lower as well.
BioSocieties advance online publication, 7 February 2011; doi:10.1057/biosoc.2010.40

Table 3: Most new drugs and new indications for older drugs do not represent any significant therapeutic advantage – 1996–2006

Category	Number	Per cent
Major therapeutic innovation in an area where previously no treatment was available	2	0.2
Important therapeutic innovation but has limitations	38	3.9
Some value but does not fundamentally change the present therapeutic practice	106	10.8
Minimal additional value and should not change prescribing habits except in rare circumstances	251	25.5
May be new molecule but is superfluous because does not add to clinical possibilities offered by previously available products	442	45.0
Without evident benefit but with potential or real disadvantages	77	7.8
Decision postponed until better data and more thorough evaluation	67	6.8
Total	983	100.0

A look back at pharmaceuticals in 2006: Aggressive advertising cannot hide the absence of therapeutic advances (Prescrire International 2007; 16: 80–86).

Measuring Quality of Life in Health

ROD O'CONNOR

Quality of Life

Quality of Life Better or Worse?

BEFORE   AFTER

Michael Jackson
1958-2009

Liposuction
Medical
Anesthetic

wedge of skin below belly button to be removed

areas for lipo-suction

before   after

THE CURx̃e
THE END OF AIDS

CURE

THE CURx̃e
THERE'S A SHOT FOR THAT

AIDS

SPECIAL REPORT 35 PAGES THAT WILL HEAL THE WORLD
DISCOVER
HOW TO
CURE
EVERYTHING
SCIENCE DELIVERS SOLUTIONS TO LIFE'S MOST VEXING PROBLEMS
Obesity • Addiction • Diabetes • Heart Attacks • Cancer • Traffic • Unsolved Murders
Mosquitoes • Nuclear Fallout • Information Overload • The End of AIDS (At Last)
Biohackers: The Self-Made Experts Who Are Reinventing Medicine

Marlboro

THERE'S A SHOT FOR THAT

RARE = EQUAL

RARE DISEASE DAY — February 2011

Public health

Drug development for neglected diseases: a deficient market and a public-health policy failure

Patrice Trouiller, Piero Olliaro, Els Torreele, James Orbinski, Richard Laing, Nathan Ford

There is a lack of effective, safe, and affordable pharmaceuticals to control infectious diseases that cause high mortality and morbidity among poor people in the developing world. We analysed outcomes of pharmaceutical research and development over the past 25 years, and reviewed current public and private initiatives aimed at correcting the imbalance in research and development that leaves diseases that occur predominantly in the developing world largely unaddressed. We compiled data by searches of Medline and databases of the US Food and Drug Administration and the European Agency for the Evaluation of Medicinal Products, and reviewed current public and private initiatives through an analysis of recently published studies. We found that, of 1393 new chemical entities marketed between 1975 and 1999, only 16 were for tropical diseases and tuberculosis. There is a 13-fold greater chance of a drug being brought to market for central-nervous-system disorders or cancer than for a neglected disease. The pharmaceutical industry argues that research and development is too costly and risky to invest in low-return neglected diseases, and public and private initiatives have tried to overcome this market limitation through incentive packages and public-private partnerships. The lack of drug research and development for "non-profitable" infectious diseases will require new strategies. No sustainable solution will result for diseases that predominantly affect poor people in the South without the establishment of an international pharmaceutical policy for all neglected diseases. Private-sector research obligations should be explored, and a public-sector not-for-profit research and development capacity promoted.

Clinical research for rare disease: Opportunities, challenges, and solutions

Robert C. Griggs [a,*], Mark Batshaw [b], Mary Dunkle [c], Rashmi Gopal-Srivastava [d], Edward Kaye [e], Jeffrey Krischer [f], Tan Nguyen [g], Kathleen Paulus [f], Peter A. Merkel [h], For the rare diseases clinical research network

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[g] Office of In Vitro Diagnostics, Food and Drug Administration Parklawn Building, Room 6A55, HF-35, 5600 Fishers Lane, Rockville, MD 20857, USA
[h] Section of Rheumatology and the Clinical Epidemiology Unit, Vasculitis Center, E5, Boston University School of Medicine, 715 Albany Street, Boston, MA 02118, USA

ABSTRACT

Over 7000 rare diseases, each <200,000 US residents, affect nearly 30 million people in the United States. Furthermore, for the 10% of people with a rare disease and for their families, these disorders no longer seem rare. Molecular genetics have characterized the cause of many rare diseases and provide unprecedented opportunities for identifying patients, determining phenotypes, and devising treatments to prevent, stabilize, or improve each disease. Rare disease research poses challenges to investigators requiring specific approaches to: (1) the design of clinical studies; (2) the funding of research programs; (3) the discovery, testing, and approval of new treatments, and (4) the training of clinical scientists. Rigorous, statistically-valid, natural history-controlled, cross-over, and n-of-1 trials can establish efficacy and support regulatory approval of new treatments for rare diseases. The U.S. Orphan Drug Act of the U.S. FDA has stimulated industry investment in clinical trials to develop treatments for rare diseases. For trainees interested in finding a treatment for a rare disease, a commitment to longitudinal care of patients provides a base for the characterization of phenotype and natural history, a stimulus for innovation, a target population for research and helps fund training and research. The scientific methodology, financial resources, and logistics of clinical research for rare diseases have changed dramatically in the past two decades resulting in increased understanding of the pathophysiology of these disorders and direct benefit to patients.

HAVE NO FEAR

REVIEW ARTICLE

CURRENT CONCEPTS

Control of Neglected Tropical Diseases

Peter J. Hotez, M.D., Ph.D., David H. Molyneux, Ph.D., D.Sc.,
Alan Fenwick, Ph.D., Jacob Kumaresan, M.B., B.S., Dr.P.H.,
Sonia Ehrlich Sachs, M.D., Jeffrey D. Sachs, Ph.D., and Lorenzo Savioli, M.D.

Table 1. The Major Neglected Tropical Diseases Ranked by Prevalence.

Disease	Global Prevalence (millions)	Population at Risk	Regions of Highest Prevalence	Source
Ascariasis	807	4.2 billion	East Asia and Pacific Islands, sub-Saharan Africa, India, South Asia, China, Latin America and Caribbean	Bethony et al., de Silva et al.
Trichuriasis	604	3.2 billion	Sub-Saharan Africa, East Asia and Pacific Islands, Latin America and Caribbean, India, South Asia	Bethony et al., de Silva et al.
Hookworm infection	576	3.2 billion	Sub-Saharan Africa, East Asia and Pacific Islands, India, South Asia, Latin America and Caribbean	Bethony et al., de Silva et al.
Schistosomiasis	207	779 million	Sub-Saharan Africa, Latin America and Caribbean	Steinmann et al.
Lymphatic filariasis	120	1.3 billion	India, South Asia, East Asia and Pacific Islands, sub-Saharan Africa	Ottesen, WHO
Trachoma	84	590 million	Sub-Saharan Africa, Middle East and North Africa	International Trachoma Initiative, Médecins sans Frontières
Onchocerciasis	37	90 million	Sub-Saharan Africa, Latin America and Caribbean	Basáñez et al.
Leishmaniasis	12	350 million	India, South Asia, sub-Saharan Africa, Latin America and Caribbean	Desjeux
Chagas' disease	8–9	25 million	Latin America and Caribbean	WHO
Leprosy	0.4	ND	India, sub-Saharan Africa, Latin America and Caribbean	International Federation of Anti-Leprosy Associations
Human African trypanosomiasis	0.3	60 million	Sub-Saharan Africa	Fèvre et al.
Dracunculiasis	0.01	ND	Sub-Saharan Africa	Carter Center
Buruli ulcer	ND	ND	Sub-Saharan Africa	Global Buruli Ulcer Initiative

* ND denotes not determined.

Integrating Neglected Tropical Diseases into AIDS, Tuberculosis, and Malaria Control

Peter J. Hotez, M.D., Ph.D., Neeraj Mistry, M.D., M.P.H., Joanna Rubinstein, D.D.S., Ph.D., and Jeffrey D. Sachs, Ph.D.

| Cause	Disability-Adjusted Life-Years (millions)
Lower respiratory tract infections	91.4
HIV–AIDS	84.5
Unipolar depression	67.3
Diarrheal diseases	62.0
Ischemic heart disease	58.6
Neglected tropical diseases	56.6
Cerebrovascular diseases	49.2
Malaria	46.5
Road traffic accidents	38.7
Tuberculosis	34.7

Figure 1. The 10 Leading Causes of Life-Years Lost to Disability and Premature Death.

The number of years lost to disability and premature death (disability-adjusted life-years) for the 13 major neglected tropical diseases were calculated according to a method we described previously.[4] The disability-adjusted life-years for the other conditions are based on data from the World Health Organization.[23] The ranking of disease burdens is based on data in Hotez.[5]

NEGLECTED DISEASES: HAVE OUR GOVERNMENTS GOT SLEEPING SICKNESS?

DNDi
Drugs for Neglected Diseases initiative

U.S. Agency for International Development control programs for neglected tropical diseases

Geneva Global grant countries

Other countries with five or more neglected tropical diseases

Neglected Tropical Diseases
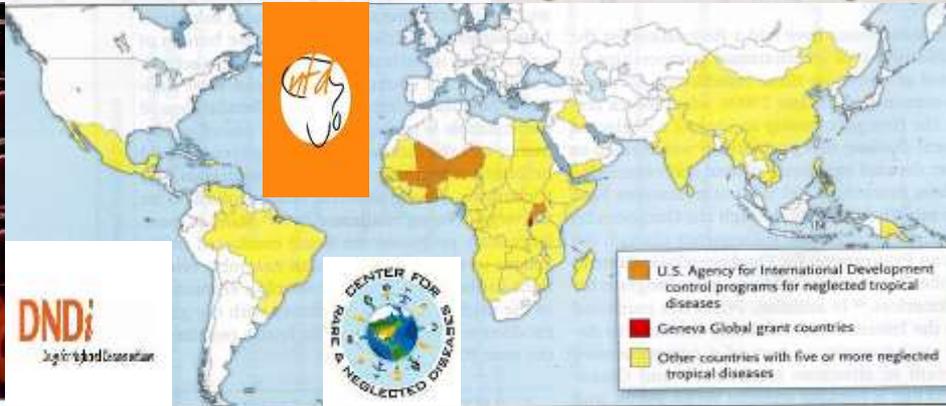
Figure 2. Nations with Five, Six, or Seven Neglected Tropical Diseases to Be Targeted for Integrated Preventive Chemotherapy.

Of the 56 nations to be targeted with a rapid-impact package, shown in yellow, 37 are located in the World Health Organization (WHO) African region, 5 in the WHO Region of the Americas, 5 in the WHO Eastern Mediterranean region, 3 in the WHO South-East Asia region, and 6 in the WHO Western Pacific region. Data regarding the occurrence of lymphatic filariasis, onchocerciasis, schistosomiasis, and the three soil-transmitted helminth infections are derived from the WHO.[44] Data regarding the occurrence of trachoma are derived from the WHO.[45] The five nations shown in orange — Burkina Faso, Ghana, Mali, Niger, and Uganda — will be targeted for integrated control in national programs through the support of the U.S. Agency for International Development Neglected Tropical Disease Control Program beginning this year. The two nations shown in red — Rwanda and Burundi — will be targeted for integrated control in national programs through the support of Geneva Global beginning this year.

**Neglected Diseases in the Developing World**
PROGRESS, CURRENT CHALLENGES, AND PROMISING APPROACHES

Learn More

POLICY SUMMARY I
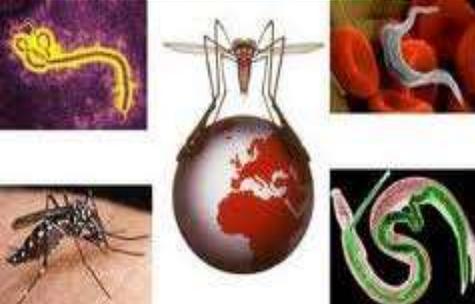
Addressing financial sustainability in health systems

Sarah Thomson, Tom Foubister, Josep Figueras, Joseph Kutzin, Govin Permanand, Lucie Bryndová

Observatory — HEN
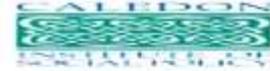
Population Aging, Health Care Spending and Sustainability: Do we really have a crisis?

by

Joe Ruggeri

September 2002
ISBN: 1-55382-027-4

⦿ " … a inovação tecnológica é o factor mais importante de incremento dos custos em saúde …o envelhecimento "apenas" contribui com cerca de 10% …"
(EU Report, 2009)

⦿ " … no debate acerca da sustentabilidade financeira relativa à prestação dos cuidados de saúde, talvez seja importante levantar a seguinte questão: Como é que ninguém questiona o facto de se gastar praticamente 12 % do PIB em actividades recreativas, mas simultaneamente toda a gente reclamar que se torna incomportável despender cerca de 10 % em saúde? … "
**(Canada Report, 2002)**

The Road to Sustainability

Integrating Sustainability

think sustainability!

World Population Ageing 2009

United Nations

Figure 44. Coverage as measured by active members of mandatory pension systems as a share of labour force: world 2000-2008

Contributors as % of Labour Force
- 0 to 5
- 5 to 25
- 25 to 50
- 50 to 100
- No data

Source: Hinz, Pallares-Miralles and Romero (forthcoming, 2010).

Figure old38. Distribution of economically active population aged 65 or over by sex: world, 1980-2020

Figure 17. Potential support ratio: world and development regions, 1950-2050
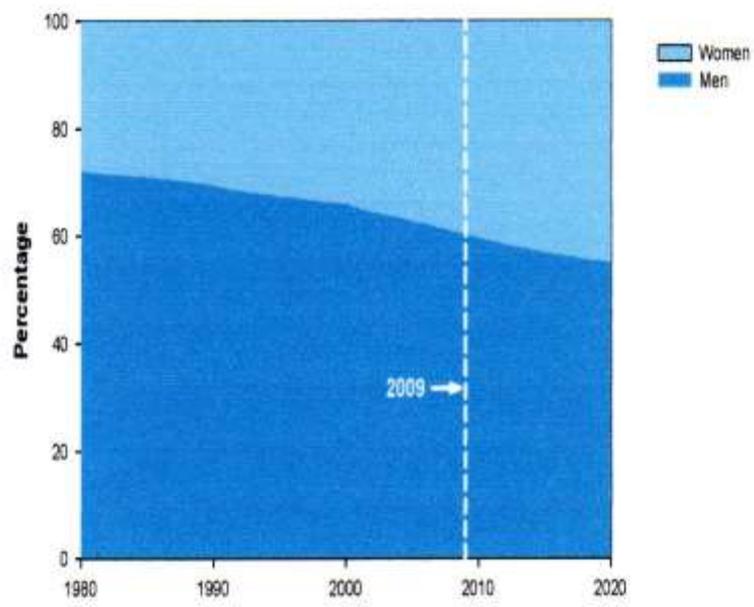
Os PRINCIPAIS PROBLEMAS ACTUAIS

**Envelhecimento
Doenças Crónicas
Deficiências
Dependências
As Falsas Profecias**

The World Health Report 1998

Life in the 21st century
**A vision for all**

Report of the Director-General

World Health Organization
Geneva
1998

An Aging World: 2008
International Population Reports

Issued June 2009
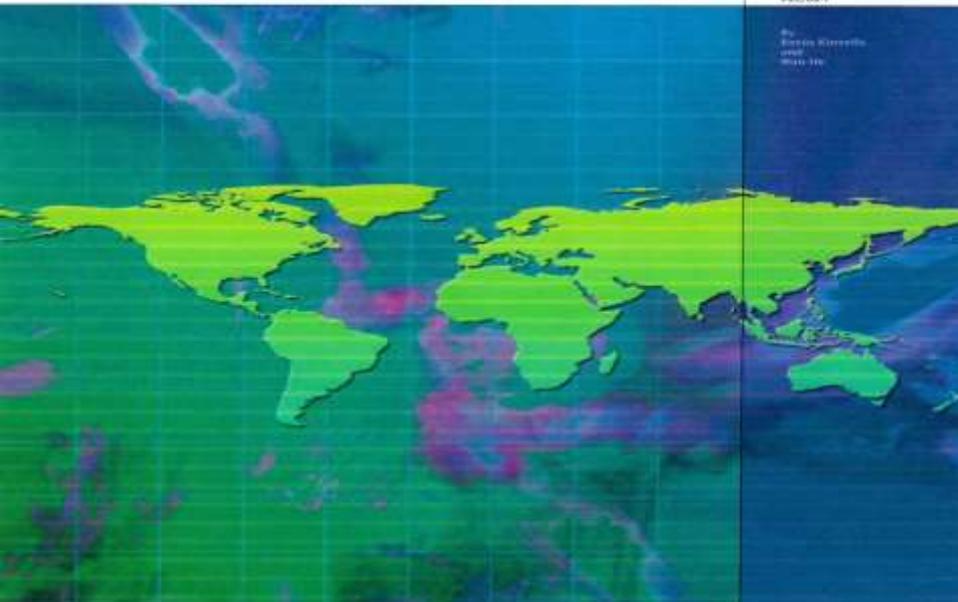
World Health Organization

U.S. Department of Health & Human Services

⊙ " … nós estamos presentemente a aprender lentamente uma das lições mais importantes: não só como viver mais anos, mas também como ficar mais tempo com um bom estado de saúde e com menor dependência dos outros … o padrão de progresso mais importante que emerge na actualidade, é o da maior sobrevivência com qualidade de vida … " *(sic.)* *(Hiroshi Nakajima, Secretário Geral da OMS, 2008)*

Directorate-General for Health & Consumers

Major and Chronic Diseases
REPORT 2007

1950-1955 estimate

1990-1995 estimate

0.8 1.5 2 2.5 3 3.5 4 4.5 5 5.5 6 6.5 7 7.5 8 8.5 9.25

0.8 1.5 2 2.5 3 3.5 4 4.5 5 5.5 6 6.5 7 7.5 8 8.5 9.25
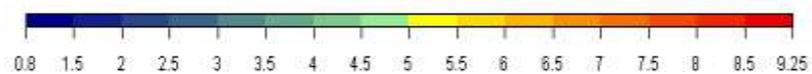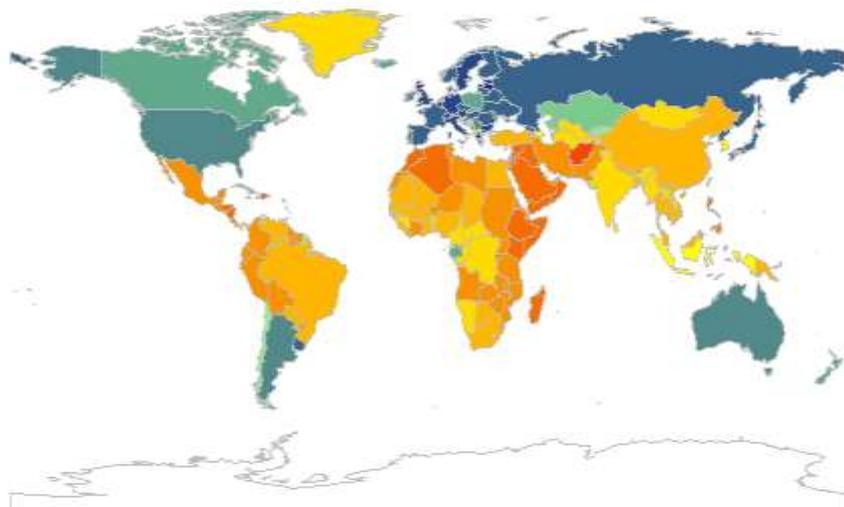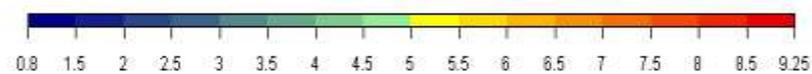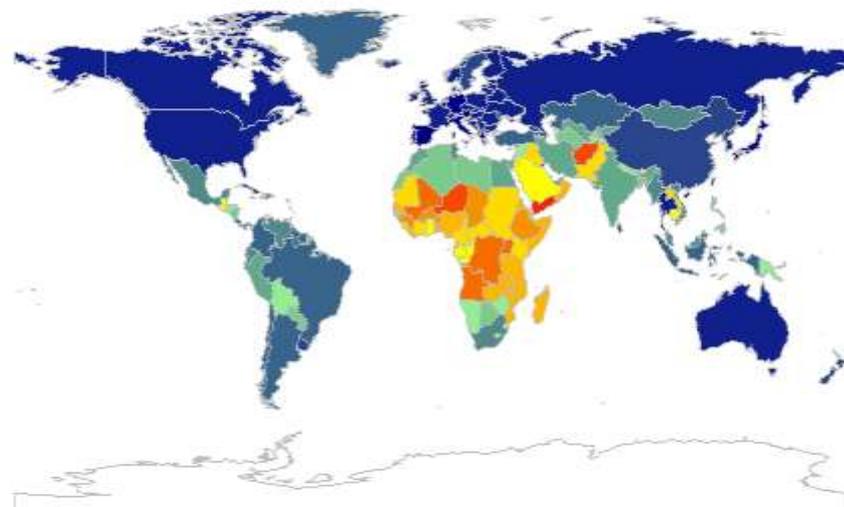
2015-2020 median projection
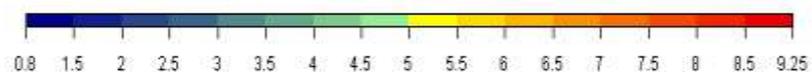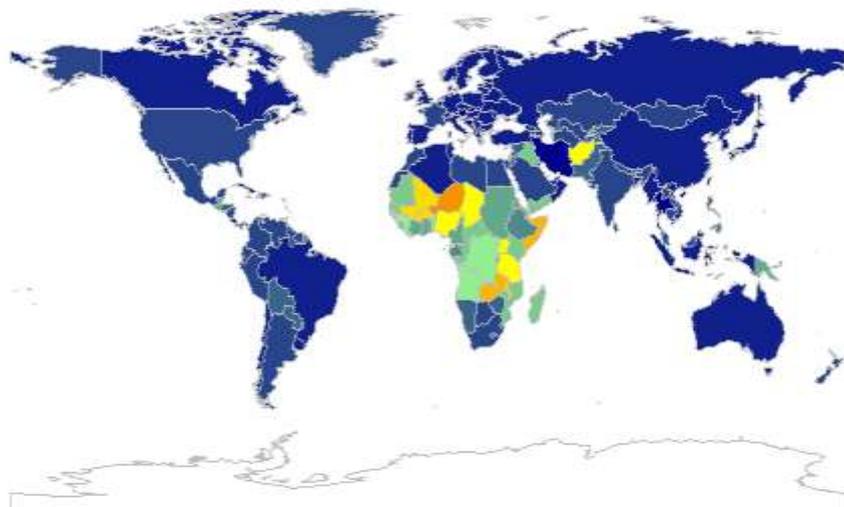
2095-2100 median projection

0.8 1.5 2 2.5 3 3.5 4 4.5 5 5.5 6 6.5 7 7.5 8 8.5 9.25

0.8 1.5 2 2.5 3 3.5 4 4.5 5 5.5 6 6.5 7 7.5 8 8.5 9.25

The World Health Report 1998
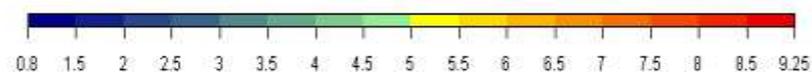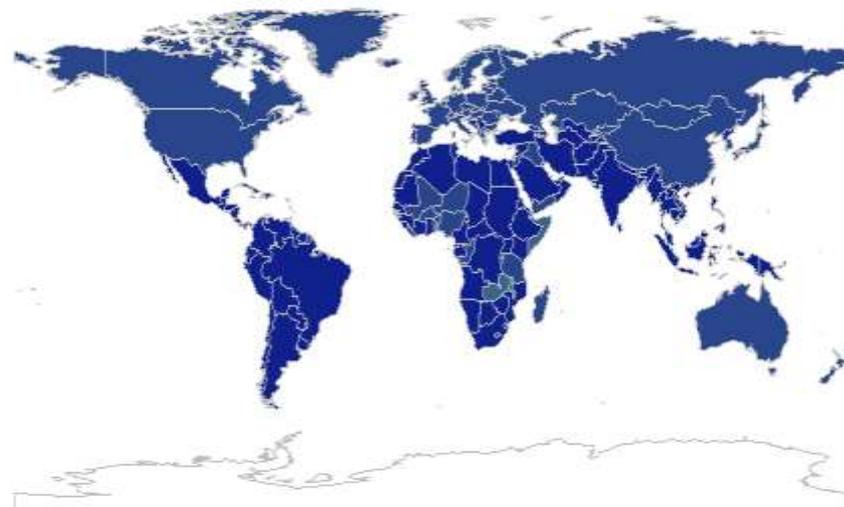
Fig. 13. Population by age and sex, 1995 and 2025

World

□ 2025
■ 1995

Age: 80+, 75-79, 70-74, 65-69, 60-64, 55-59, 50-54, 45-49, 40-44, 35-39, 30-34, 25-29, 20-24, 15-19, 10-14, 5-9, 0-4

Developed market economies

Developing countries (excluding LDCs)

Least developed countries

Males | **Females**

Percentage of total population by level of development
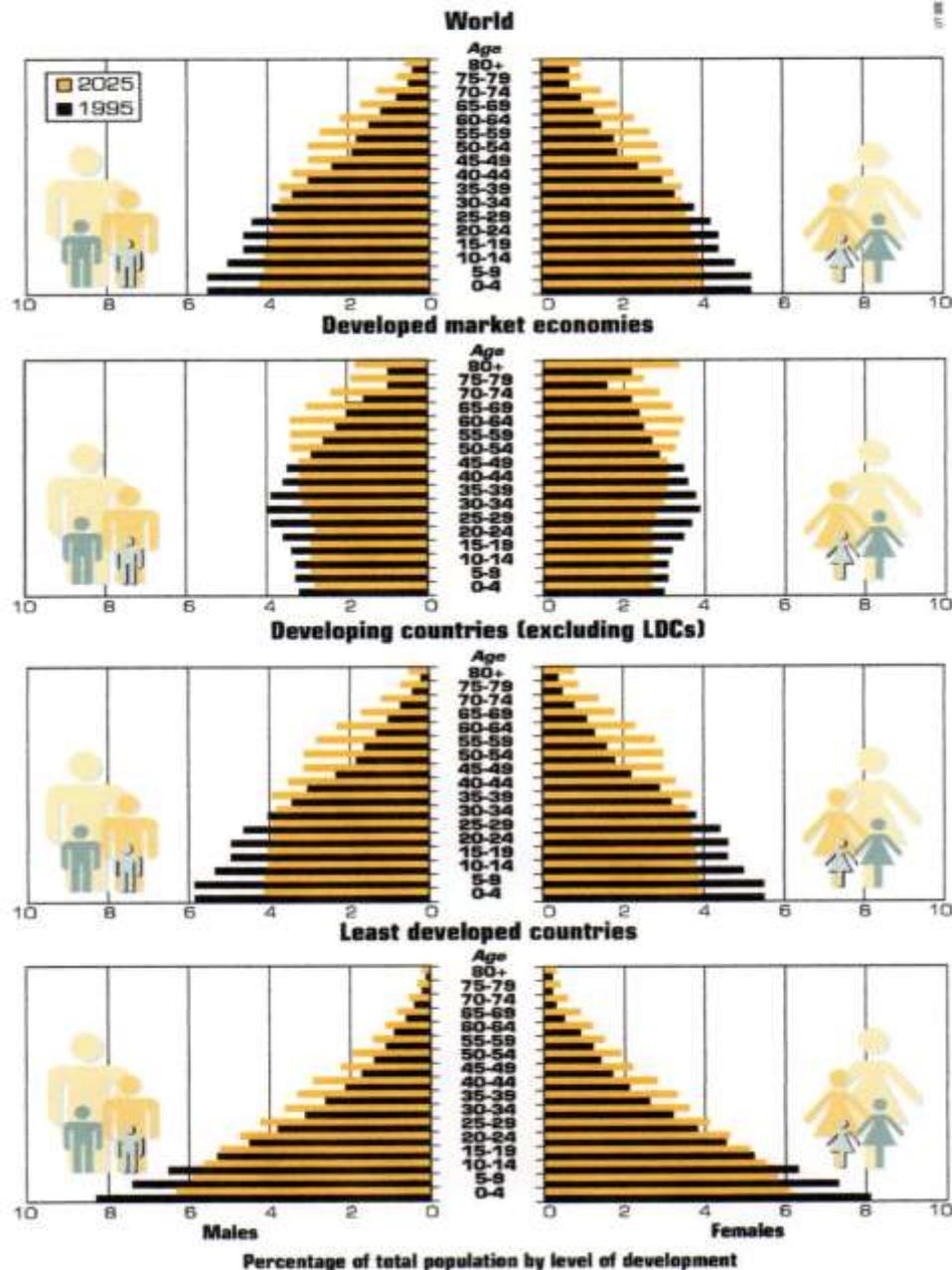
Population Age Shifts
Reshape Global Economics and Geopolitics

Adele Hayutin, Ph.D.
Director, Global Aging Program
Stanford Center on Longevity
ahayutin@stanford.edu

Populations Shift to Older People: Pyramid to Cube

As fertility falls and longevity increases, a country's age profile changes from a pyramid with a broad base of young people to a cube with a more even distribution across age brackets. If fertility remains low, the population will become top-heavy with older people.

The share of older people (65+) is increasing almost everywhere, but at different rates depending on changes in fertility and longevity. These population age shifts have profound impacts on everything from a country's workforce growth and economic prospects to public and personal budgets, security risks, cultural institutions and family structures.

Oldest Countries: Working-age populations shrink; age structures become top-heavy with old people.

- Fertility has been below replacement level of 2.1 births per woman for the last 10-15 years.
- Working-age population (15-64) is shrinking.
- Declining workforce growth combined with the increasing fiscal burden of retirees threatens economic growth and current living standards.
- Selected countries: Germany, Italy, Japan, Singapore, South Korea, Spain.

Japan

Median age 22 — 1950 — 84 million
43 — 2005 — 128 million
55 — 2050 — 103 million

65+, 15-64, 0-14
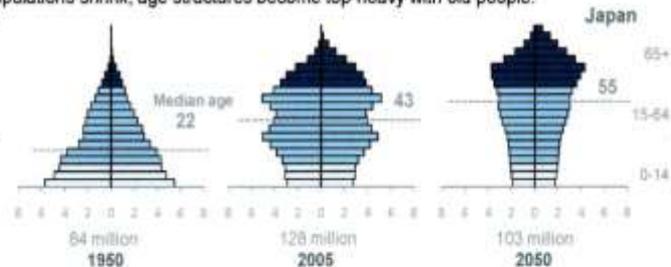
Middle-Age Countries: Workforce growth continues but at a slower pace; age structures become cube-like.

- Fertility rates are approaching or already below replacement rate.
- Workforce growth will be slow or moderate, eventually becoming negative in some countries.
- Sustained economic growth will require productivity gains to offset slower workforce growth.
- Selected countries: Australia, Brazil, Canada, China, Ireland, Mexico, Thailand, Tunisia, United States, Vietnam.

US

Median age 30 — 1950 — 158 million
36 — 2005 — 300 million
41 — 2050 — 402 million

65+, 15-64, 0-14
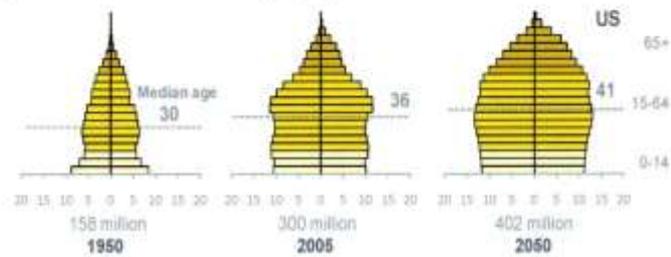
Young Countries: Rapid growth in working-age population threatens economic and political stability.

- Fertility rates are falling but still relatively high; age structure remains pyramidal.
- Working-age population will continue to grow rapidly, though the pace will slow.
- "Youth bulges" increase the risk of violent conflict; the challenge is to provide work for a large share of young adults.
- Selected countries: Afghanistan, Bolivia, Nigeria, India, Iraq, Pakistan, Philippines, Uganda.

Pakistan

Median age 21 — 1950 — 37 million
20 — 2005 — 158 million
34 — 2050 — 292 million

65+, 15-64, 0-14
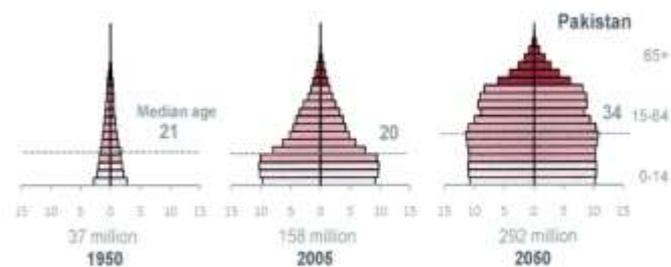
Population in millions by five-year age bracket. Males on left, females on right.

Global Aging
Share of Population Age 65 and Older

**Dramatic and unprecedented population age shifts are occurring at vastly different rates around the world.**

% 65+, 1950–2050

The worldwide share of population age 65 and older is projected to more than double to over 16% by 2050.

World
40%
30%
20%
10%
0%
1950 2000 2050

Map chart labels: Canada, US, Mexico, Colombia, Brazil, Argentina, Chile, UK, Sweden, Poland, Russia, South Korea, EU 27, Germany, Italy, Turkey, Iran, China, Japan, Ethiopia, Saudi Arabia, Pakistan, India, Vietnam, Philippines, Nigeria, Dem. Rep. of Congo, South Africa, Singapore, Indonesia, Australia

15 Oldest Countries, 2030

Europe and Japan are currently the oldest; by 2030 several Asian Tigers join the ranks of the oldest.

Country	2005	2030	2050
1. Japan	19.7 %	30.6 %	37.7 %
2. Singapore	8.5 %	27.4 %	32.6 %
3. Germany	18.8 %	27.3 %	30.2 %
4. Italy	19.7 %	27.0 %	32.6 %
5. Hong Kong	12.0 %	25.8 %	32.6 %
6. Slovenia	15.6 %	25.7 %	33.1 %
7. Finland	15.9 %	25.0 %	25.6 %
8. Austria	16.2 %	24.6 %	29.0 %
9. Belgium	17.3 %	24.4 %	27.1 %
10. Greece	18.3 %	24.2 %	31.7 %
11. Netherlands	14.2 %	24.1 %	25.2 %
12. Croatia	17.2 %	24.1 %	28.5 %
13. Spain	16.8 %	23.9 %	33.2 %
14. Switzerland	15.4 %	23.9 %	25.0 %
15. South Korea	9.4 %	23.4 %	35.1 %
US	12.3 %	19.4 %	21.0 %
EU 27	16.7 %	23.9 %	28.7 %
World	7.3 %	11.7 %	16.2 %

Declining Fertility Rates

Fertility has decreased worldwide, but at different rates.

(Chart: lines for Nigeria, Pakistan, US, China, Japan, Italy; y-axis 0–8, x-axis 1950–2050)
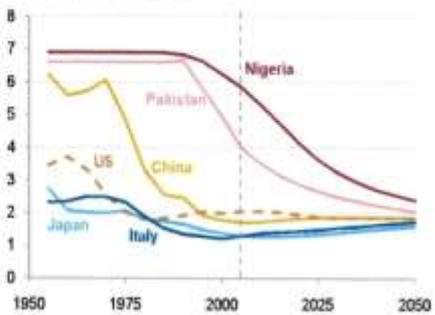
Rising Median Age

The pace of aging varies dramatically; many countries face steep gains in median age.

(Chart: lines for Japan, Italy, China, US, Pakistan, Nigeria; y-axis 15–55, x-axis 1950–2050)
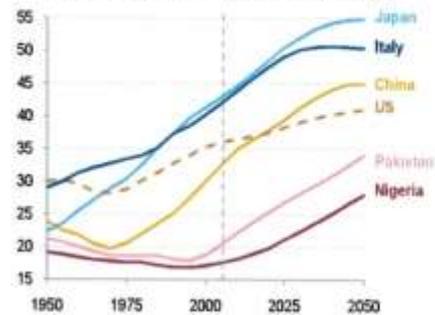
Declining Worker to Retiree Ratios

As countries age, the ratio of potential workers (15–64) to retirees (65+) declines, increasing the fiscal burden on workers.

(Chart: lines for Nigeria, Pakistan, Japan, US, China, Italy; y-axis 0–20, x-axis 1950–2050)

STANFORD CENTER ON LONGEVITY
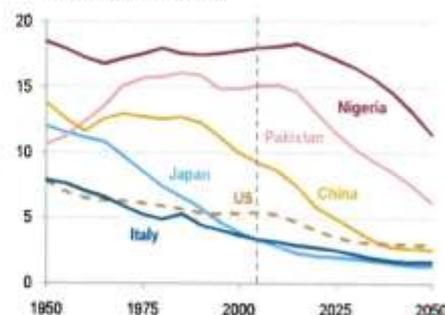
Figure 4-2.
Life Expectancy at Birth: 2008

Life expectancy in years
Less than 64
64 to 72
73 to 77
78 or more
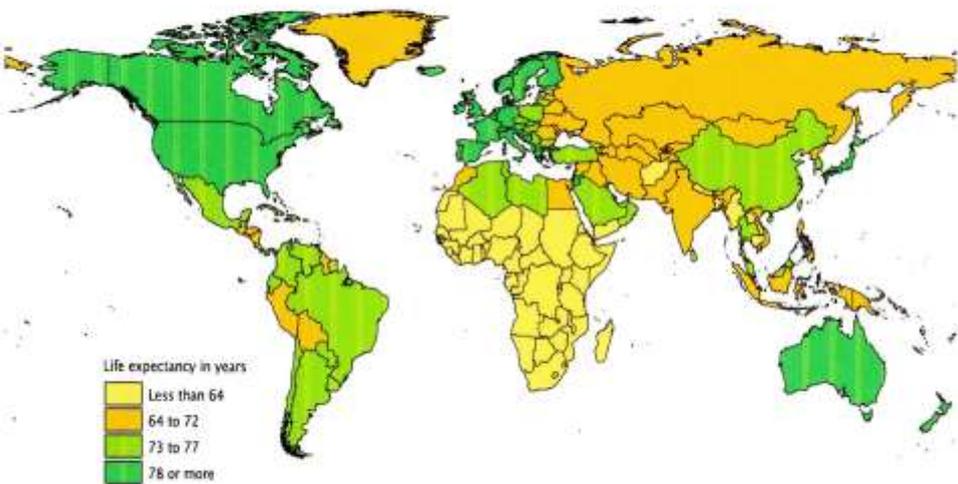
Figure 2-1.
Young Children and Older People as a Percentage of Global Population: 1950 to 2050

Percent

Under 5

65 and over
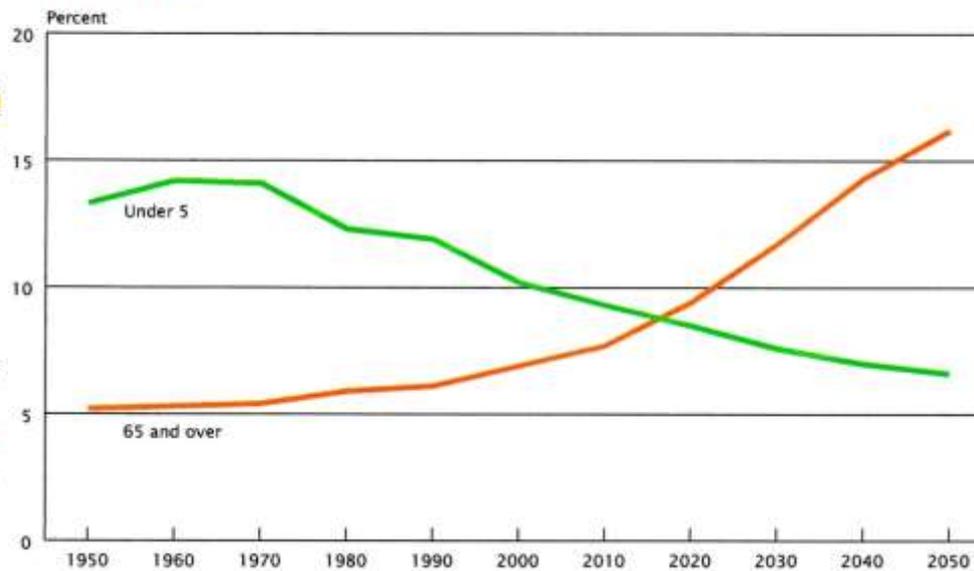
Figure 1. Total fertility rate and life expectancy at birth: world, 1950-2050

Total fertility
Life expectancy

Figure 13. Median age of the population: world and development regions, 1950-2050

(years)

World
More developed regions
Less developed regions
Least developed countries

Figure 20. Distribution of population aged 60 or over by age: world, 1950-2050

80+
70-79
60-69
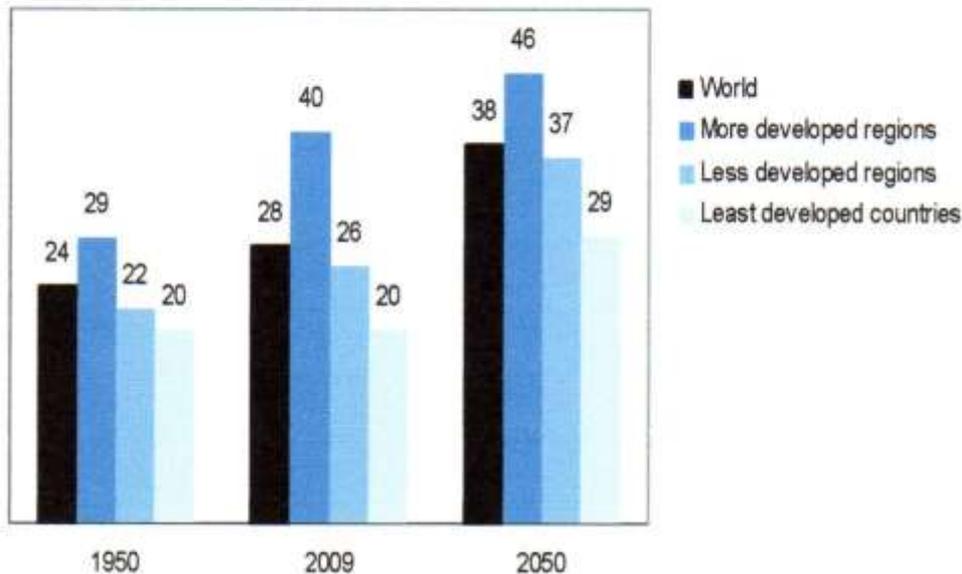
Figure 23. Distribution of world centenarians by development regions, 2009-2050

Less developed regions
More developed regions
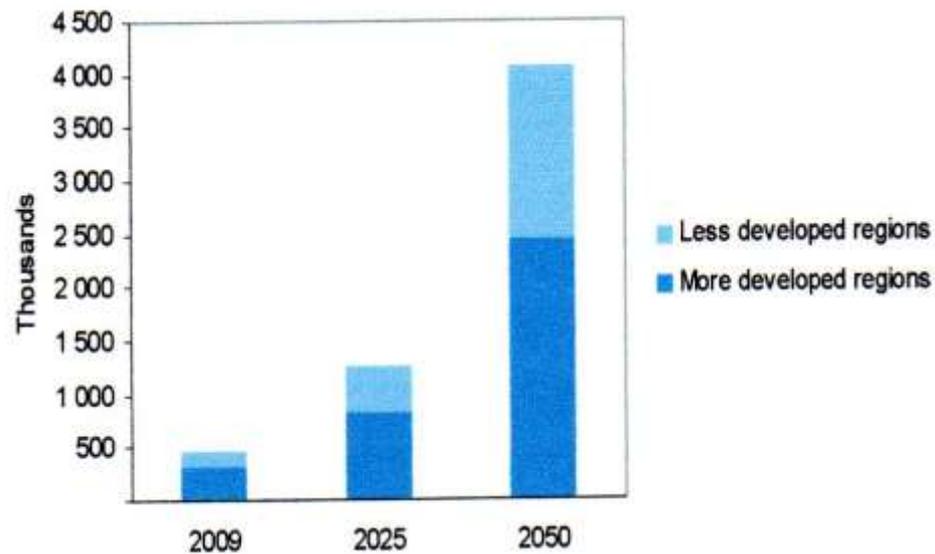
Preventing CHRONIC DISEASES
a vital investment

World Health Organization

## CHRONIC DISEASES ARE THE MAJOR CAUSE OF DEATH IN ALMOST ALL COUNTRIES

Chronic diseases include heart disease, stroke, cancer, chronic respiratory diseases and diabetes. Visual impairment and blindness, hearing impairment and deafness, oral diseases and genetic disorders are other chronic conditions that account for a substantial portion of the global burden of disease.

From a projected total of 58 million deaths from all causes in 2005, it is estimated that chronic diseases will account for 35 million, which is double the number of deaths from all infectious diseases (including HIV/AIDS, tuberculosis and malaria), maternal and perinatal conditions, and nutritional deficiencies combined.

Projected global deaths by cause, all ages, 2005

- 2 830 000 deaths — HIV/AIDS
- 1 607 000 deaths — Tuberculosis
- 883 000 deaths — Malaria
- 17 528 000 deaths — Cardiovascular diseases
- 7 586 000 deaths — Cancer
- 4 057 000 deaths — Chronic respiratory diseases
- 1 125 000 deaths — Diabetes

## 35 000 000 people will die from chronic diseases in 2005
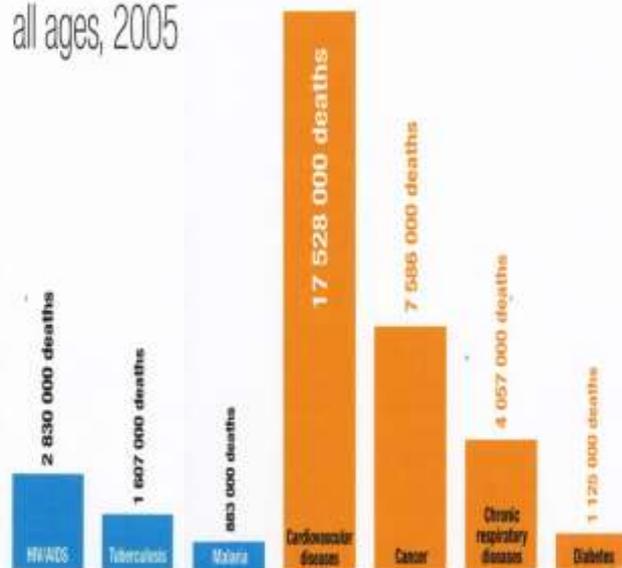
60% of all deaths are due to chronic diseases

THE POOREST COUNTRIES ARE THE WORST AFFECTED

Only 20% of chronic disease deaths occur in high income countries – while 80% occur in low and middle income countries, where most of the world's population lives. As this report will show, even least developed countries such as the United Republic of Tanzania are not immune to the growing problem.

Projected deaths by major cause and World Bank income group, all ages, 2005
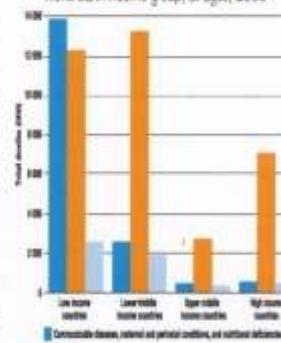
80% of chronic disease deaths occur in low and middle income countries

Projected foregone national income due to heart disease, stroke and diabetes in selected countries, 2005–2015

Brazil, Canada, China, India, Nigeria, Pakistan, Russian Federation, United Kingdom, United Republic of Tanzania
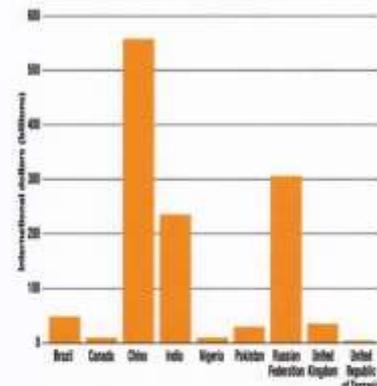
THE PROBLEM HAS SERIOUS IMPACT

The burden of chronic disease:
- has major adverse effects on the quality of life of affected individuals;
- causes premature death;
- creates large adverse – and underappreciated – economic effects on families, communities and societies in general.

$558 billion

The estimated amount China will forego in national income over the next 10 years as a result of premature deaths caused by heart disease, stroke and diabetes

CHRONIC DISEASE

Chronic Diseases 1

The burden and costs of chronic diseases in low-income and middle-income countries

Dele O Abegunde, Colin D Mathers, Taghreed Adam, Monica Ortegon, Kathleen Strong

This paper estimates the disease burden and loss of economic output associated with chronic diseases—mainly cardiovascular diseases, cancer, chronic respiratory diseases, and diabetes—in 23 selected countries which account for around 80% of the total burden of chronic disease mortality in developing countries. In these 23 selected low-income and middle-income countries, chronic diseases were responsible for 50% of the total disease burden in 2005. For 15 of the selected countries where death registration data are available, the estimated age-standardised death rates for chronic diseases in 2005 were 54% higher for men and 86% higher for women than those for men and women in high-income countries. If nothing is done to reduce the risk of chronic diseases, an estimated US$84 billion of economic production will be lost from heart disease, stroke, and diabetes alone in these 23 countries between 2006 and 2015. Achievement of a global goal for chronic disease prevention and control—an additional 2% yearly reduction in chronic disease death rates over the next 10 years—would avert 24 million deaths in these countries, and would save an estimated $8 billion, which is almost 10% of the projected loss in national income over the next 10 years.

FIGURE 8: **Preventable Cases of Chronic Diseases**
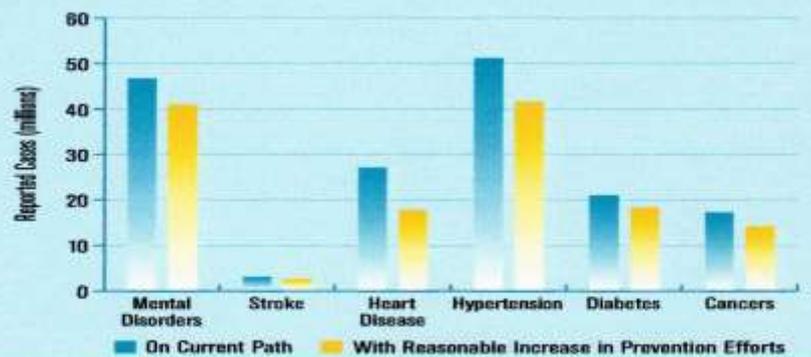
SOURCE: Adapted from R. DeVol, et al., An Unhealthy America: The Economic Burden of Chronic Disease, (Santa Monica, CA: Milken Institute, October 2007), www.milkeninstitute.org/publications/publications.taf?function=detail&ID=38801019&cat=ResRep (accessed 6 February 2009).
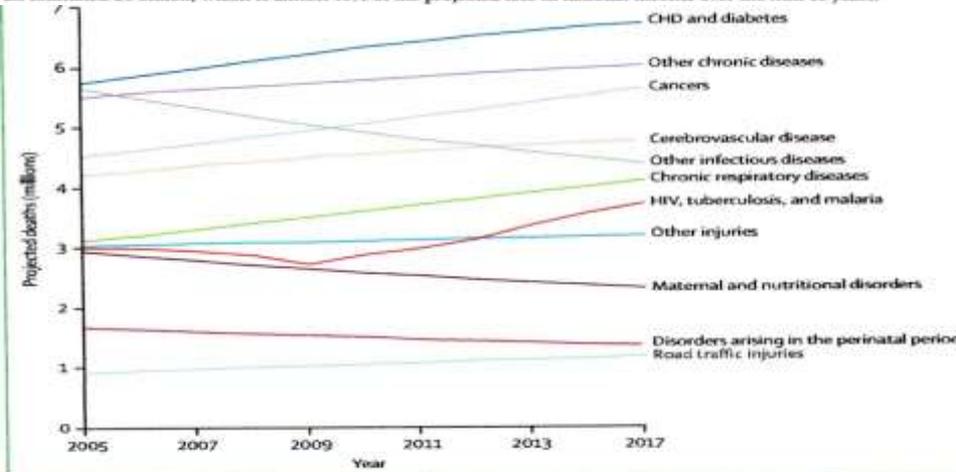

Figure 3: Projected global deaths (millions) for major chronic disease groups and other causes of death in 23 selected countries, 2005–15
CHD=coronary heart disease.

PROJECTED DEATHS BY CAUSE FOR HIGH-, MIDDLE- AND LOW-INCOME COUNTRIES
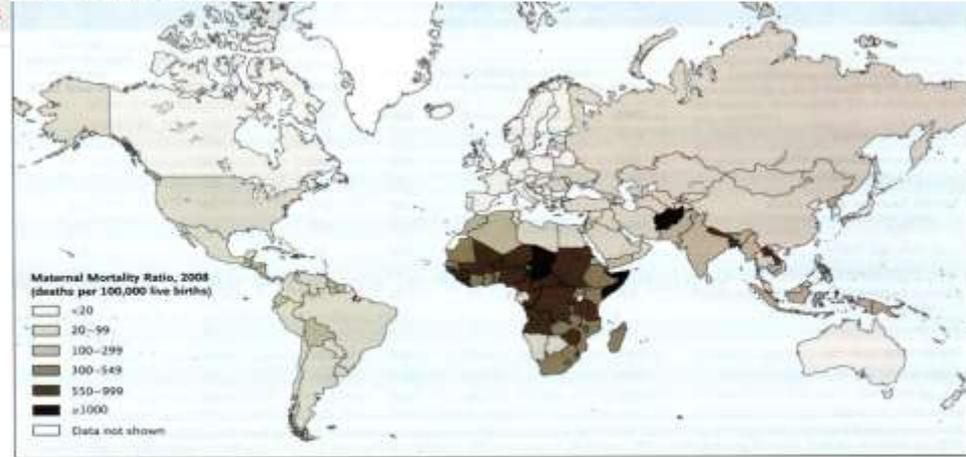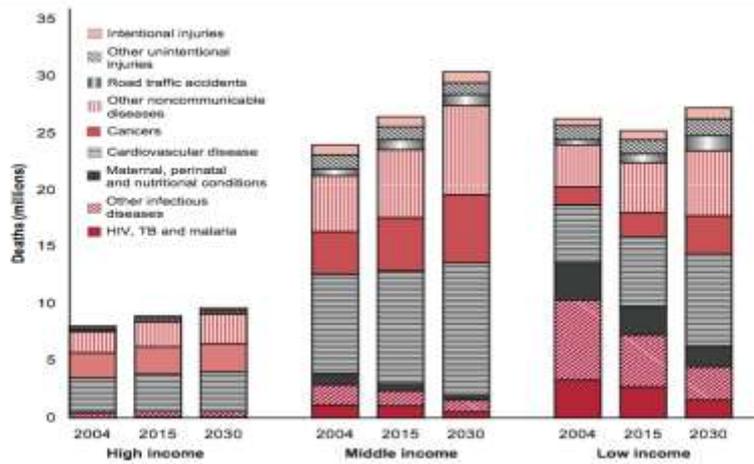
Maternal Mortality Ratios, 2008.

Data are the numbers of maternal deaths per 100,000 live births. Data are from the WHO, UNICEF, the United Nations Population Fund, and the World Bank. The boundaries used do not imply official endorsement or acceptance by the United Nations. An interactive map showing changes in maternal mortality ratios over time is available with the full text of this article at NEJM.org.

European
Observatory
on Health Systems and Policies

Virtuous Cycle

Prevent the preventable

Sustain through prevention

Towards a sustainable social policy

A safer future

Better health, better use of resources

Prevention through sustainability

Manage unavoidable disease

Figure 1

TACKLING CHRONIC DISEASE IN EUROPE

Strategies, interventions and challenges

Reinhard Busse, Miriam Blümel,
David Scheller-Kreinsen,
Annette Zentner

Table 2.1 Disease burden and deaths from noncommunicable diseases in the WHO European Region by cause (2005)

Groups of causes	Disease burden		Deaths	
	DALYs (millions)	Proportion from all causes (%)	Number (millions)	Proportion from all causes (%)
Selected noncommunicable diseases				
Cardiovascular diseases	34.42	23	5.07	52
Neuropsychiatric conditions	29.37	20	0.26	3
Cancer (malignant neoplasms)	17.03	11	1.86	19
Digestive diseases	7.12	5	0.39	4
Respiratory diseases	6.84	5	0.42	4
Sense organ diseases	6.34	4	0	0
Musculoskeletal diseases	5.75	4	0.03	0
Diabetes mellitus	2.32	2	0.15	2
Oral conditions	1.02	1	0	2
All noncommunicable diseases	*115.34*	*77*	*8.21*	*86*
All causes	*150.32*	*100*	*9.56*	*100*

Source: Adapted from Singh 2008.

	2005	2015	2030
Deaths (all ages)			
CVD and diabetes	12·4 (33%)	14·3 (35%)	17·3 (36%)
Cancers	4·5 (12%)	5·6 (14%)	7·5 (15%)
Chronic respiratory	3·1 (8%)	4·1 (10%)	5·9 (12%)
All chronic diseases	23·1 (61%)	27·2 (66%)	34·3 (71%)
Deaths in people younger than 70 years			
CVD and diabetes	5·2 (21%)	5·6 (23%)	5·9 (23%)
Cancers	2·8 (12%)	3·4 (14%)	4·1 (16%)
Chronic respiratory	1·1 (5%)	1·4 (6%)	1·9 (7%)
All chronic diseases	11·2 (46%)	12·4 (50%)	13·7 (53%)
DALYs			
CVD and diabetes	121 (12%)	131 (13%)	145 (14%)
Cancers	51 (5%)	59 (6%)	71 (7%)
Chronic respiratory	42 (4%)	49 (5%)	62 (6%)
All chronic diseases	496 (50%)	538 (55%)	597 (59%)

CVD=cardiovascular disease. DALYs=disability-adjusted life-years.

Table 1: Millions of projected deaths and DALYs for chronic diseases as a proportion of deaths and DALYs for all causes in 23 selected countries for 2005, 2015, and 2030

Measuring Disability Prevalence

Daniel Mont

March 2007

TOGETHER WE CAN
TOGETHER WE WILL

Social **Protection**
THE WORLD BANK

Figure 1: The ICF Model

Health Condition
(disorder/disease)

Body Function & Structure *(Impairment)* ⟷ Activities *(Limitation)* ⟷ Participation *(Restriction)*

Environmental Factors

Personal Factors
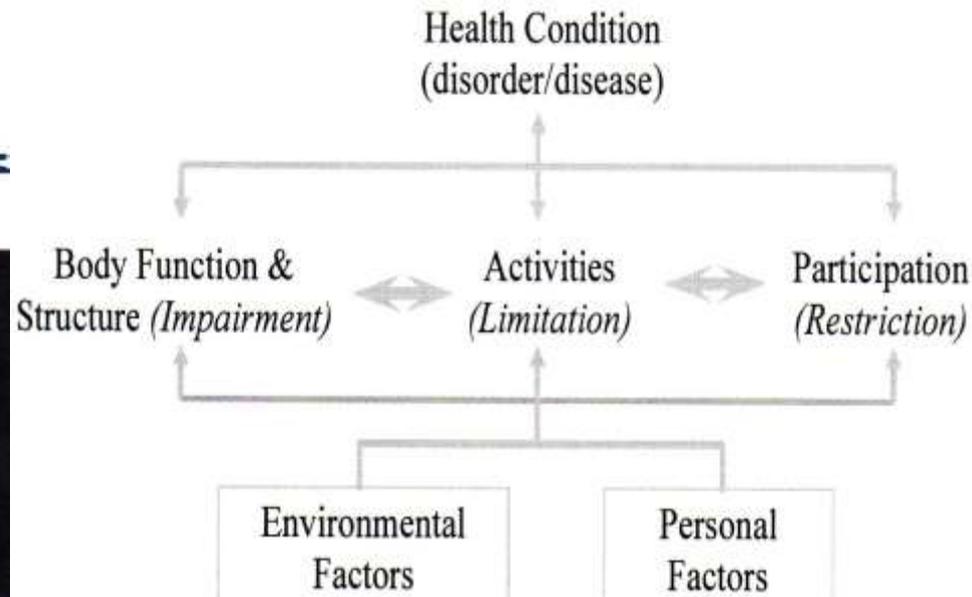
Box 1: WHO-Disability Assessment Schedule -- 12-Question Set

In the last 30 days how much difficulty did you have in:
(None Mild Moderate Severe Extreme Cannot Do)

Standing for long periods such as 30 minutes?

Taking care of your household responsibilities?

Learning a new task, for example, learning how to get to a new place?

How much of a problem did you have joining in community activities (for example, festivities, religious or other activities) in the same way as anyone else can?

How much have you been emotionally affected by your health problems?

Concentrating on doing something for ten minutes?

Walking a long distance such as a kilometre [or equivalent]?

Washing your whole body?

Getting dressed?

Dealing with people you do not know?

Maintaining a friendship?

Your day to day work?

Source: World Health Organization: http://www.who.int/icidh/whodas/index.html

Current and future worldwide prevalence of dependency, its relationship to total population, and dependency ratios

Rowan H. Harwood,[1] Avan Aihie Sayer,[2] & Miriam Hirschfeld[3]

Objective To estimate the number of people worldwide requiring daily assistance from another person in carrying out health, domestic or personal tasks.

Methods Data from the Global Burden of Disease Study were used to calculate the prevalence of severe levels of disability, and consequently, to estimate dependency. Population projections were used to forecast changes over the next 50 years.

Findings The greatest burden of dependency currently falls in sub-Saharan Africa, where the "dependency ratio" (ratio of dependent people to the population of working age) is about 10%, compared with 7–8% elsewhere. Large increases in prevalence are predicted in sub-Saharan Africa, the Middle East, Asia and Latin America of up to 5-fold or 6-fold in some cases. These increases will occur in the context of generally increasing populations, and dependency ratios will increase modestly to about 10%. The dependency ratio will increase more in China (14%) and India (12%) than in other areas with large prevalence increases. Established market economies, especially Europe and Japan, will experience modest increases in the prevalence of dependency (30%), and in the dependency ratio (up to 10%). Former Socialist economies of Europe will have static or declining numbers of dependent people, but will have large increases in the dependency ratio (up to 13%).

Conclusion Many countries will be greatly affected by the increasing number of dependent people and will need to identify the human and financial resources to support them. Much improved collection of data on disability and on the needs of caregivers is required. The prevention of disability and provision of support for caregivers needs greater priority.

Keywords Dependency (Psychology); Population dynamics; Aging; Chronic disease; Disabled persons; Health services needs and demand; Activities of daily living; Cost of illness; Forecasting (source: MeSH, NLM).

Mots clés Dépendance (Psychologie); Dynamique population; Vieillissement; Maladie chronique; Handicapé; Besoins et demande services santé; Activité quotidienne; Coût maladie; Prévision (source: MeSH, INSERM).

Palabras clave Dependencia (Psicología); Dinámica de población; Envejecimiento; Enfermedad crónica; Evaluación de la incapacidad; Necesidades y demanda de servicios de salud; Actividades cotidianas; Costo de la enfermedad; Predicción (fuente: DeCS, BIREME).

الكلمات المفتاحية: ترمية الحياة الحالة الصحية: مؤشرات الحالة الصحية؛ قياس الأله الإفصاح الذاتي عن المعلومات؛ العوامل الاجتماعية الاقتصادية الخصائص الثانية؛ تحليل التحوُّل؛ حزب أفريقيا، رلضمن؛ زرى المؤهزعات الفئة- ذلك الإقليمي لشرق فلسطين

Table 1. **Twenty-two indicator conditions used to determine disability severity scores, with their disability classes, and the frequency of care needs as rated by health professionals**

Short description of condition	Disability class	Median dependency rating[a]
Active psychosis	7	1
Dementia	7	1
Quadriplegia	7	1
Severe continuous migraine	7	1
Blind	6	1
Paraplegic	6	1
Severe depression	6	1
Down syndrome	5	1
Mild mental retardation	5	1
Recto-vaginal fistula	5	2
Below-knee amputation	4	2
Deafness	4	2
Infertility	3	3
Fracture radius	3	2
Rheumatoid arthritis	3	2
Impotence	3	3
Angina after walking 50 m	3	2
Severe continuous sore throat	2	2,5
Anaemia	2	2
Diarrhoea	2	2
Severe thinness	1	2.5
Vitiligo	1	3

* 1, daily help; 2, weekly help; 3, less than weekly help.

The Assessment of Chronic Health Conditions on Work Performance, Absence, and Total Economic Impact for Employers

James J. Collins, PhD
Catherine M. Baase, MD
Claire E. Sharda, RN, MBA
Ronald J. Ozminkowski, PhD
Sean Nicholson, PhD
Gary M. Billotti, MS
Robin S. Turpin, PhD
Michael Olson, PhD
Marc L. Berger, MD

C

Objective: The objective of this study was to determine the prevalence and estimate total costs for chronic health conditions in the U.S. workforce for the Dow Chemical Company (Dow). **Methods:** Using the Stanford Presenteeism Scale, information was collected from workers at five locations on work impairment and absenteeism based on self-reported "primary" chronic health conditions. Survey data were merged with employee demographics, medical and pharmaceutical claims, smoking status, biometric health risk factors, payroll records, and job type. **Results:** Almost 65% of respondents reported having one or more of the surveyed chronic conditions. The most common were allergies, arthritis/joint pain or stiffness, and back or neck disorders. The associated absenteeism by chronic condition ranged from 0.9 to 5.9 hours in a 4-week period, and on-the-job work impairment ranged from a 17.8% to 36.4% decrement in ability to function at work. The presence of a chronic condition was the most important determinant of the reported levels of work impairment and absence after adjusting for other factors ($P < 0.000$). The total cost of chronic conditions was estimated to be 10.7% of the total labor costs for Dow in the United States; 6.8% was attributable to work impairment alone. **Conclusion:** For all chronic conditions studied, the cost associated with performance based work loss or "presenteeism" greatly exceeded the combined costs of absenteeism and medical treatment combined. (J Occup Environ Med. 2005;47:547-557)
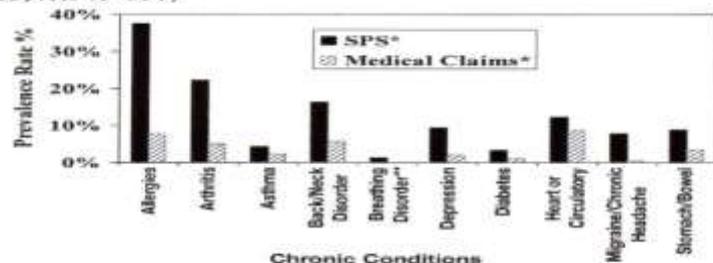
Fig. 2. *Prevalence rates from the SPS of any chronic conditions in the last 4 weeks and chronic conditions reported from medical and pharmacy claims in the last year. **Breathing disorders (bronchitis and emphysema).
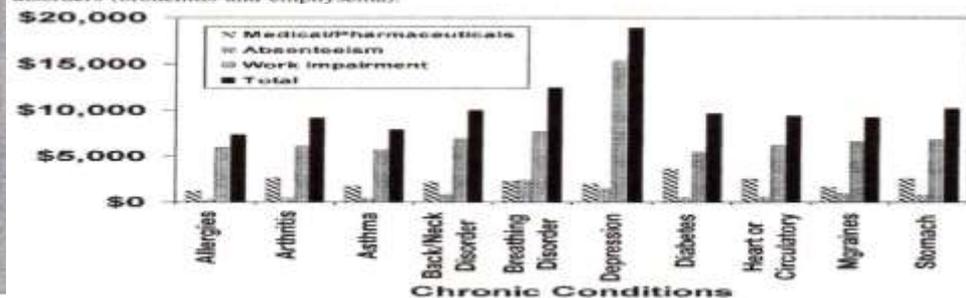

Fig. 3. Chronic Conditions.

Old Age Dependency Ratio by Region
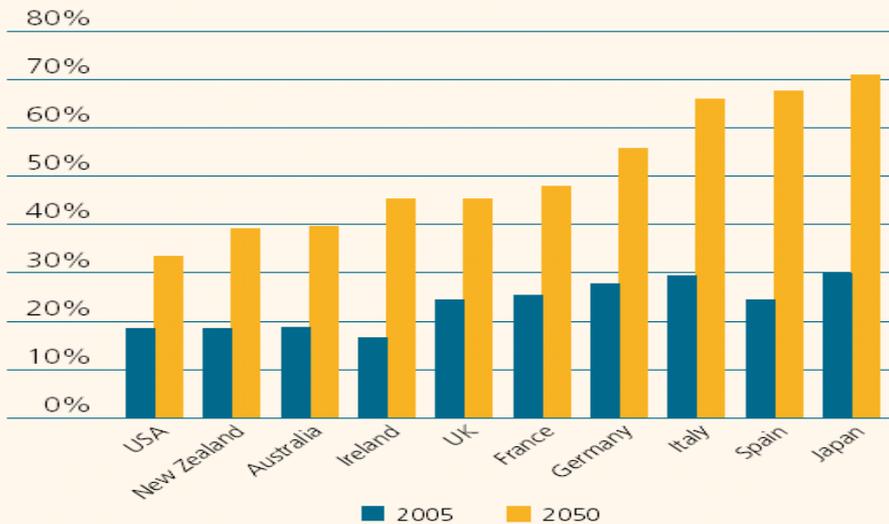
Figure 16. Old-age dependency ratio: major areas, 2009 and 2050
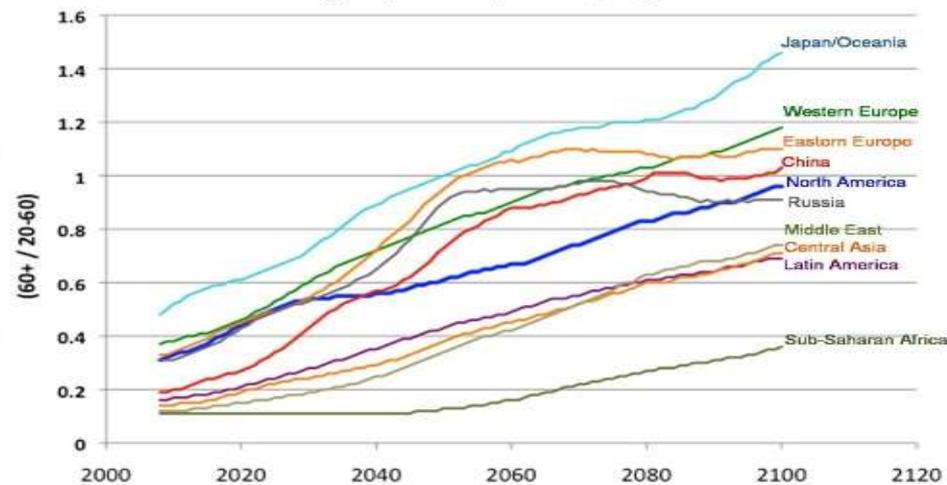
Figure 14. Total dependency ratio: world and development regions, 1950-2050
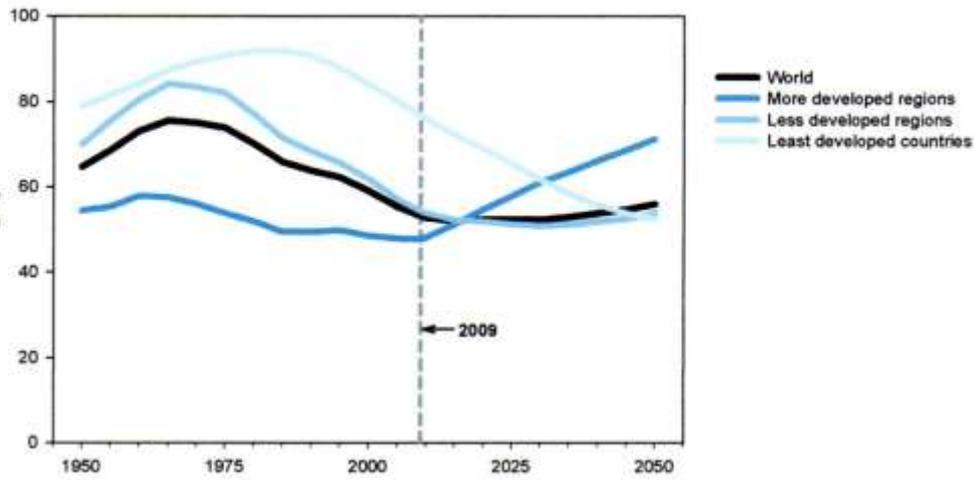
Figure 7-2.
Older Dependency Ratio: 2008
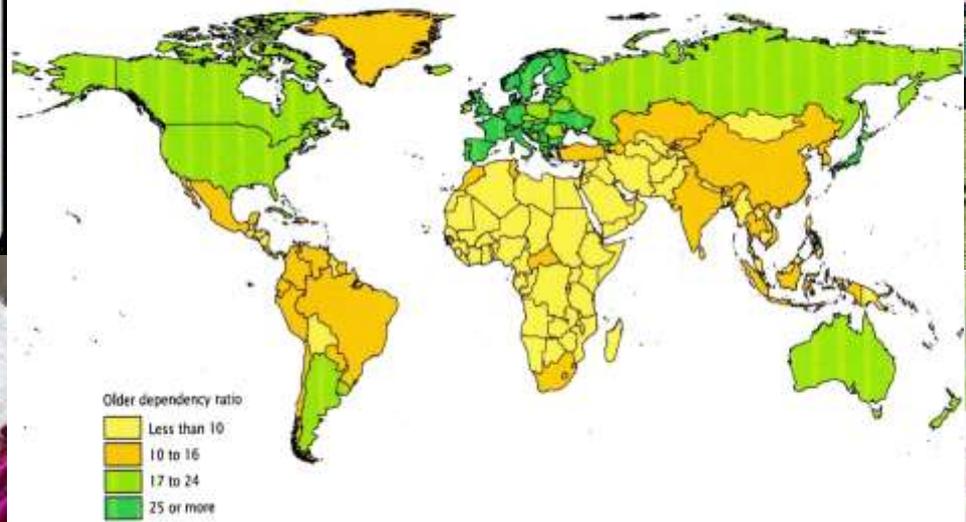
Older dependency ratio
- Less than 10
- 10 to 16
- 17 to 24
- 25 or more

Note: Older dependency ratio is the number of people aged 65 and over per 100 people aged 20 to 64.
Source: U.S. Census Bureau, International Data Base, accessed on May 27, 2008.

Figure 7-3.
Older Dependency Ratio for World Regions: 2000, 2020, and 2040
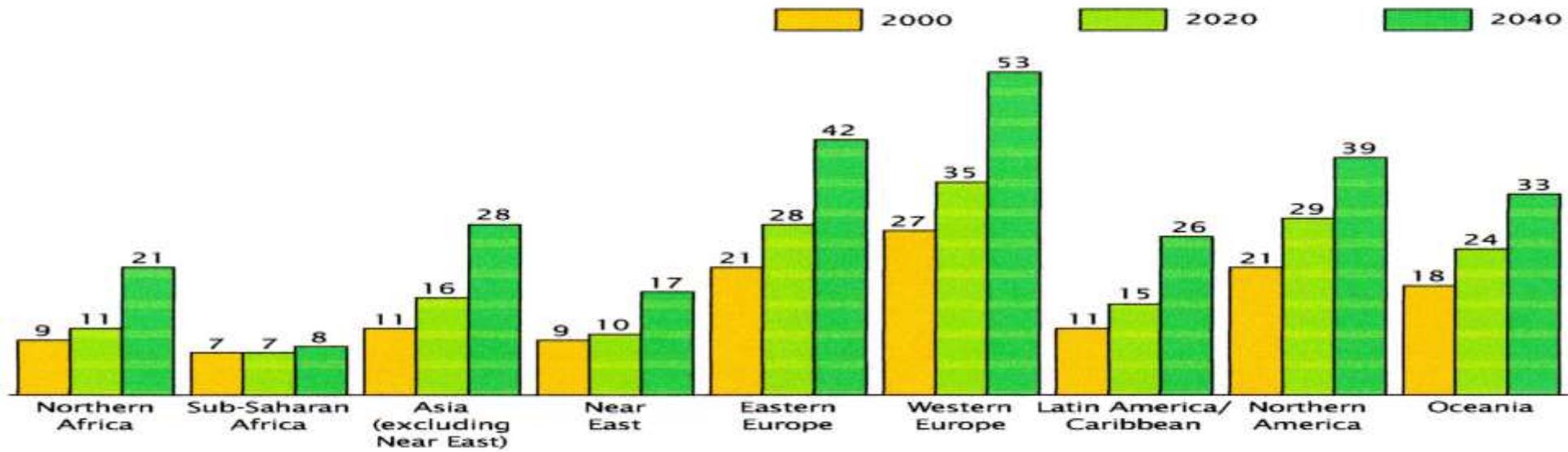
- 2000
- 2020
- 2040

Region	2000	2020	2040
Northern Africa	9	11	21
Sub-Saharan Africa	7	7	8
Asia (excluding Near East)	11	16	28
Near East	9	10	17
Eastern Europe	21	28	42
Western Europe	27	35	53
Latin America/ Caribbean	11	15	26
Northern America	21	29	39
Oceania	18	24	33

Note: Older dependency ratio is the number of people aged 65 and over per 100 people aged 20 to 64.
Source: U.S. Census Bureau, International Data Base, accessed on January 10, 2008.

DISABILITY ISSUES, TRENDS AND RECOMMENDATIONS FOR THE WORLD BANK

(FULL TEXT AND ANNEXES)

Figure 1.1: The Disablement Phenomena as Conceptualized in the Original ICIDH
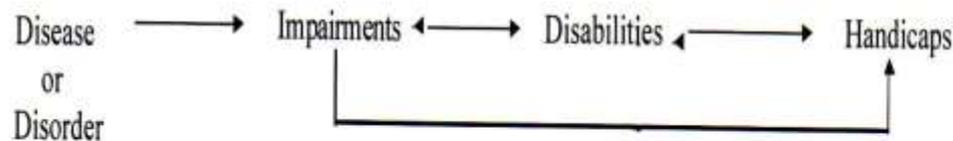
Source: World Health Organization, *ICIDH-2*. 11.

Figure 1.2: Current Understanding of Interactions within ICIDH-2 Dimensions
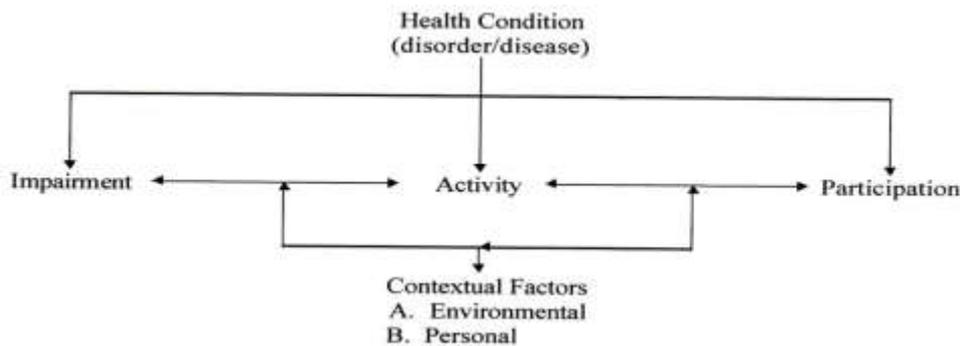
Source: World Health Organization, *ICIDH-2*. 12.

ROBERT L. METTS, PH.D.

FEBRUARY, 2000

Table 1.1: Estimated Ranges of Populations of People with Disabilities

Human Development Category	Low Estimate	High Estimate
High Human Development Countries	124,226,190	124,226,190
Medium Human Development Countries	93,517,500	250,222,500
Low Human Development Countries	17,650,000	174,735,000
TOTAL	235,393,690	549,183,690

Table 1.2: Total Annual Value of GDP Lost Due to Disability

Value of GDP Lost (US Dollars)	High Estimate	Low Estimate
High Income Countries	1,264,232,430,105	891,283,863,224
Medium Income Countries	480,206,038,845	338,545,257,386
Low Income Countries	192,002,986,035	135,362,105,155
TOTAL	$ 1,936,441,454,985	$ 1,365,191,225,765

Figure 15. Composition of the total dependency ratio: world and development regions, 1950-2050
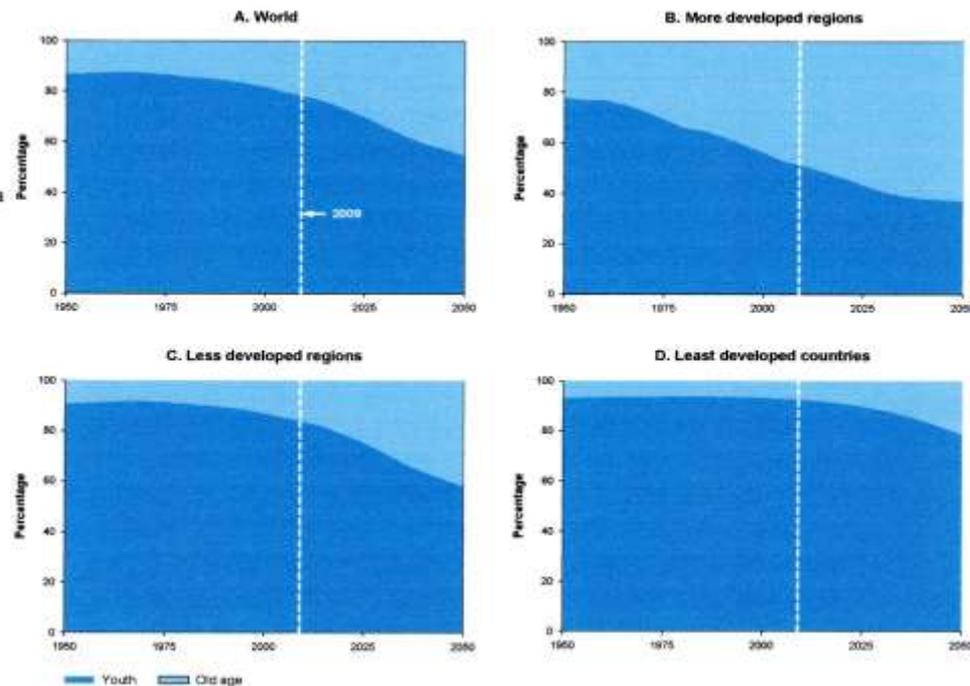
Futurologia Falaciosa

- "… the war against infectious diseases is over, because we have won…" (sic.)(US Surgeon General, 1967)

- "… no new diseases are going to be discovered …" (sic.) (L. Thomas Dean of Yale Medical School, 1976)

- WHO (2008)
  - D. Infecciosas 15.000.000 mortes / ano
    - 2ª causa de Mortalidade Mundial
      - 1ª Causa nos Países em Desenvolvimento
  - Novos Agentes Patogénicos para a Espécie Humanas descritos nos últimos 40 anos: 50
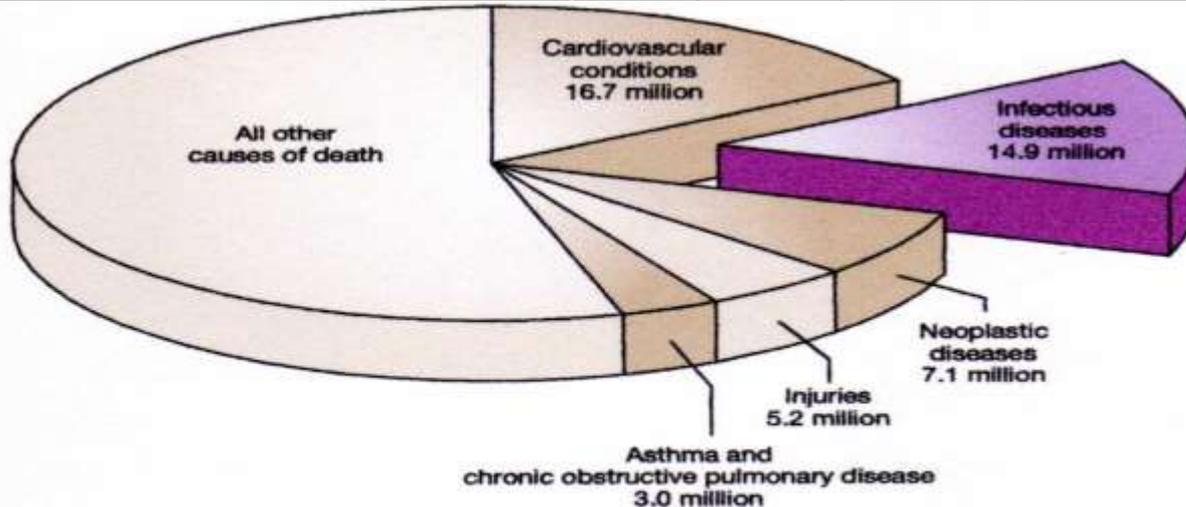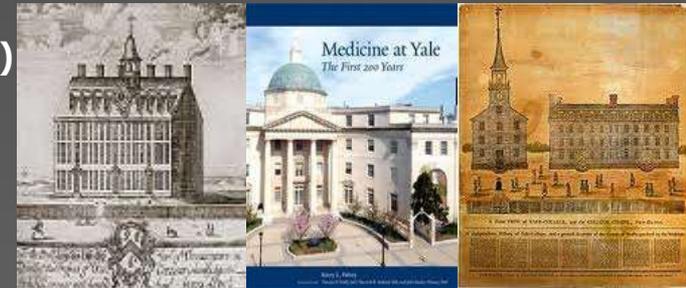
Infectious diseases	Annual deaths (million)
Respiratory infections	3.96
HIV/AIDS	2.77
Diarrhoeal diseases	1.80
Tuberculosis	1.56
Vaccine-preventable childhood diseases	1.12
Malaria	1.27
STDs (other than HIV)	0.18
Meningitis	0.17
Hepatitis B and C	0.16
Tropical parasitic diseases	0.13
Dengue	0.02
Other infectious diseases	1.76

Figure 2 Leading causes of death worldwide. About 15 million (>25%) of 57 million annual deaths worldwide are the direct result of infectious disease. Figures published by the World Health Organization (see http://www.who.int/whr/en and ref. 7).

"...nowadays we see maladies unknown to our forefathers springing up around us".

Ullrich von Hutten, 1519[36]

"After our fathers and grandfathers succumbed... dogs and vultures devoured the bodies. So it was that we became orphans, oh, my sons! So we became when we were young. All of us were thus. We were born to die!"

Cakchiquel Mayan on the hueyzahuatl epidemic of 1520–21[44]

"If there is, within medicine, any subject worthy of... investigation... it is, without doubt, epidemic pestilential diseases, obscure and hidden in their causes, rapid in their progression, frightening in their symptoms, and deadly in their consequences."

Félix Vicq-d'Azyr, 1776[51]
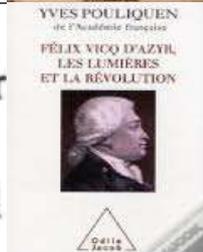
"France and its capital have been visited by a fearful pestilence... It comes without any known cause; it disappears without any revealed reason. The bodies of its victims are in vain examined; death is interrogated: death betrays nothing."

Louis-François Benoiston de Châteauneuf, 1834[69]

"Not a single year passes without [which]... we can tell the world: here is a new disease!"

Rudolf Virchow, 1867[1]

"The future of microbes and mankind will probably unfold as episodes of a suspense thriller that could be entitled Our wits versus their genes".

J. Lederberg, Science 2000[33]
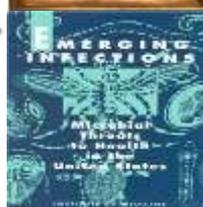
Les grandes étapes dela recherche

■ Par Marine Benoiste, avec la collaboration de Jérôme Pierrel, historien des sciences au laboratoire Épistémé de l'université Bordeaux-I

DU MICROBE ENNEMI AU MICROBE ALLIÉ

▶1878
Le médecin Charles Sédillot crée le terme de « microbe » pour nommer l'ensemble des organismes vivants invisibles à l'œil nu. Plus tard, ils seront qualifiés de micro-organismes.

▶1880
Les Français Louis Pasteur, Charles Chamberland et Émile Roux, développent un vaccin contre le choléra des poules, à l'aide de cultures du microbe (lire « Principes de mort et principes de vie », p. 64).

▶1882
Le médecin allemand Robert Koch pense identifier le bacille de la tuberculose. Pour confirmer sa découverte, il le met en culture et l'utilise pour reproduire la maladie chez des animaux.

▶1884
Robert Koch et le bactériologiste allemand Friedrich Löffler définissent les critères pour établir un lien de cause à effet entre un micro-organisme et une maladie.

▶1885
En injectant de la moelle épinière de lapin atteint de la rage à un enfant victime de morsures par un chien enragé, Pasteur prévient tout développement de la maladie. C'est le premier vaccin contre la rage.

▶1892
Après filtration de la sève de feuilles atteintes de la mosaïque du tabac, le Russe Dmitri I. Ivanovski montre que l'agent responsable de cette maladie végétale est trop petit pour être une bactérie.

▶1898
Le botaniste néerlandais Martinus W. Beijerinck attribue la maladie de la mosaïque du tabac à un agent infectieux « vivant » et « fluide ». Pour désigner ce dernier, il emploie le terme « virus ».

▶1907
Elie Metchnikoff, immunologiste russe, étudie les bactéries produisant le lait fermenté ; il suggère que ces bactéries sont bénéfiques pour la santé humaine.

▶1915
L'Anglais Frederick Twort découvre des virus qui infectent spécifiquement les bactéries.

▶1917
Le Français Félix d'Hérelle les redécouvre et les nomme « bactériophages » (lire « Des virus pour lutter contre les bactéries », p.42).

▶1928
Une substance antibactérienne produite par des moisissures *Penicillium notatum* est isolée inopinément par le biologiste

écossais Alexander Fleming. C'est la pénicilline.

▶1939
Grâce à l'un des premiers microscopes électroniques, Gustav-Adolf Kausche, Edgar Pfankuch et Helmut Ruska visualisent l'agent de la mosaïque du tabac. C'est le premier virus observé.

▶1944
La société pharmaceutique américaine Pfizer inaugure la première usine de production de pénicilline. Les soldats alliés disposent de cet antibiotique lors du Débarquement en Normandie.

▶1972
Le biochimiste américain Paul Berg et ses collègues construisent un ADN constitué d'ADN bactérien et d'ADN viral : c'est le premier ADN recombinant (lire « La peur conjurée du génie génétique », p. 60).

ALEXANDER FLEMING a isolé la pénicilline à partir du champignon *Penicillium notatum*. Le biologiste écossais reçut le prix Nobel en 1945 pour cette découverte de ce qui allait être le premier antibiotique.
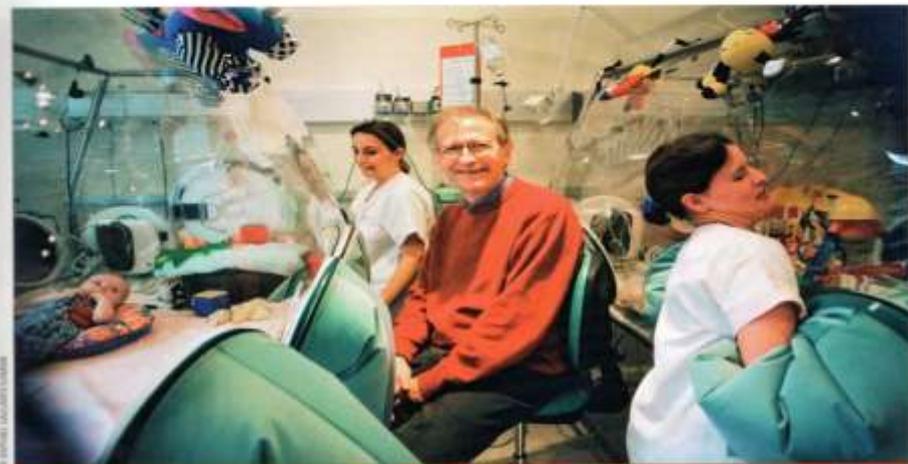
LE MÉDECIN ALAIN FISCHER et ses collègues de l'hôpital Necker ont utilisé avec succès la thérapie génique en 2000 pour traiter des nourrissons atteints de déficit immunitaire combiné sévère. Ces enfants étaient auparavant contraints de vivre sous une bulle stérile.

▶1973
Les Américains Stanley Cohen et Herbert Boyer insèrent un ADN recombinant portant des gènes de résistance à des antibiotiques dans une bactérie *Escherichia coli*. C'est le premier organisme génétiquement modifié.

▶1978
La compagnie pharmaceutique Genentech, fondée

par Herbert Boyer et Robert Swanson, transfère le gène de l'insuline humaine dans une bactérie. C'est l'une des premières fabrications d'hormone par un organisme transformé génétiquement.

▶1982
L'agence américaine de sécurité sanitaire, la FDA, donne son feu vert à la commercialisation d'insuline

produite par génie génétique. Il s'agit du premier médicament ainsi fabriqué industriellement (lire « Des mini-usines à médicaments », p. 46).

▶2000
Alain Fischer et Marina Cavazzana-Calvo, médecins à l'hôpital Necker à Paris, annoncent que deux nourrissons ont été traités avec succès par thérapie

génique : une première (lire « Comment on corrige des gènes », p. 46).

▶2006
Jeffrey Gordon et ses collaborateurs de l'université Washington, à Saint Louis, aux États-Unis, montrent que le microbiote intestinal (auparavant appelé « flore intestinale ») joue un rôle dans l'obésité

(lire « Des bactéries qui stockent des graisses », p. 28).

▶2010
Les premiers résultats du projet MetaHIT, mis en place pour caractériser génétiquement le microbiote intestinal humain, sont publiés. 3,3 millions de gènes bactériens ont été identifiés (lire « Un autre génome pour l'homme », p. 22).
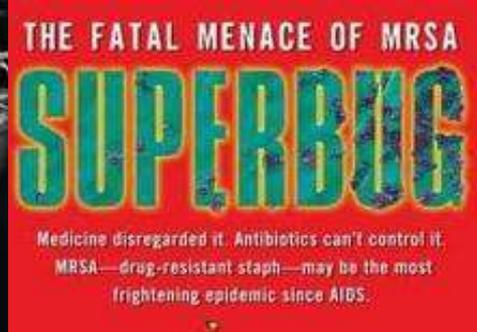
GET SMART
Know When Antibiotics Work

Who's hogging our antibiotics?
Up to 70% of U.S. antibiotics go to farm animals that aren't sick.
SaveAntibiotics.org   PEW

THE FATAL MENACE OF MRSA
SUPERBUG
Medicine disregarded it. Antibiotics can't control it.
MRSA—drug-resistant staph—may be the most frightening epidemic since AIDS.

The Danger of Antibiotic Overuse

ELSEVIER

REVIEW ARTICLE

Resistance to Antibiotics: Are We in the Post-Antibiotic Era?

Alfonso J. Alanis

Lilly Research Laboratories, Eli Lilly and Company, Indianapolis, Indiana

Serious infections caused by bacteria that have become resistant to commonly used antibiotics have become a major global healthcare problem in the 21st century. They not only are more severe and require longer and more complex treatments, but they are also significantly more expensive to diagnose and to treat. Antibiotic resistance, initially a problem of the hospital setting associated with an increased number of hospital-acquired infections usually in critically ill and immunosuppressed patients, has now extended into the community causing severe infections difficult to diagnose and treat. The molecular mechanisms by which bacteria have become resistant to antibiotics are diverse and complex. Bacteria have developed resistance to all different classes of antibiotics discovered to date. The most frequent type of resistance is acquired and transmitted horizontally via the conjugation of a plasmid. In recent times new mechanisms of resistance have resulted in the simultaneous development of resistance to several antibiotic classes creating very dangerous multidrug-resistant (MDR) bacterial strains, some also known as "superbugs". The indiscriminate and inappropriate use of antibiotics in outpatient clinics, hospitalized patients and in the food industry is the single largest factor leading to antibiotic resistance. In recent years, the number of new antibiotics licensed for human use in different parts of the world has been lower than in the recent past. In addition, there has been less innovation in the field of antimicrobial discovery research and development. The pharmaceutical industry, large academic institutions or the government are not investing the necessary resources to produce the next generation of newer safe and effective antimicrobial drugs. In many cases, large pharmaceutical companies have terminated their anti-infective research programs altogether due to economic reasons. The potential negative consequences of all these events are relevant because they put society at risk for the spread of potentially serious MDR bacterial infections. © 2005 IMSS. Published by Elsevier Inc.

Key Words: Antibiotic resistance, Bacterial resistance, New antibiotics, Antibiotic research.
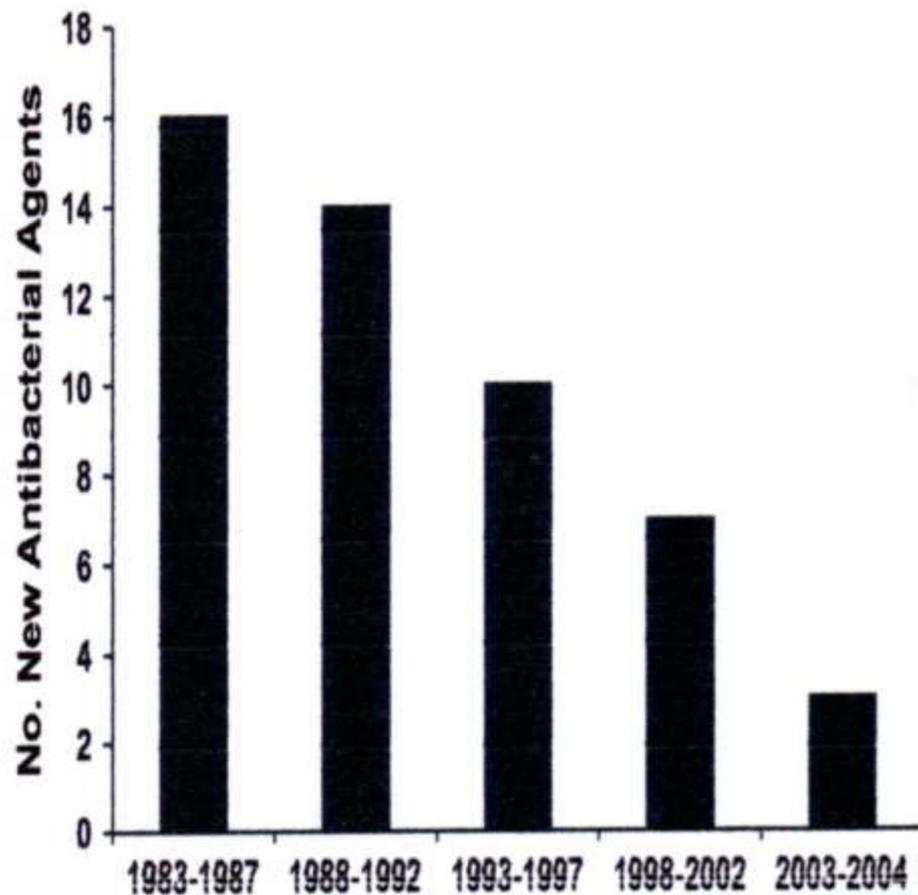
Figure 1. New antibacterial agents approved by the FDA in the U.S from 1983 to 2003. Data from Reference 70.

Os PRINICIPAIS PROBLEMAS no FUTURO

A Explosão Demográfica
A Limitação dos Recursos Naturais

- " … o crescimento populacional vai inevitavelmente tornar insuficiente a capacidade mundial de produção alimentar, razão pela qual irá imperar futuramente a fome e a miséria … " (sic.) (Thomas Malthus, 1803)

- " … calcula-se que entre 30 a 40 % do total de alimentos sejam desperdiçados … " (FAO Report, 2004)

- " … calcula-se que cerca de 20 % das emissões de gases nocivos para a atmosfera (designadamente metano e oxido nítrico), resultam das actividades relacionadas directa ou indirectamente com a agricultura … " (sic.) (H. Charles J. Godfrey, et al, 2010)

- " … nós não seremos capazes de alimentar todos os habitantes do mundo até ao final do presente milénio utilizando apenas os actuais meios tecnológicos e práticas. Insistir nessa falsa ideia de que isso será possível, só condenará milhões de pessoas à condição de famintos, bem como produzirá o caos social, económico e político … " (sic.) (Norman Borlaug)

- " … a prática actual da agricultura tem sido dominada por um discurso centrado nas questões tecnológicas, ignorando aspectos muito mais importantes de natureza económica e social, e negligenciado outros domínios do conhecimento, designadamente o saber tradicional e a experiência acumulada das populações. O resultado desta postura materializou-se na imposição inadequada de modelos de desenvolvimento menos consentâneos, de que a biotecnologia é uma das suas últimas variantes … " (sic.) (Miguel Altieri e Peter Rosset)

Em 2100, seremos um planeta de dez mil milhões de pessoas

Europa será o único continente a ver a sua população
decrescer nos próximos 90 anos

População mundial
Por região do globo

2011 / 2100

EUROPA 738.199 / 674.796

AMÉRICA DO NORTE 344.529 / 526.435

ÁSIA 4.164.252 / 4.596.234

AMÉRICA DO SUL 590.082 / 671.527

ÁFRICA 1.022.234 / 3.574.141

OCEANIA 36.593 / 65.818

MUNDO 6.895.889 / 10.124.926

Índia 1.550.899
China 1.341.335 / 941.042
Nigéria 310.384
EUA 478.026

Alemanha · Indonésia · Brasil · Japão · África do Sul · México

310.384 / 254.178
239.871 / 177.349
194.946 / 127.081
126.536 / 91.330
113.423 / 70.392
82.302 / 54.477
90.133

2010 · 2020 · 2030 · 2040 · 2050 · 2060 · 2070 · 2080 · 2090 · **2100**

Fonte: World Population Prospects, 2010 Revision. Organização das Nações Unidas

Change in Working-Age Population, 2005-30

Most advanced economies face shrinking workforces, while many young countries face
explosive growth.

**Change in Working-Age
Population, 2005-30**

- Shrinking (-25% to 0%)
- Slow Growth (0% to 25%)
- Moderate Growth (25% to 50%)
- Too Fast to Absorb (50% to 100%)
- Dangerously Fast (100% to 145%)

The new urban world

Key

US
246.2
81%
New York
18.9

Mexico
84.295
77%

Mexico City
18.1

Brazil
162.6
85%

São Paulo
18.4

Russia
103.6
73%
Moscow
11.4

China
559.2
42%

Shanghai
16.6

Beijing
11.7

Japan
84.7
66%

India
329.3
29%

Indonesia
114.1
50%

Philippines
55.0
63%

Turkey
51.1
68%

Iran
48.4
65%

Nigeria
68.6
50%

Cairo
15.9

3,307,950,000

UK
54.0
90%

London
12.0

Netherlands
13.2
81%

Poland
23.9
62%

Ukraine
30.9
68%

103.6
73%

13.4

Germany
62.0
75%

Czech
Republic
7.4

Belgium
10.2
27%

France
46.9
77%

Italy
39.6
68%

Romania
11.6
54%

Istanbul
11.7

Tehran
12.1

Kazakh-
stan
8.6

Uzbekistan
10.1
37%

Spain
33.6
77%

Portugal

Greece

Turkey
51.1
68%

Iran
48.4
68%

Afghan-
istan
7.8

Pakistan
59.3
36%

Cairo
15.9

Syria
10.2
51%

Iraq
20.3
67%

Morocco
19.4
60%

Algeria
22.0
65%

Tunisia

Egypt
33.1
43%

Saudi Arabia
20.9
81%

Karachi
14.8

Lagos
10.0

Nigeria

Sudan

Ethiopia

Yemen

Fig. 16. Urban and rural population, world, 1955-2015

De 2004 para cá, o índice dos preços da comida da FAO mais do que duplicou, enquanto os preços do petróleo foram de uns modestos 40 dólares o barril em 2000 para um pico de quase 140 em 2008. Os biocombustíveis estão a desviar comida da boca das pessoas para os depósitos automóveis. A água escasseia. O planeta está a consumir por ano, segundo a ONG World Wide Fund, 50% mais do que a terra consegue suportar. **É neste cenário que vamos bater mais um recorde de crescimento populacional**

Han, o bebé sete mil milhões...

... Haverá lugar para ele?

FEEDING THE FUTURE

Every 3.6 seconds a person dies of hunger

75% of them are children

The Future of Food and Farming:
Challenges and choices for global sustainability

UM FUTURO SEM 'SOL VERDE'

População (milhares de milhões de habitantes)

Ração diária (calorias disponíveis por pessoa e por dia)

Estimativas

1970 1990 2010 2030 2050
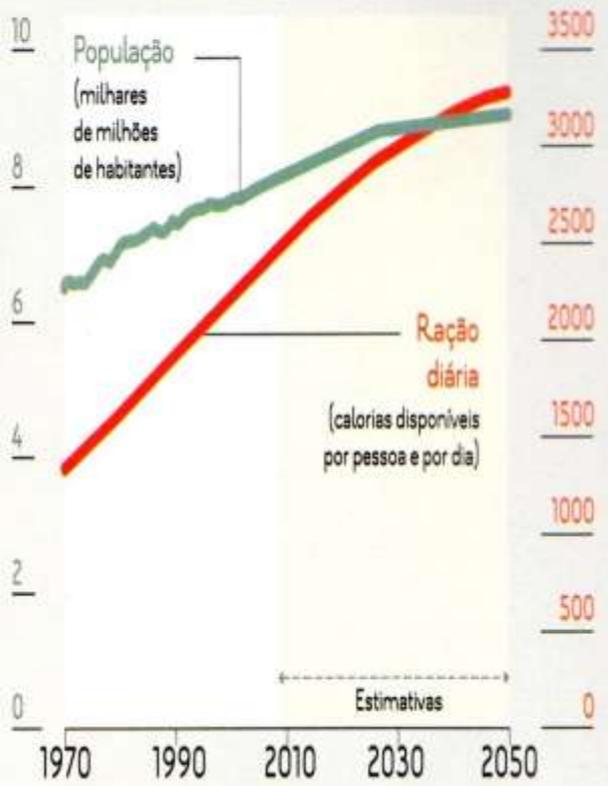
A bomba demográfica há muito tempo temida não chegará a explodir. Segundo as mais recentes projeções da ONU, a população mundial deverá estabilizar em redor dos nove mil milhões de habitantes, no ano 2050. Ao mesmo tempo, a produção agrícola e, logo, a ração diária disponível, deverá aumentar. Produzir comida suficiente para satisfazer as necessidades da população tornar-se-á possível, mas não sem ameaçar gravemente outros recursos, nomeadamente a água.

Fontes: Nature, Divisão de população da ONU, FAO

ORÇAMENTO FAMILIAR

Peso da alimentação nas despesas das famílias (percentagem, 2005)

País	Percentagem
EUA	6,2
França	10,6
Japão	12,3
Brasil	15,5
Irão	23,4
China	24,1
Tunísia	24,8
Roménia	25
Rússia	25,5
Índia	33,7
Egito	41,6
Mali	46,7
Arménia	65,1

Fonte: World Food Demand, Grace Gao, Universidade da Austrália Ocidental, 2010
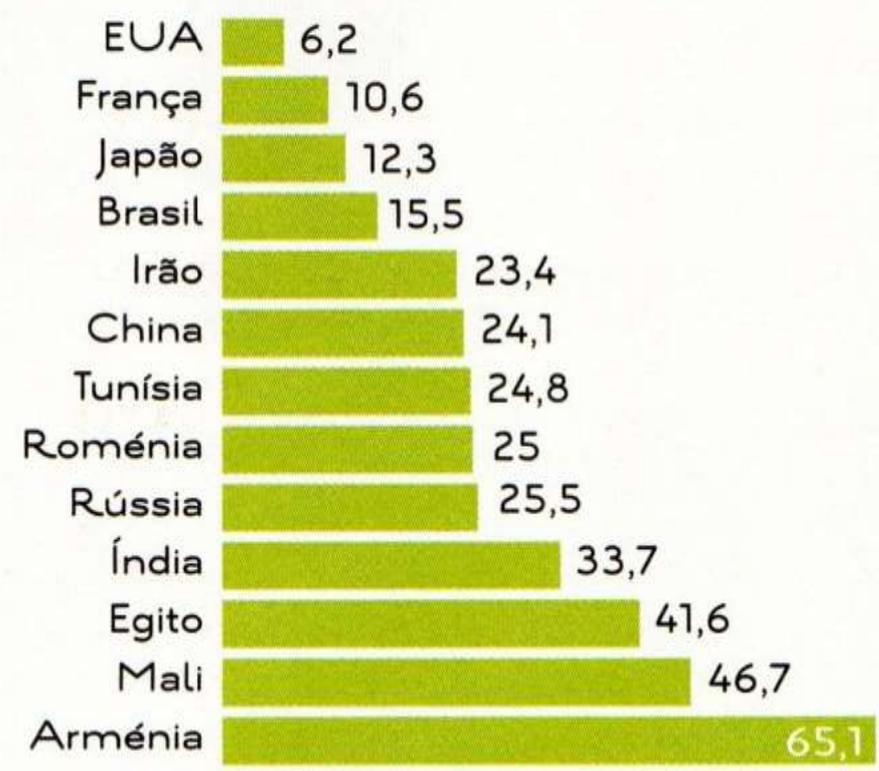
E para comer, quer cobra, canguru ou girafa?

Os portugueses passaram a ter à mesa novos sabores, exóticos e desconhecidos.
Fomos provar onze tipos de carne 'estranha', do kudu à zebra, passando
pela cobra, girafa ou canguru. E contamos a história de produtos
mentas nacionais. TEXTOS DE **KATYA DELIMBEUF**

alimentação | europa

Insetos procuram um lugar no menu dos europeus

Desde 2008 que a FAO
(Organização das Nações Unidas
para a Agricultura e Alimentação)
recomenda o consumo de insetos
ao invés da carne, por motivos tanto
económicos como ecológicos.
Mas na Europa, a mensagem não
é bem recebida e as iniciativas nesse
sentido são marginais.

Girafa

Cavalo

Kudu

Cobra pitão

Zebra

Veado

Órix

Bisonte

Crocodilo

Camelo

Population With Sustainable Access to an Improved Water Source (Percent) 2008
(Go to Table or Notes and Sources below)

0% - 13%   13% - 25%   25% - 38%   38% - 51%   51% - 63%   63% - 76%   76% - 88%
88% - 101%
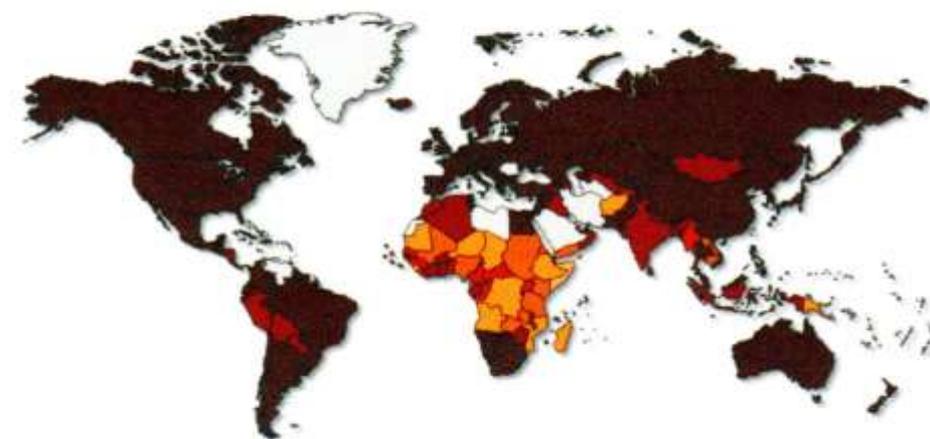
Availability of Freshwater in 2000
Average River Flows and Groundwater Recharge

Hyper arid
Arid
Semi-arid
Dry Subhumid
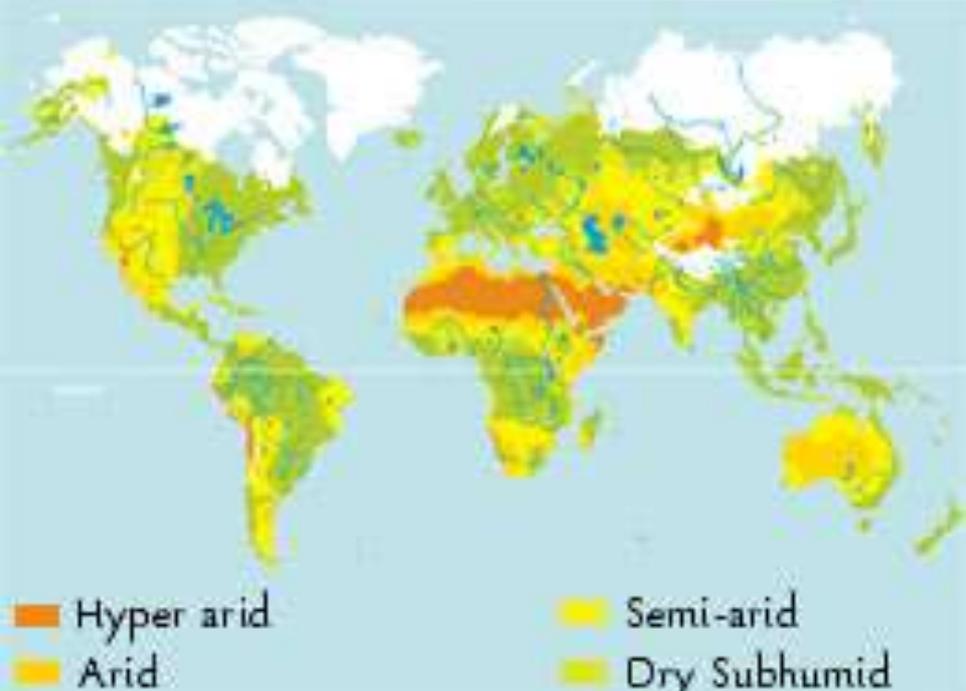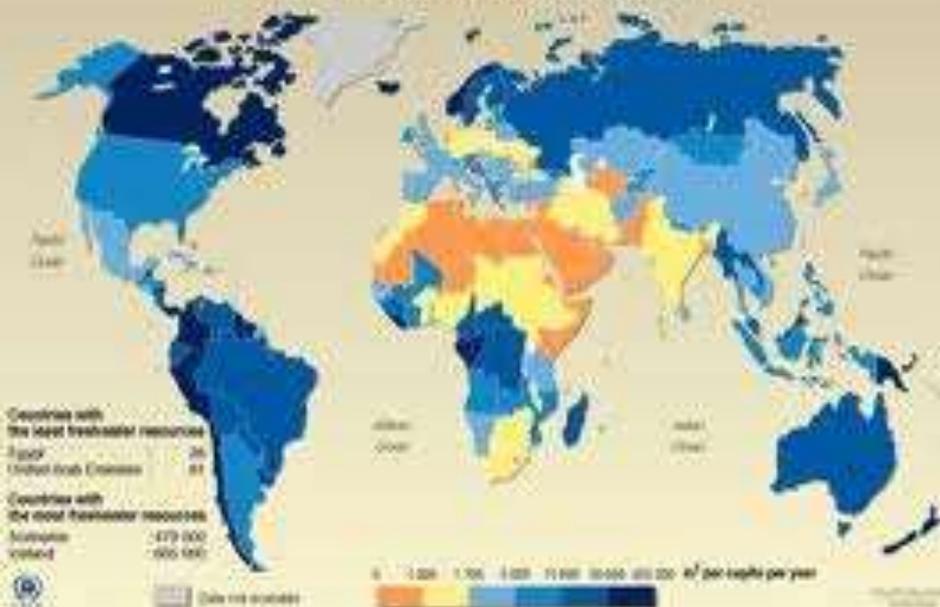
ENTÃO, E EM PORTUGAL?

Escrito na pedra

O país não precisa de quem diga o que está errado; precisa de quem saiba o que está certo.

Agustina Bessa-Luís (1922-), escritora portuguesa

Life expectancies

Life expectancy at birth
2002-2004, NUTS 2

- < 74.17
- 74.17 to 76.84
- 76.84 to 78.21
- 78.21 to 78.65
- 78.65 to 78.95
- 78.95 to 79.30
- 79.30 to 79.70
- 79.70 to 80.17
- 80.17 to 80.62
- > 80.62
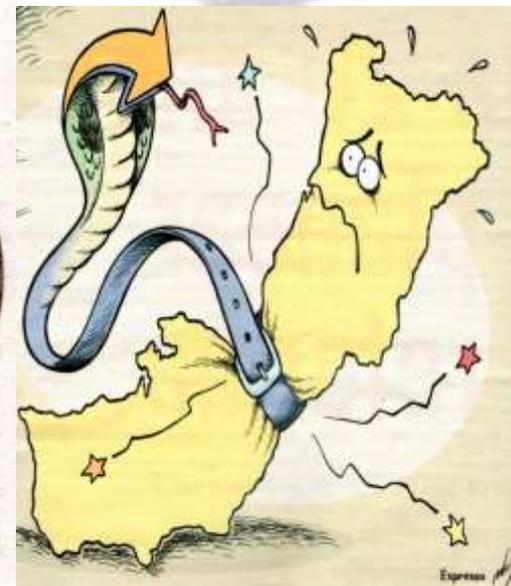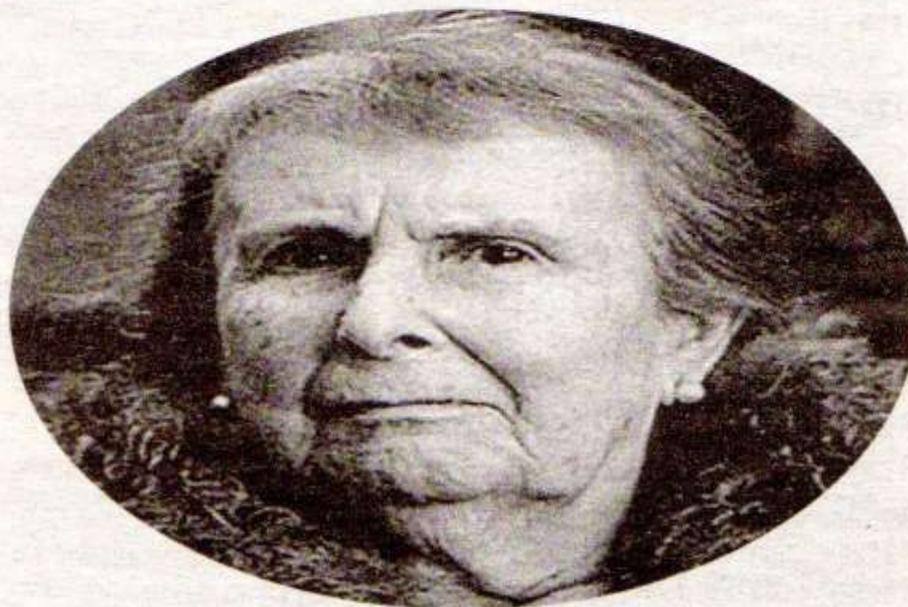
Figure 2-4.
The World's 25 Oldest Countries: 2008
(Percent of population aged 65 years and over)
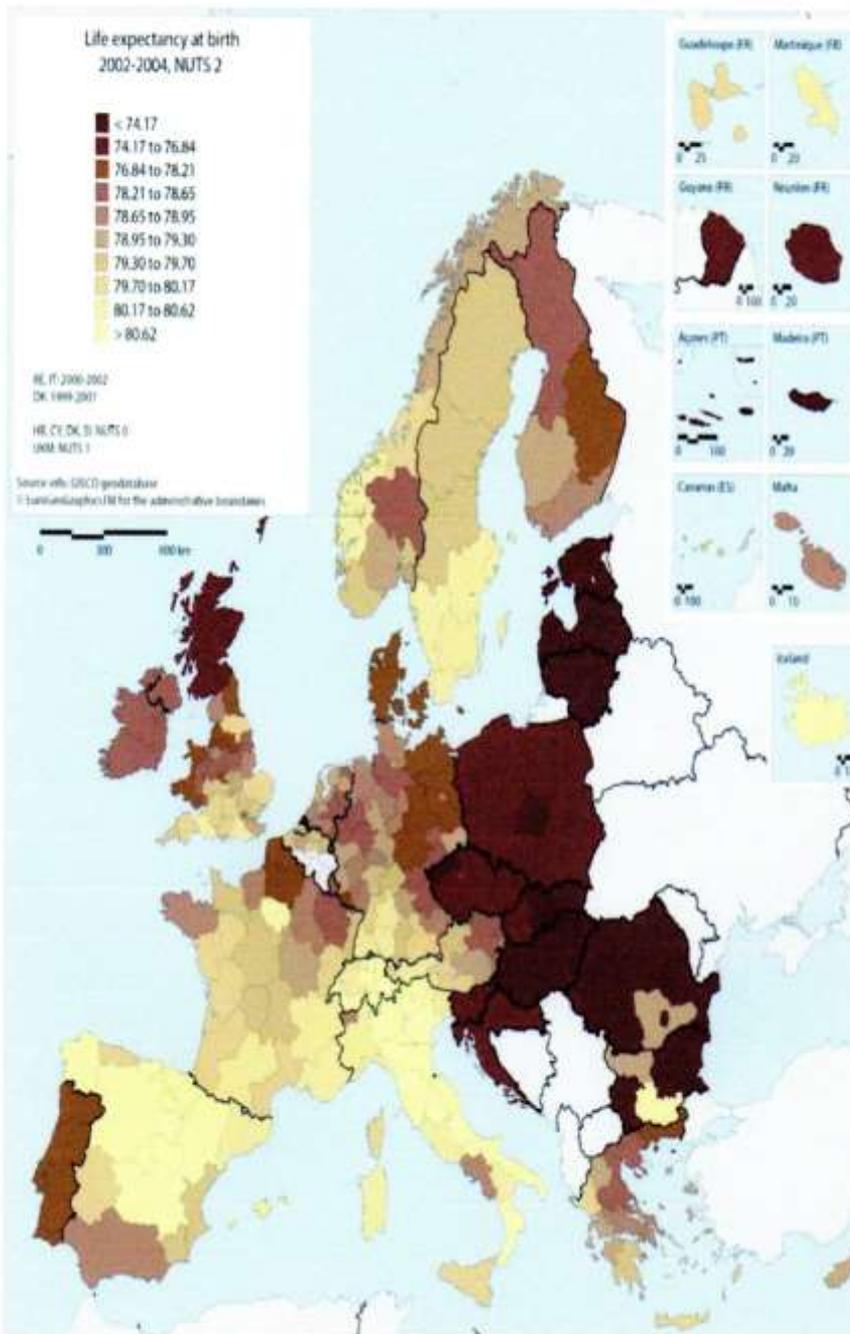
Country	Percent
Japan	21.6
Italy	20.0
Germany	20.0
Greece	19.1
Sweden	18.3
Spain	17.9
Austria	17.7
Bulgaria	17.6
Estonia	17.6
Belgium	17.5
Portugal	17.4
Croatia	17.0
Latvia	16.9
Georgia	16.6
Finland	16.6
France	16.3
Slovenia	16.3
Ukraine	16.1
United Kingdom	16.0
Switzerland	16.0
Lithuania	16.0
Denmark	15.7
Hungary	15.6
Serbia	15.2
Norway	15.0

Figure 6-8.
People Aged 65 and Over Living Alone in Ten European Nations by Sex: 2001
(In percent)

Male   Female

Nation	Male	Female
Denmark	26.4	55.1
Slovakia	18.1	49.1
Germany	16.6	47.4
Czech Republic	18.3	44.1
Netherlands	17.7	43.9
Switzerland	16.7	41.7
Italy	13.6	36.0
Poland	13.8	33.7
Latvia	12.7	28.5
Portugal	10.1	25.3
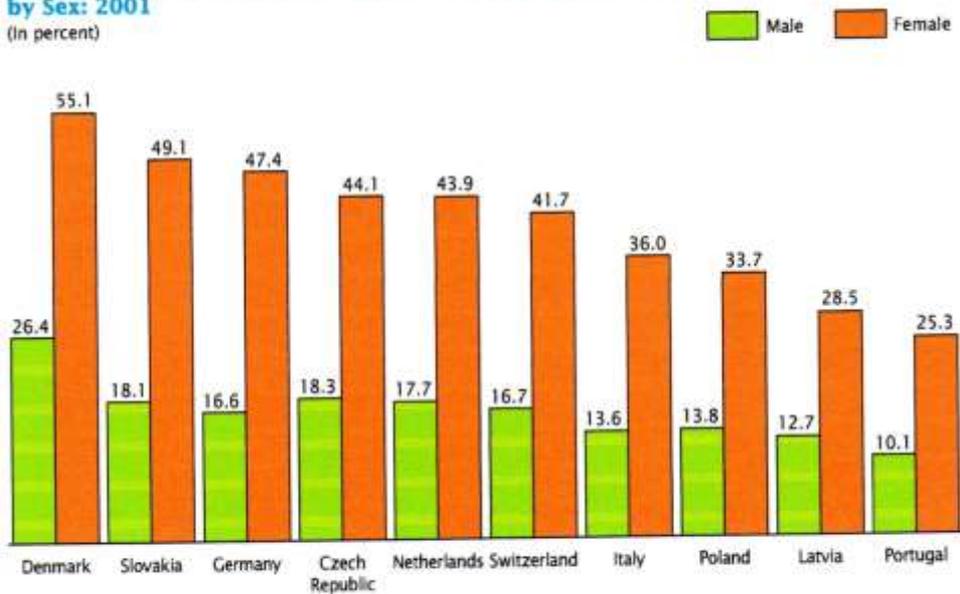
Figure 9-10.
Years of Life Expectancy After Retirement in Ten Countries by Sex: 2004 Versus 1970
(Based on average actual age at retirement)

2004
1970

Male

Country	2004	1970
France	21.4	10.8
Italy	20.6	13.1
Spain	19.6	10.7
Finland	19.6	10.8
Australia	18.9	10.9
Germany	18.9	10.5
Sweden	17.8	11.9
United States	17.1	11.0
Portugal	15.2	7.1
Japan	14.8	8.5

Female

Country	2004	1970
France	26.2	13.4
Italy	23.9	18.4
Spain	22.6	11.1
Finland	23.8	16.4
Australia	21.2	12.4
Germany	23.8	13.9
Sweden	22.9	14.9
United States	21.0	14.7
Portugal	18.6	9.4
Japan	22.0	13.1

Source: Organisation for Economic Co-Operation and Development (OECD), 2006b.

Figure 6. Premature mortality: percentage of deaths in the age group 20-64 years - 2002

men

%	Country
55,8%	Russia
45,9%	Latvia
44,7%	Estonia
43,3%	Lithuania
38,4%	Hungary
38,1%	Poland
37,9%	Slovakia
36,7%	EU10
35,5%	Romania
34,5%	Slovenia
32,8%	Czech Rep.
30,0%	Bulgaria
29,4%	Finland
28,6%	Luxembourg
27,1%	Germany
26,6%	Austria
26,4%	France
25,2%	Portugal
25,0%	Ireland
23,8%	Belgium
23,7%	Denmark
23,6%	EU15
22,6%	Netherlands
22,5%	Spain
21,0%	UK
20,8%	Greece
19,7%	Italy
17,0%	Sweden

women

Country	%
Russia	23,7%
Romania	18,8%
Hungary	18,6%
Latvia	18,3%
Lithuania	18,2%
Estonia	17,6%
EU10	17,3%
Poland	17,3%
Slovakia	16,9%
Bulgaria	15,2%
Slovenia	14,7%
Czech Rep.	14,6%
Denmark	14,3%
Netherlands	14,1%
Ireland	14,0%
Luxembourg	13,9%
Portugal	12,4%
Belgium	12,4%
Finland	12,1%
France	12,0%
UK	11,8%
Germany	11,4%
EU15	11,4%
Austria	11,1%
Italy	10,1%
Greece	10,0%
Spain	9,9%
Sweden	9,6%

Belgium 1997, Denmark 2001

Source: WHO mortality statistics (http://www.who.int/whosis/mort/download/en/index.html)

Portugal em quebra

Em 2100, a população portuguesa poderá ser de 6,7 milhões de habitantes segundo o cenário médio das projecções da ONU

10.676.000
2010

Por sexo

5,17   5,50   3,34   3,41

População total	10,67	6,75
Média etária	41 anos	47,7 anos
	2010	2100

milhões de habitantes

Idades

2010	
Mais de 65 anos **17,9%**	0-14 **15,1%**
	15-24 **10,9%**
25-64 **43,7%**	

2100	
Mais de 65 **30,6%**	0-14 **15,4%**
	15-24 **10,3%**
25-64 **56,1%**	

6.754.000
2100

Nascimentos

96 mil — 2010
70 mil — 2100

N.º de crianças por mulher

1,36 — 2010
1,99 — 2100
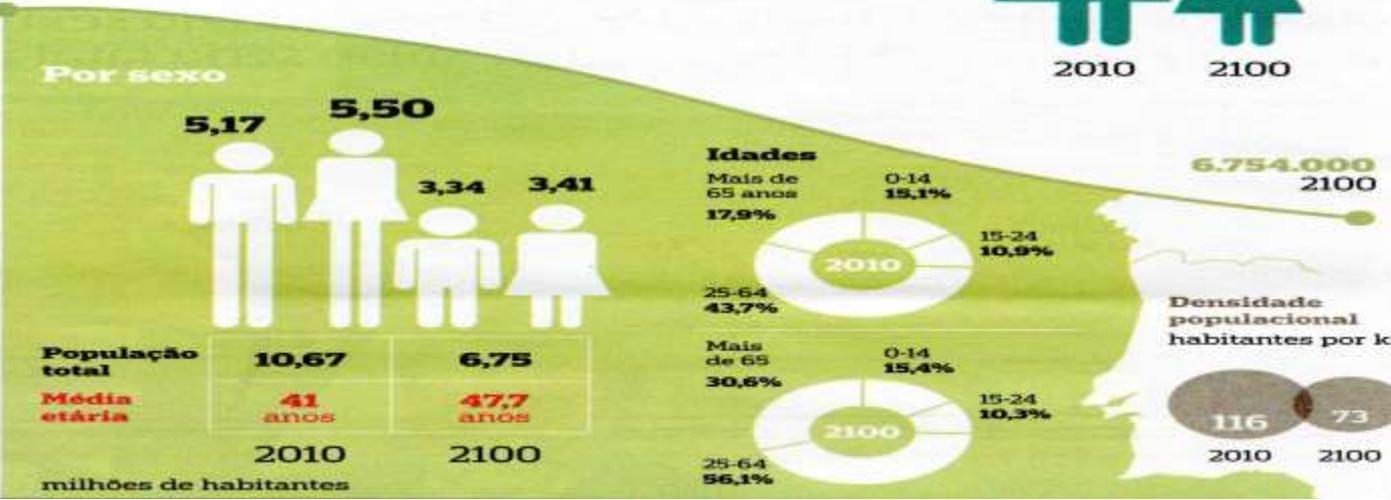
Esperança de vida

Média de idades por sexo

Homens		Mulheres	
76,8	85,1	82,8	90,4
2010	2100	2010	2100

Mortes

108 mil — 2010
97 mil — 2100

Densidade populacional habitantes por km²

116 — 2010
73 — 2100

Portugal

Legenda:
- median
- PI 80
- PI 95
- +/- 0.5 child
- observed TFR

Eixo Y: TFR
Eixo X: Year (1950–2100)

ESTATÍSTICAS

MONICA
Monograph and
Multimedia Sourcebook
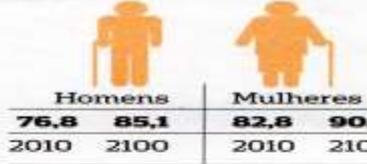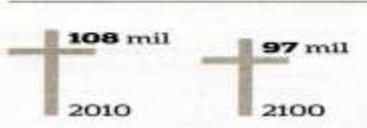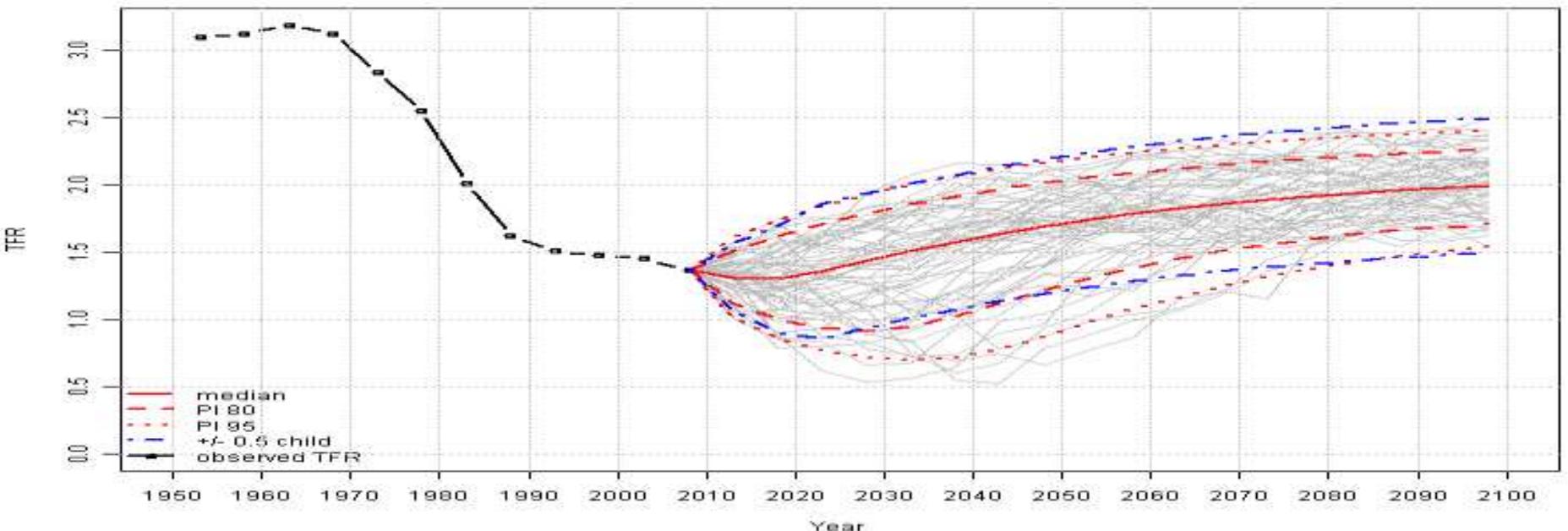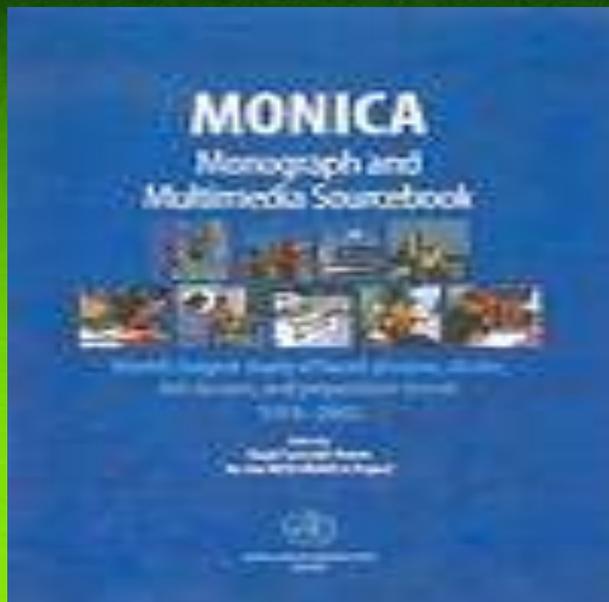
ELEMENTOS ESTATÍSTICOS
INFORMAÇÃO GERAL
SAÚDE / 2008

Direcção-Geral da Saúde
www.dgs.pt

Ministério da Saúde

DIRECÇÃO-GERAL DA SAÚDE

Gráfico nº2
Pirâmide etária, Portugal, 2008

■ Homens   ■ Mulheres
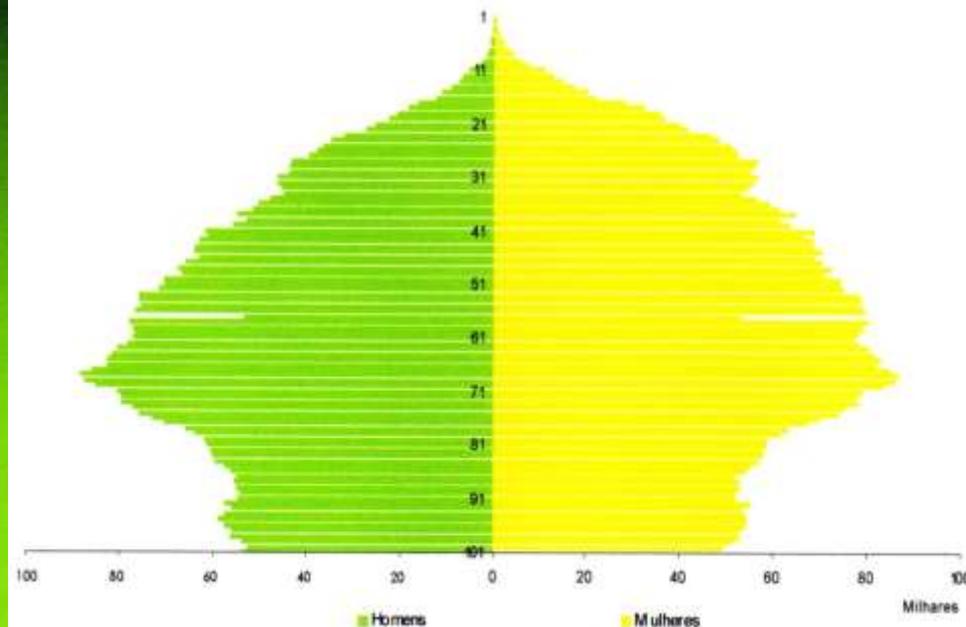
Gráfico nº 4
Esperança de vida à nascença, por sexos, Portugal, 1990-2006/2008

HM   H   M

Fonte: INE, Estatísticas Demográficas
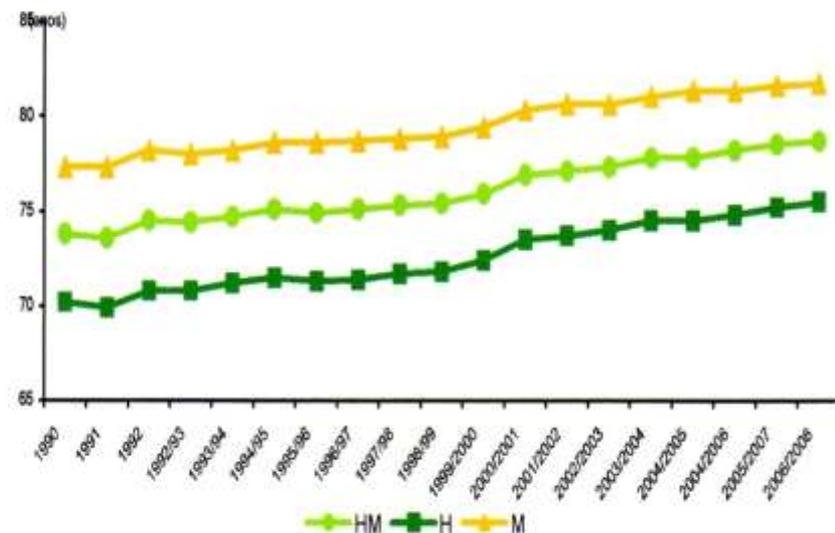
Cartograma nº 1
Percentagem da população com 65 e mais anos, por sub-regiões de saúde e RA, 2008

AÇORES

MADEIRA

13,0 - 16,1
16,2 - 19,3
19,4 - 22,4
22,5 - 25,5
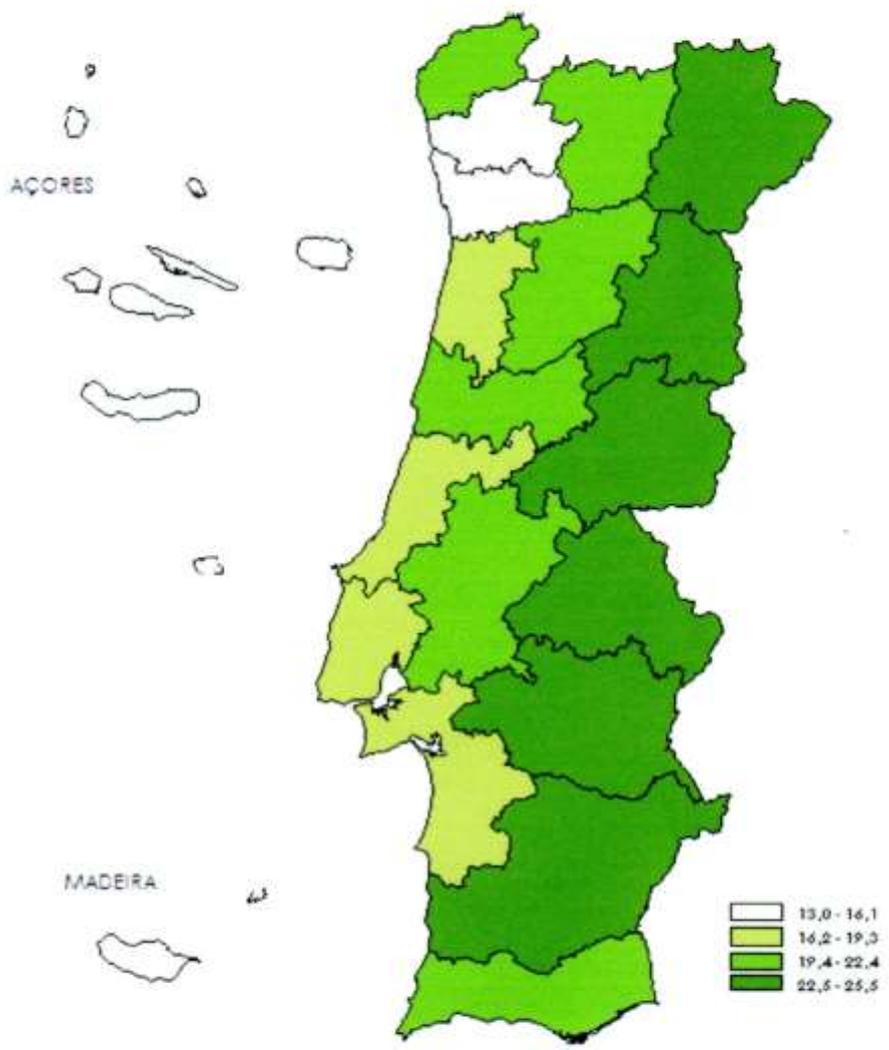
Fonte: INE, Estimativas da População Residente

Cartograma nº 2
Crescimento natural da população, por sub-regiões de saúde e RA, 2008

AÇORES

MADEIRA

Positivo
Negativo

Fonte: INE, Estatísticas Demográficas

Cartograma nº 3
Taxa de fecundidade, por sub-regiões de saúde e RA, 2008

□	28,1 - 33,6
▨	33,7 - 39,1
▨	39,2 - 44,6
■	44,7 - 50,1
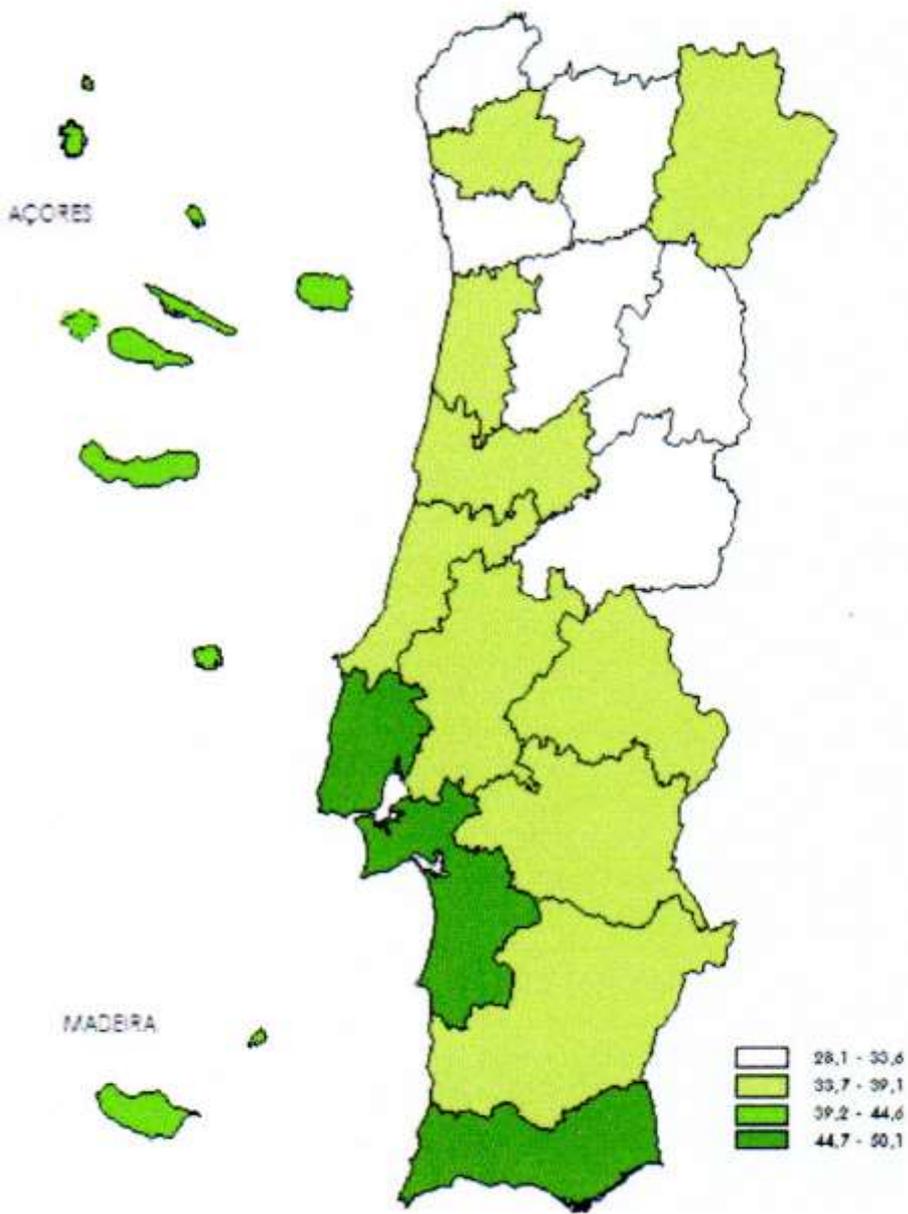
Fonte: INE

Cartograma nº 4
Taxa de mortalidade geral, por sub-regiões de saúde e RA, 2008

□	7,0 - 9,3
▨	9,4 - 11,6
▨	11,7 - 13,5
■	14,0 - 16,2
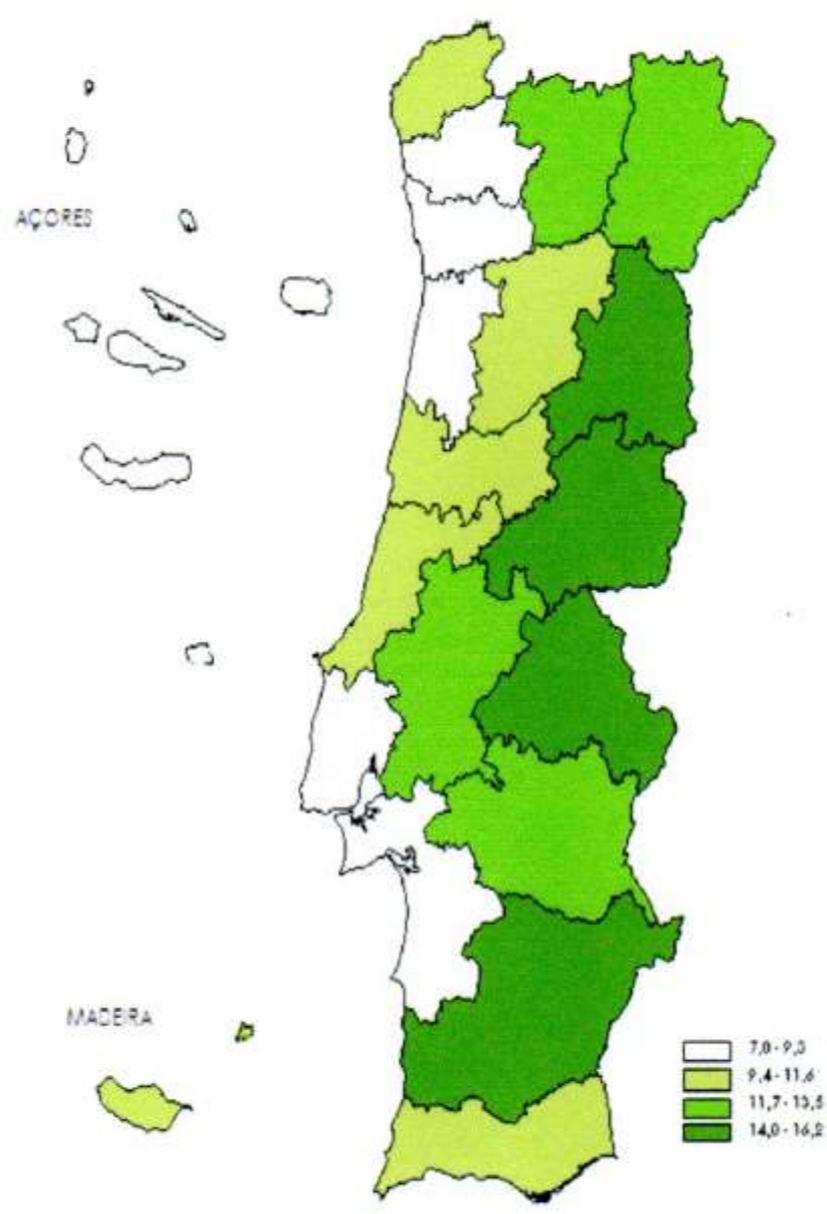
Fonte: INE

Gráfico nº 1
Evolução do índice de dependência total, índice de dependência dos jovens e índice de envelhecimento, Portugal, 1990-2008

(%)

Legend: Índ.dep.Total — Índ.dep.Jovens — Índ. Envelhecimento
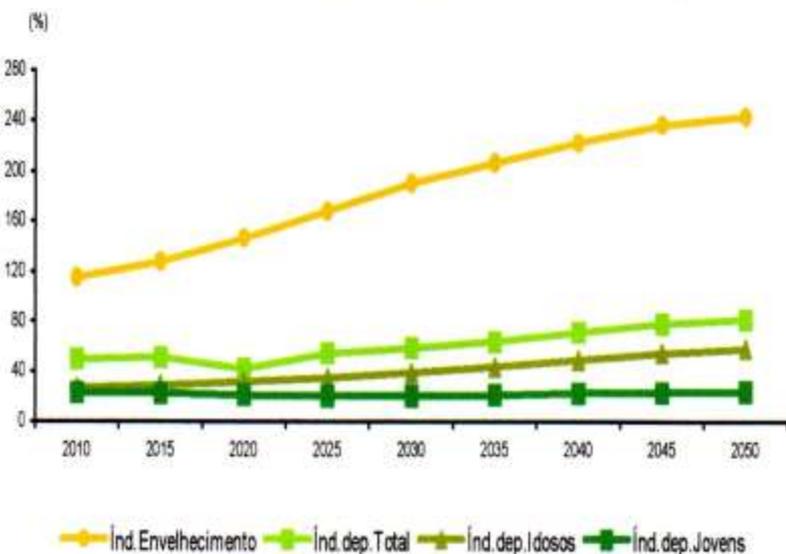
Fonte: INE, Estimativas da População Residente

COUNTRY	GDP ($US)	UR%	Year	%GDP Lost High	%GDP Lost Low	$GDP Lost High	$GDP Lost Low
Andorra	1,200,000,000	0.0	96	0.00	0.00	-	-
Aruba	1,400,000,000	0.5	94	0.46	0.32	6,405,000	4,515,525
Australia	332,000,000,000	8.5	96	7.78	5.48	25,821,300,000	18,204,016,500
Austria	197,000,000,000	6.2	96	5.67	4.00	11,175,810,000	7,878,946,050
Bahamas	4,600,000,000	15.0	95	13.73	9.68	658,800,000	464,454,000
Belgium	228,000,000,000	9.5	96	8.69	6.13	19,818,900,000	13,972,324,500
Bermuda	1,800,000,000	0.0	95	0.00	0.00	-	-
Brunei Darussalam	4,600,000,000	4.8	94	4.39	3.10	202,032,000	142,432,560
Canada	543,000,000,000	9.5	96	8.69	6.13	47,200,275,000	33,276,193,875
Cayman Islands	860,000,000	7.0	92	6.41	4.52	55,083,000	38,833,515
Cyprus	9,336,000,000	5.9	95	5.40	3.81	504,003,960	355,322,792
Denmark	146,000,000,000	8.2	96	7.50	5.29	10,954,380,000	7,722,837,900
Faeroe Islands	800,000,000	11.0	96	10.07	7.10	80,520,000	56,766,600
Finland	96,000,000,000	17.1	96	15.65	11.03	15,333,570,000	10,810,168,850
France	1,330,000,000,000	11.6	96	10.61	7.48	141,156,200,000	99,522,171,000
French Polynesia	1,760,000,000	15.0	92	13.73	9.68	241,560,000	170,299,800
Germany	2,046,000,000,000	8.2	96	7.50	5.29	153,511,380,000	108,225,522,900
Greenland	892,000,000	10.5	95	9.61	6.77	85,699,900	60,417,725
Hong Kong	131,900,000,000	3.1	96	2.84	2.00	3,741,343,500	2,637,647,168
Iceland	5,300,000,000	5.0	96	4.58	3.23	242,475,000	170,944,875
Ireland	52,000,000,000	12.9	96	11.80	8.32	6,137,820,000	4,327,163,100
Israel	78,000,000,000	6.5	96	5.95	4.19	4,639,050,000	3,270,530,250
Italy	1,025,000,000,000	12.2	96	11.16	7.87	114,420,750,000	80,666,628,750
Japan	4,591,000,000,000	3.1	96	2.84	2.00	130,223,715,000	91,807,719,075
Kuwait	24,300,000,000	1.8	96	1.65	1.16	400,221,000	282,155,805
Liechtenstein	713,000,000	1.1	96	1.01	0.71	7,176,345	5,059,323
Luxembourg	10,000,000,000	3.0	95	2.75	1.94	274,500,000	193,522,500
Macau	6,800,000,000	2.0	92	1.83	1.29	124,440,000	87,730,200
Monaco	800,000,000	3.1	94	2.84	2.00	22,692,000	15,997,860
Netherlands	330,000,000,000	6.5	96	5.95	4.19	19,626,750,000	13,836,858,750
Netherlands Antilles	2,040,000,000	13.4	93	12.26	8.64	250,124,400	176,337,702
New Zealand	51,000,000,000	6.3	96	5.76	4.06	2,939,895,000	2,072,626,975
Norway	110,000,000,000	4.9	96	4.48	3.16	4,931,850,000	3,476,954,250
Portugal	87,000,000,000	7.1	96	6.50	4.58	5,651,955,000	3,984,628,275
Qatar	11,700,000,000	7.3	—	6.68	4.71	781,601,500	550,958,556
Singapore	68,900,000,000	2.7	96	2.47	1.74	1,702,174,500	1,200,033,023
Spain	483,000,000,000	22.7	96	20.77	14.64	100,321,515,000	70,726,668,075
Sweden	196,000,000,000	9.2	96	8.42	5.93	16,499,280,000	11,631,992,400
Switzerland	260,000,000,000	5.3	96	4.85	3.42	12,608,700,000	8,889,133,500
United Arab Emirates	35,400,000,000	7.3	—	6.68	4.71	2,364,543,000	1,667,002,815
United Kingdom	1,017,000,000,000	8.7	96	7.96	5.61	80,958,285,000	57,075,590,925
United States	6,848,000,000,000	5.4	96	4.94	3.48	328,477,680,000	231,576,764,400
Virgin Islands (U.S)	1,200,000,000	6.2	94	5.67	4.00	68,076,000	47,993,580

Gráfico nº3
Projecção da evolução dos índices de envelhecimento e dependência, Portugal 2000-2050

(%)

Legend: Índ.Envelhecimento — Índ.dep.Total — Índ.dep.Idosos — Índ.dep.Jovens

Fonte: INE, Projecções da População Residente

Gráfico nº 12
Evolução do número de pensionistas com incapacidade permanente (1), em resultado de doenças profissionais contraídas, Continente, 1990-2008
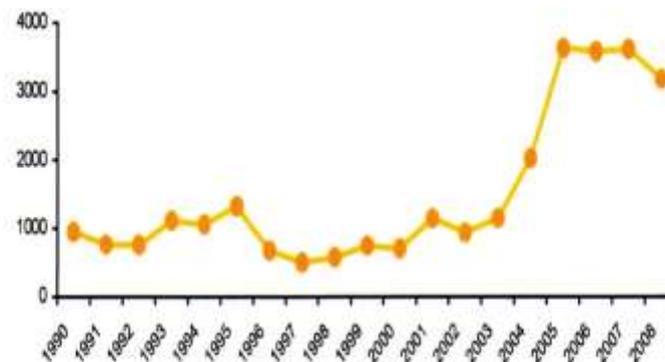
(1) Os pensionistas com incapacidade permanente incluem: pensionistas com incapacidade parcial permanente; pensionistas com incapacidade permanente absoluta para o trabalho habitual e pensionistas com incapacidade permanente e absoluta para qualquer trabalho.

Fonte: MSST, Instituto de Informática e Estatística da Solidariedade

2.1.3 DOENÇAS CRÓNICAS

População inquirida que tem ou já teve doença crónica, por tipo de doença, sexo, NUTS I, 2005/2006

(%)

Tipo de doença	Continente			R.A.Açores			R.A.Madeira		
	HM	H	M	HM	H	M	HM	H	M
Diabetes	6,5	5,9	7,1	6,7	5,2	8,2	4,6	3,4	5,7
Asma	5,5	4,8	6,2	5,8	5,2	6,4	4,4	4,7	4,1
Tensão arterial alta	20,0	16,4	23,4	16,3	12,0	20,5	13,1	9,1	16,7
Dor crónica	16,3	12,7	19,8	10,0	7,7	12,2	7,2	5,1	9,0
Doença reumática	16,3	11,4	20,9	12,9	8,4	17,3	6,0	3,6	8,2
Osteoporose	6,3	1,1	11,1	5,6	1,6	9,4	4,1	1,3	6,6
Glaucoma	0,7	0,5	0,9	0,7	0,4	0,9	0,4	0,3	0,4
Retinopatia	0,8	0,7	0,8	0,3	0,3	0,3	1,2	1,1	1,3
Tumor maligno/cancro	1,9	1,7	2,2	1,4	1,1	1,6	0,8	0,6	1,1
Pedra nos rins	4,9	4,8	5,1	3,5	2,8	4,1	0,9	1,2	0,7
Insuficiência renal	1,6	1,4	1,8	0,6	0,6	0,6	0,8	0,6	0,9
Ansiedade crónica	4,6	2,6	6,4	4,4	2,0	6,8	0,9	0,9	1,0
Ferida crónica	0,9	0,8	1,1	0,8	0,5	1,0	0,6	0,7	0,5
Enfisema, bronquite crónica	3,7	3,5	3,9	2,1	2,3	1,9	2,0	2,2	1,8
Acidente vascular cerebral	1,6	1,8	1,5	1,4	1,6	1,3	1,3	1,2	1,4
Obesidade	3,9	3,0	4,7	2,2	1,4	3,1	0,6	0,3	1,0
Depressão	8,3	4,0	12,5	5,4	2,6	8,2	4,1	2,2	5,7
Enfarte do miocárdio	1,3	1,9	0,8	1,0	1,2	0,9	0,9	1,1	0,7
Outra doença crónica	24,4	23,2	25,5	25,5	23,1	27,7	21,9	20,3	23,3

Fonte: INSA, Inquérito Nacional de Saúde, 2005/2006

Gráfico Nº 24
Distribuição percentual dos óbitos por algumas causas de morte, Portugal, 2008
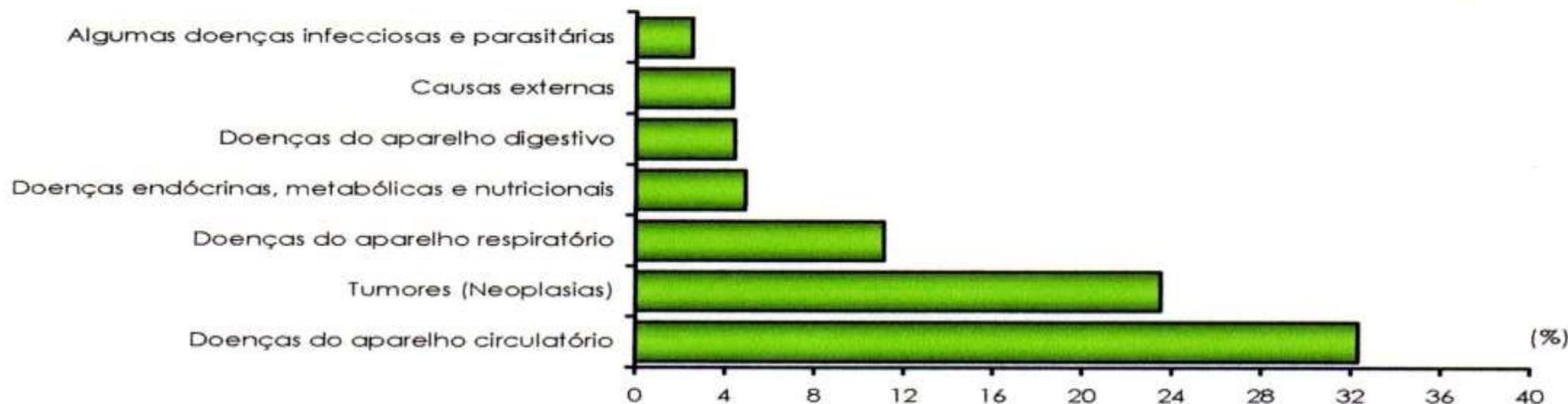
Fonte: INE, Estatísticas da Saúde; Estatísticas Demográficas

DGS/ DSEES

Prevalence of diagnosed diabetes in general population, all ages - standardised*
per 1000 individuals

Gráfico nº 8
Percentagem da população com excesso de peso, por grupos etários, Continente, 2005/2006
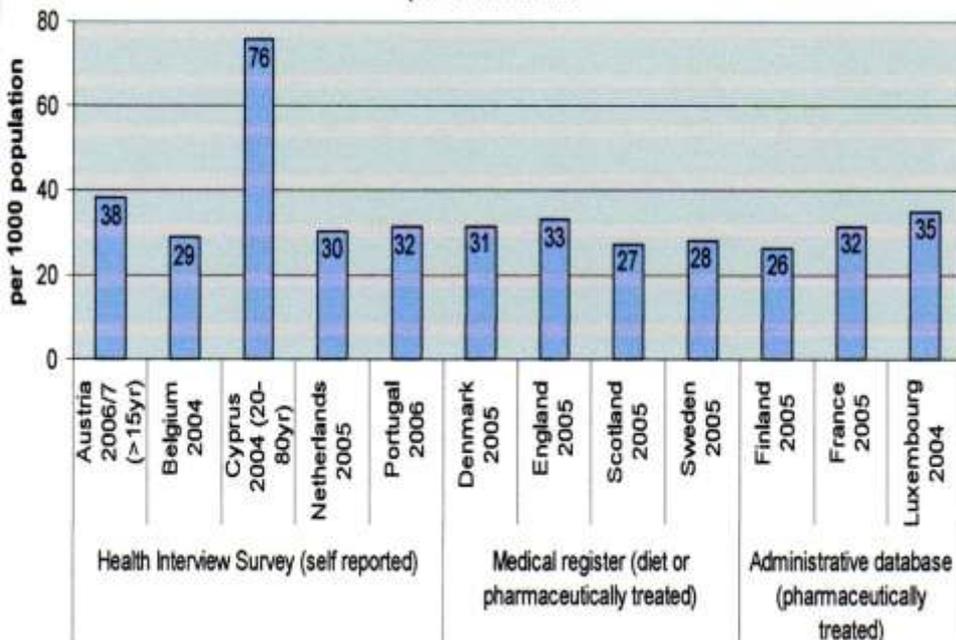
Figure 16a Standardised rates of mortality per 100.000 individuals in general population with primary or secondary cause of death of diabetes

Standardised* rates of diabetes mortality per 100.000 individuals in general population (primary and secundary cause of death)
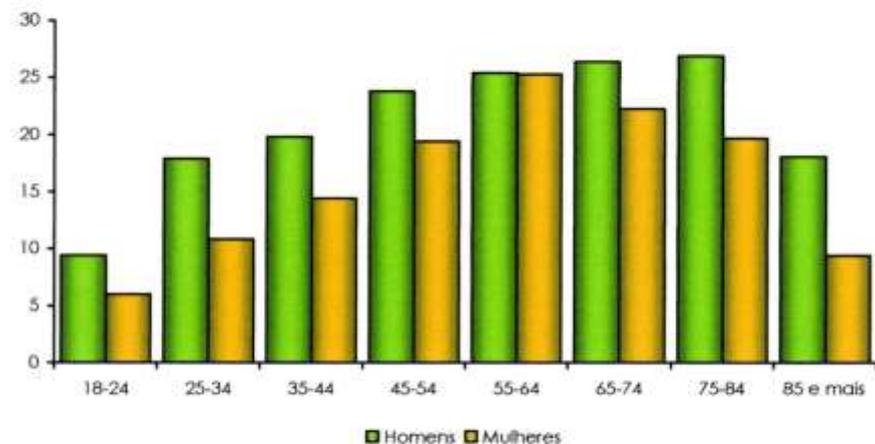
Gráfico nº 9
Percentagem da população com obesidade, por grupos etários, NUTS I, 2005/2006

Fonte: INSA, Inquérito Nacional de Saúde, 2005/2006
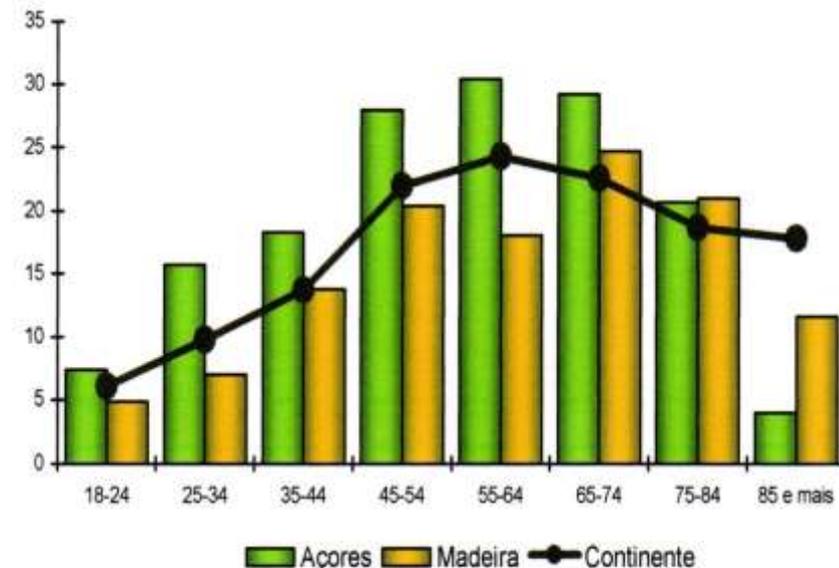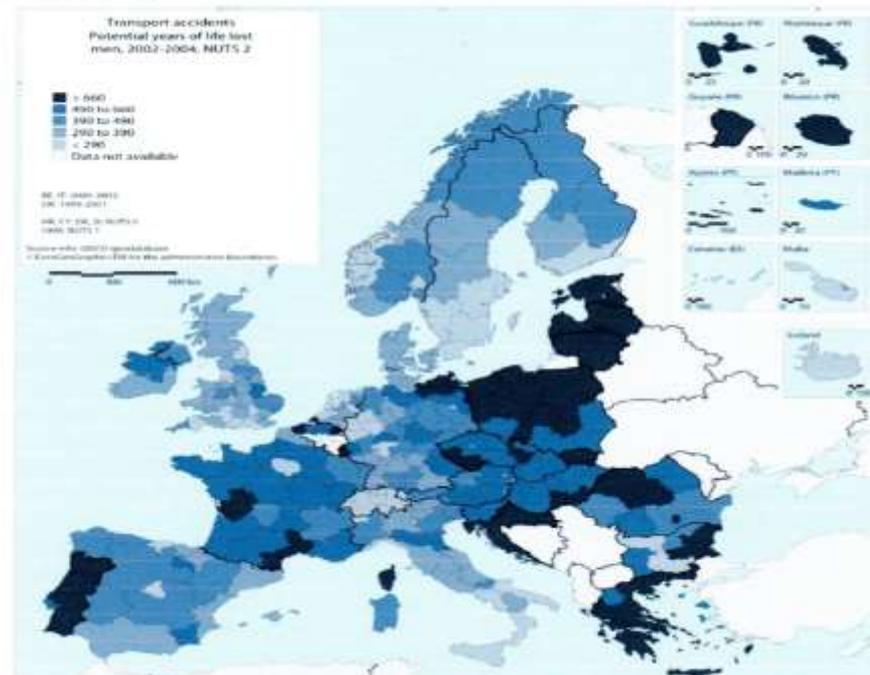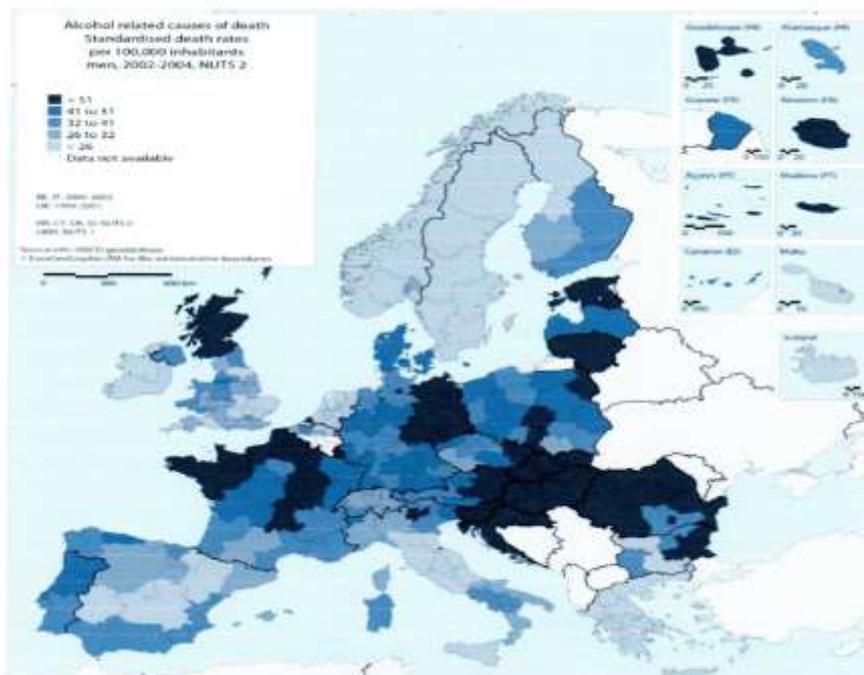
Gráfico nº 19
Índices de variação de casos novos e de óbitos relacionados com o uso ilícito de droga, 1990-2008, (1990=100)

Fonte: Instituto da Droga e da Toxicodependência
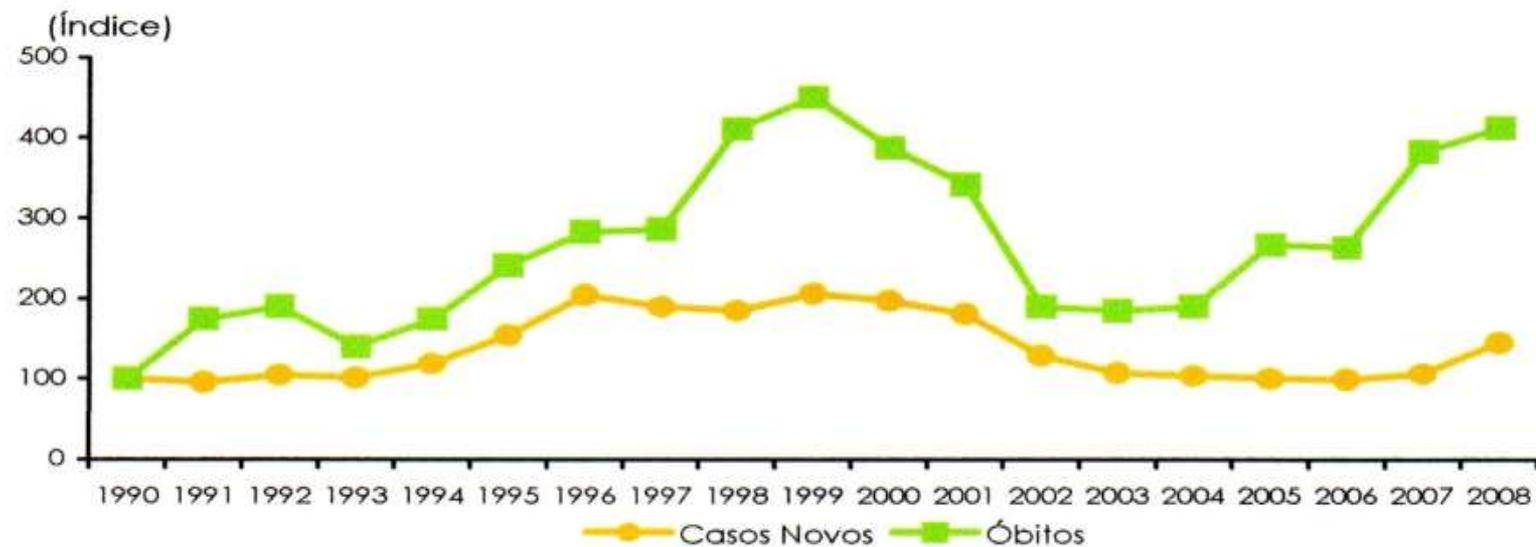
Fig. 2.5 *Burden of disease attributable to unipolar depressive disorder in selected countries in the WHO European Region (2004)*

Age-standardized DALY rate per 100 000 population

Source: WHO 2009.

Suicide

Suicide and intentional self-harm
Potential years of life lost
men, 2002-2004, NUTS 2
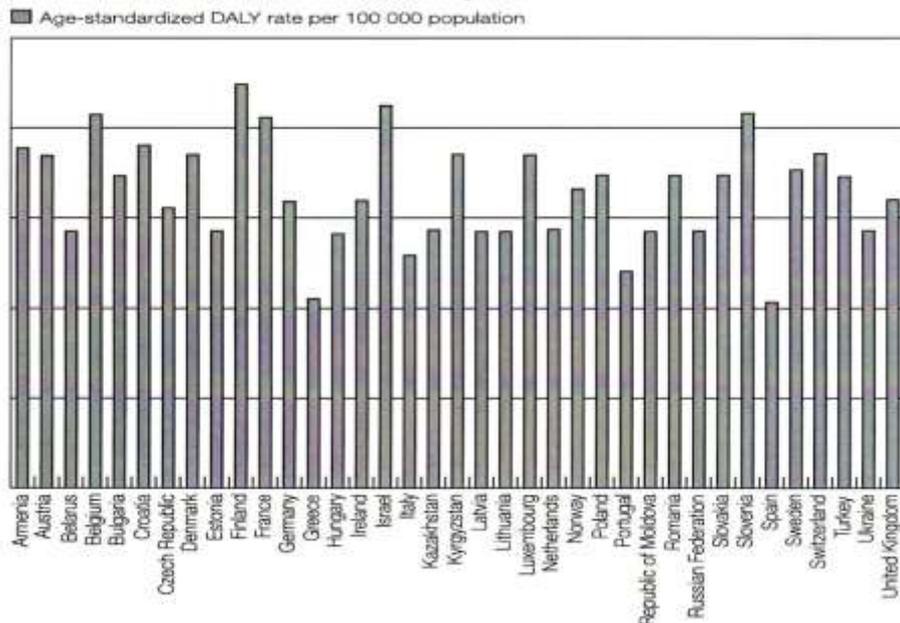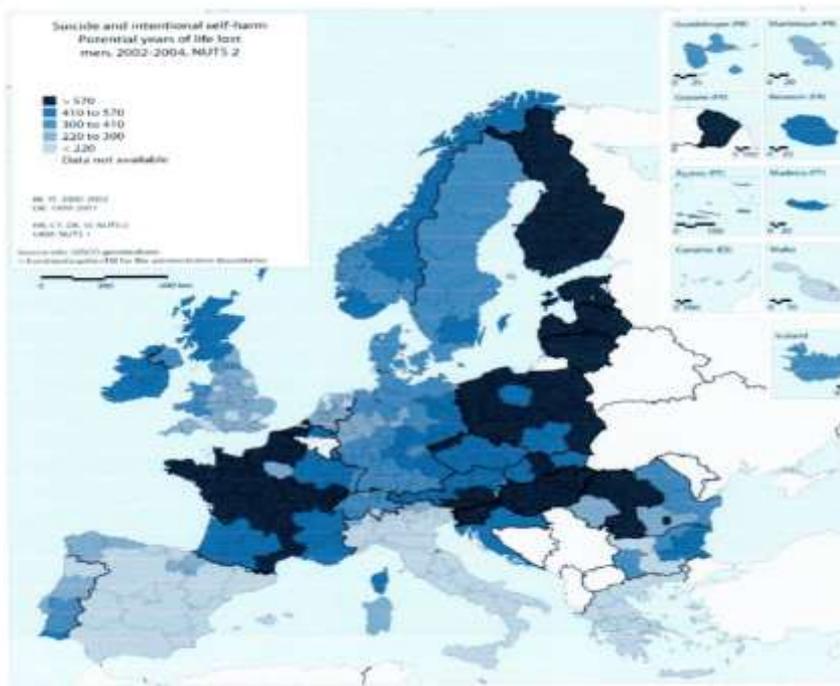
Table 1 The estimated number of people with dementia in Europe*

Country	Age group	Number of people with dementia (EURODEM)	As % of total population	Number of people with dementia (Ferri et al.)	As % of total population
Austria	30-94	104,428	1.27	94,441	1.15
Belgium	30-99	140,639	1.35	127,174	1.22
Bulgaria	30-99	87,797	1.13	76,556	0.99
Cyprus	30-99	6,725	0.9	6,054	0.81
Czech Republic	30-99	105,553	1.03	93,973	0.92
Denmark	30-99	68,430	1.26	62,318	1.15
Estonia (2004)	30-99	15,065	1.12	12,955	0.96
Finland	30-99	65,362	1.25	59,360	1.13
France	30-99	847,808	1.36	760,715	1.22
Germany	30-94	1,118,429	1.36	1,010,245	1.22
Greece	30-99	135,566	1.22	123,700	1.12
Hungary	30-89	100,567	1	88,070	0.87
Ireland	30-94	35,381	0.86	31,940	0.78
Italy	30-99	905,713	1.55	820,462	1.4
Latvia	30-99	25,969	1.13	22,509	0.98
Lithuania	30-99	35,298	1.03	30,169	0.88
Luxembourg	30-94	4,857	1.07	4,370	0.96
Malta	30-89	3,427	0.85	3,148	0.78
Netherlands	30-99	183,485	1.13	165,585	1.02
Poland	30-99	350,511	0.92	300,447	0.79
Portugal	30-94	129,916	1.23	119,308	1.13
Romania	30-99	200,893	0.93	172,130	0.79
Slovenia	30-99	21,788	1.09	19,302	0.97
Slovakia	30-99	44,813	0.83	38,232	0.71
Spain	30-99	583,208	1.36	533,388	1.24
Sweden	30-99	138,641	1.54	128,220	1.42
UK (2004)	30-89	660,573	1.11	621,717	1.04
EU27 TOTAL		**6,120,842**	**1.25**	**5,526,488**	**1.13**
Iceland	30-99	2,845	0.97	2,584	0.88
Norway	30-99	61,077	1.33	56,227	1.22
Switzerland	30-94	97,068	1.31	88,900	1.2
Turkey	30-74	129,715	0.18	78,546	0.11
other countries TOTAL		290,705		226,257	
GRAND TOTAL		**6,411,547**		**5,752,745**	

*The figures in this table are from 2005 unless stated otherwise
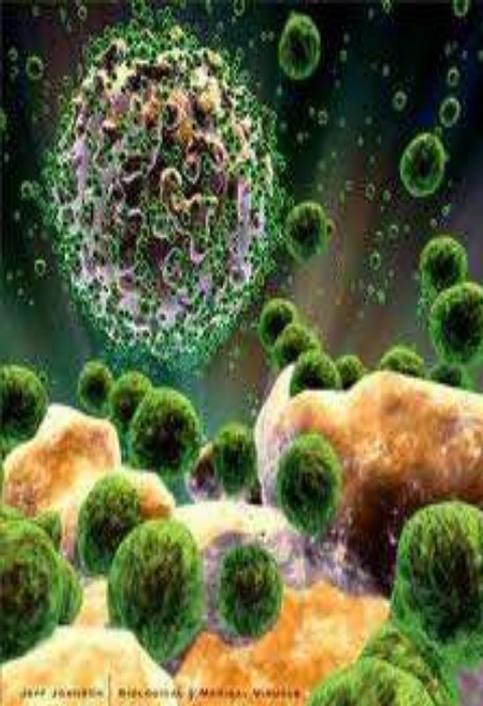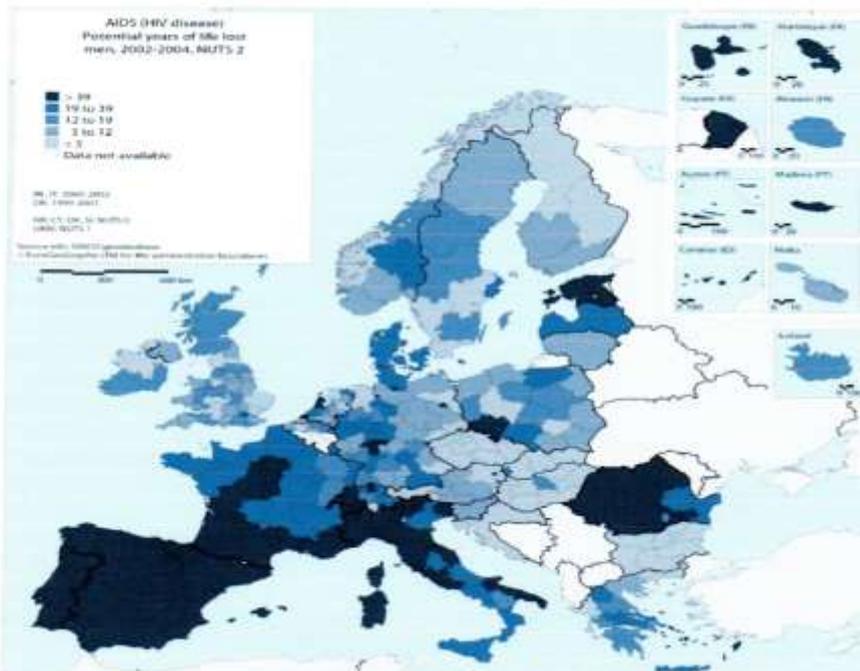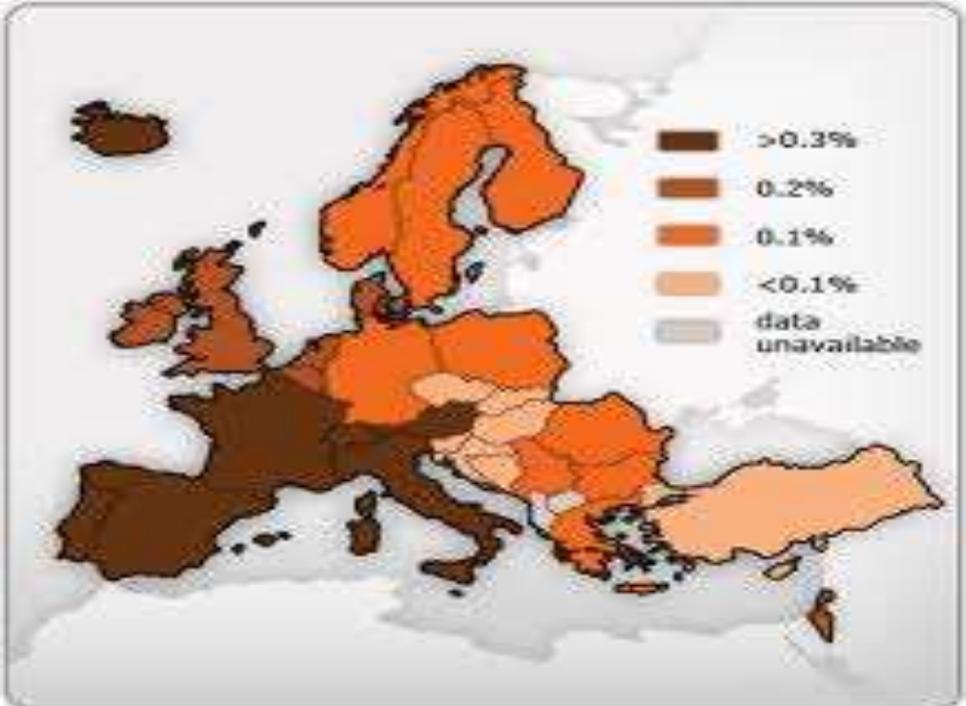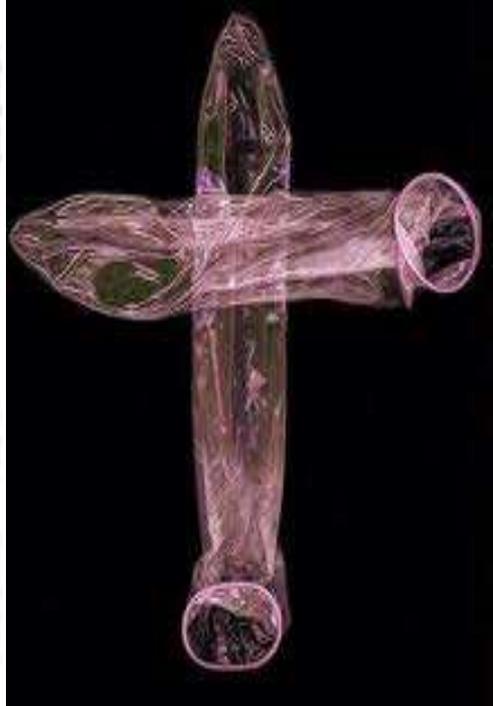
Fig. 4. Total health spending as a percent of GDP in current EU countries, 1997, 2001 and 2007
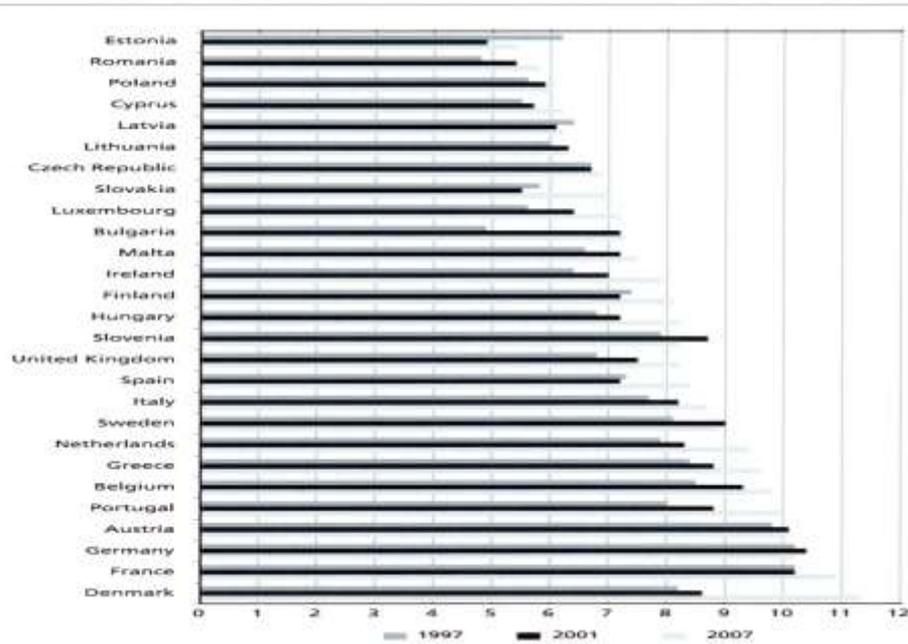
Source: WHO 2009 (81).

Fig. 5. Sources of health finance by country, 2007
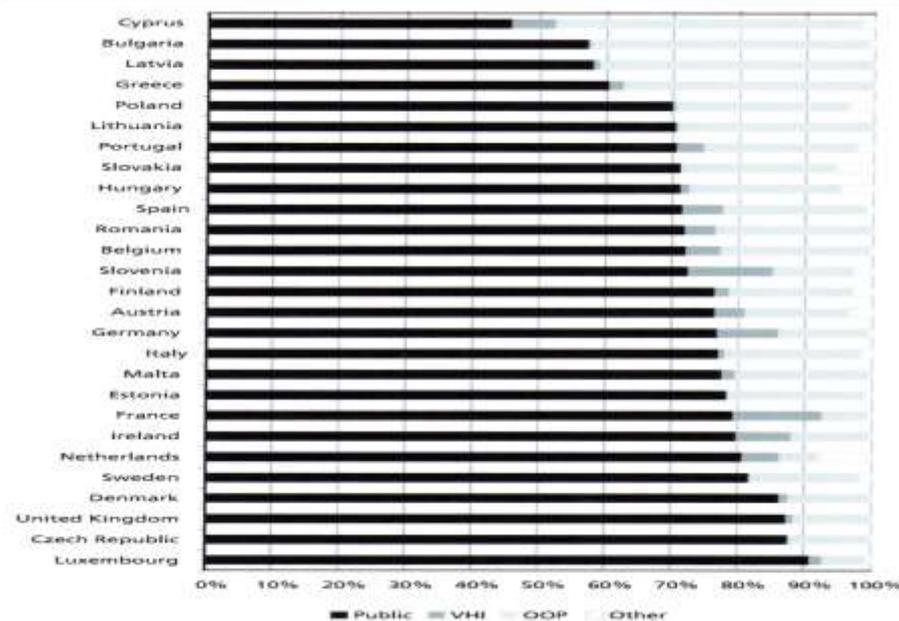
Source: WHO 2009 (81).
Notes: VHI: Voluntary health insurance; OOP: out-of-pocket (payment).

Gráfico nº 69
Despesas totais e públicas em saúde (percentagem) em relação ao PIB, na UE, 2008
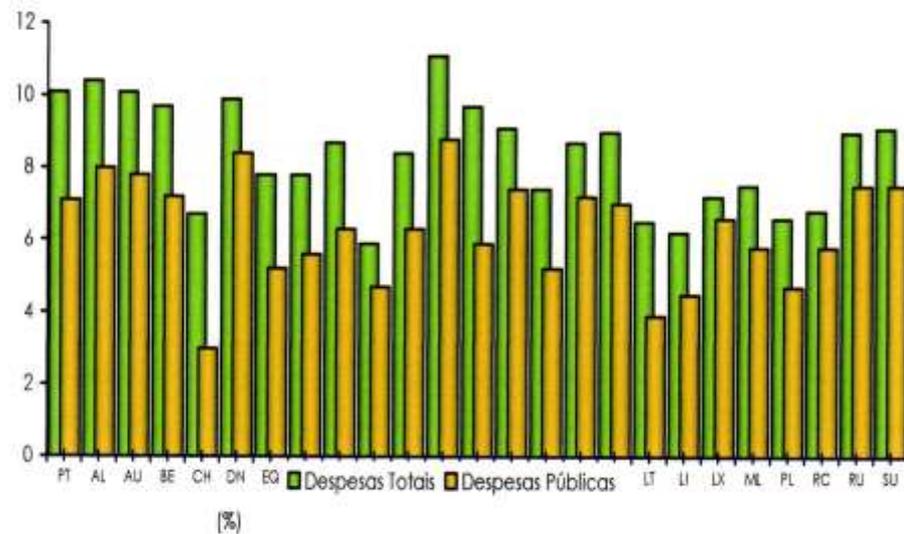
(%)

Nota: os dados do gráfico referem-se ao último ano disponível.

Fonte: OECD, Health Data 2008 (November 2010)

Gráfico nº 70
Despesas totais e públicas em saúde per capita (Dólares PPP), na UE, 2008
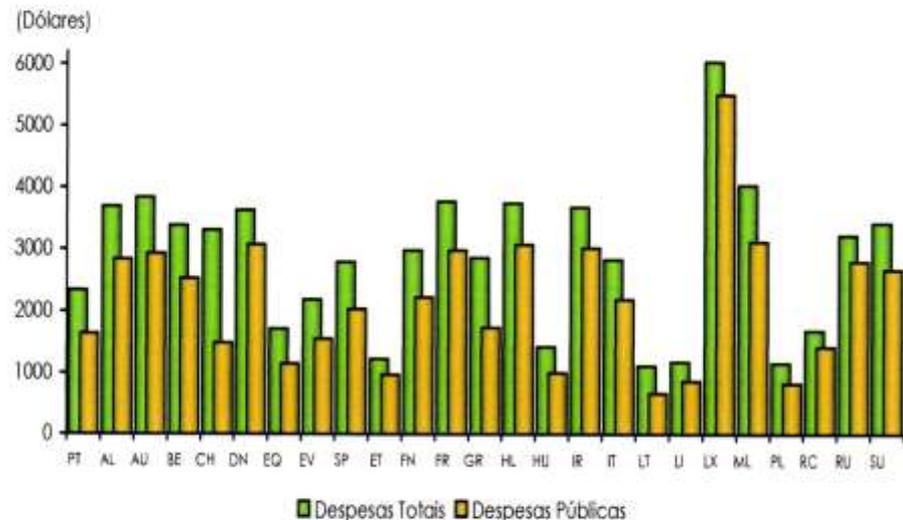
Nota: os dados do gráfico referem-se ao último ano disponível.

Fonte: OECD, Health Data 2008 (November 2010)

Figure 10-4.

Net Replacement Rate in Mandatory Pension Programs for Men in Selected Countries: 2005
(Percent)

Country	Rate
Australia	52
Austria	93
Belgium	63
Canada	57
Czech Republic	58
Finland	79
France	69
Germany	72
Greece	100
Hungary	90
Ireland	37
Italy	89
Japan	59
Mexico	45
Netherlands	84
New Zealand	40
Norway	65
Poland	70
Portugal	80
South Korea	44
Spain	88
Sweden	68
Switzerland	67
Turkey	103
United Kingdom	48
United States	51
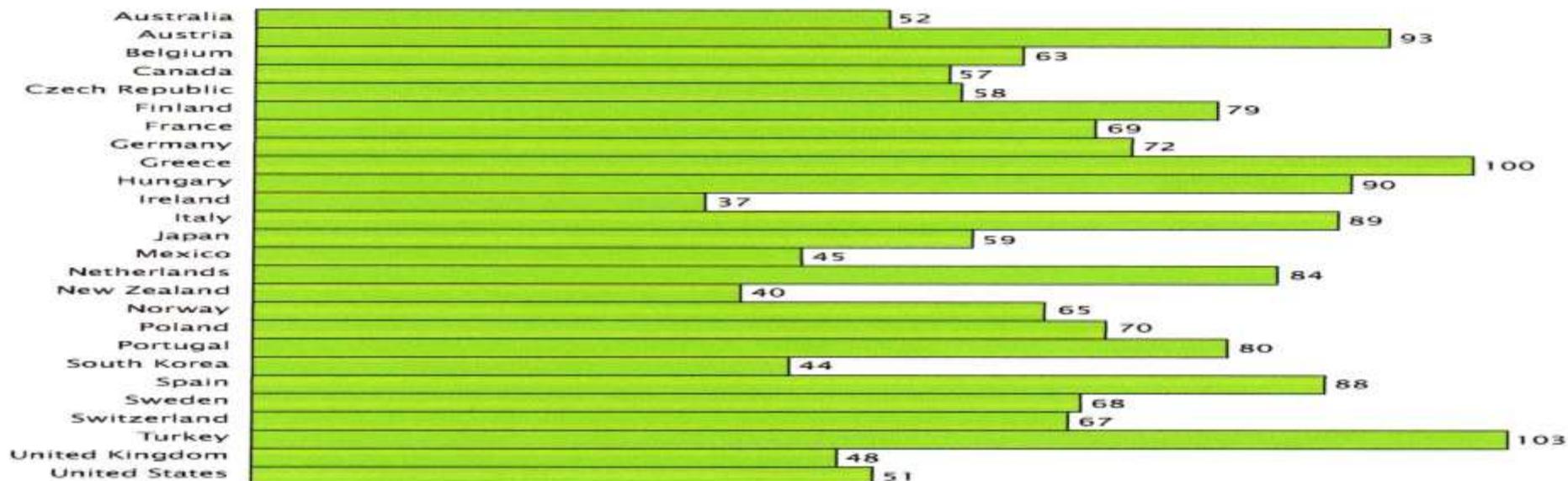
Note: Data refer to average individual pension entitlements as a percentage of preretirement earnings (net of taxes and contributions).
Source: Organisation for Economic Co-Operation and Development (OECD), 2005.

Graph 1: Sustainability gap* of the HCS by country, percent of GDP

Germany, Hungary, Austria, Czech Rep., France, Italy, Sweden, Belgium, Portugal, Finland, USA, Denmark, UK, Nederlands, Spain, Greece, Poland, Lux, Ireland, Slovakia
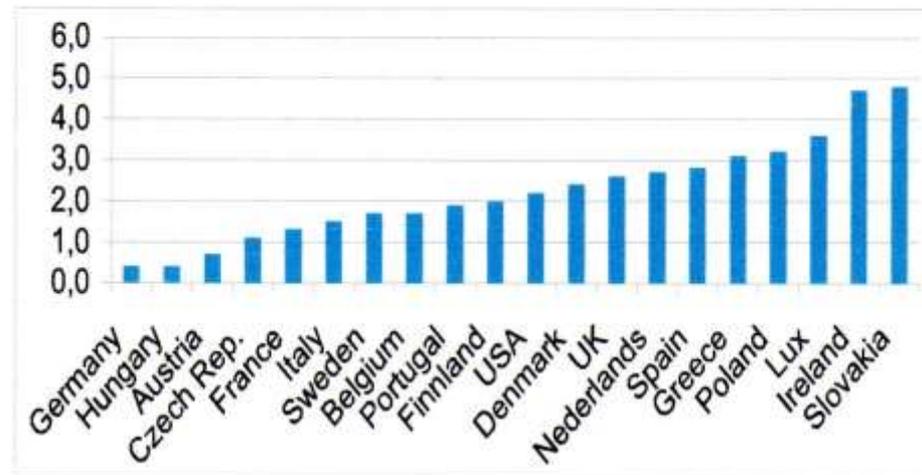
*Difference between average real growth rate of per capita total health care spending and real per capita GDP growth rate 2000-2008; Own calculations.
Source: OECD, Eurostat.

Table 1. OECD countries ranked by level of age-standardized mortality from causes amenable to health care, 1997–98 and 2002–03

Country	Amenable mortality (SDR, ages 0–74, per 100 000)		Rank in 1997–98	Rank in 2002–03	Change in rank
	1997–98	2002–03			
France	75.62	64.79	1	1	–
Spain	84.26	73.83	2	2	–
Sweden	88.44	82.09	3	5	-2
Italy	88.77	74.00	4	3	+1
Netherlands	96.89	81.86	5	4	+1
Greece	97.27	84.31	6	6	–
Germany	106.18	90.13	7	8	-1
Austria	108.92	84.48	8	7	+1
Denmark	113.01	100.84	9	10	-1
United States	114.74	109.65	10	14	-4
Finland	116.22	93.34	11	9	+2
Portugal	128.39	104.31	12	13	-1
United Kingdom	129.96	102.81	13	11	+2
Ireland	134.36	103.42	14	12	+2

Source: adapted from Nolte and McKee (33).

Notes: Amenable mortality is deaths before age 75 that are potentially preventable with timely and appropriate medical care; SDR: standardized death rate; Denmark 2000–01; Sweden 2001–02; United States 2002.

Cartograma nº 14
Índice habitante/médico, por sub-regiões de Saúde e RA, 2008

AÇORES

MADEIRA

112 - 239,5
239,6 - 367
367,1 - 494,5
494,6 - 622
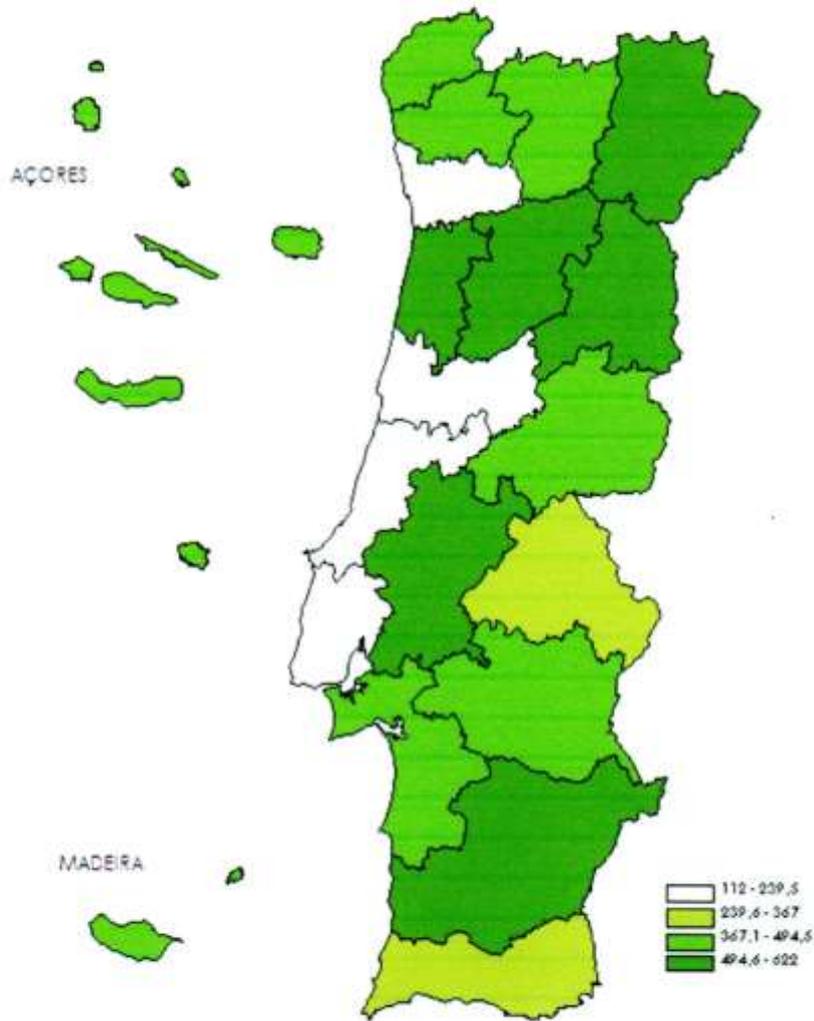
Gráfico nº 55
Distribuição dos médicos (2) por grupos etários e sexos, Portugal, 2008

HOMENS   MULHERES   Milhares
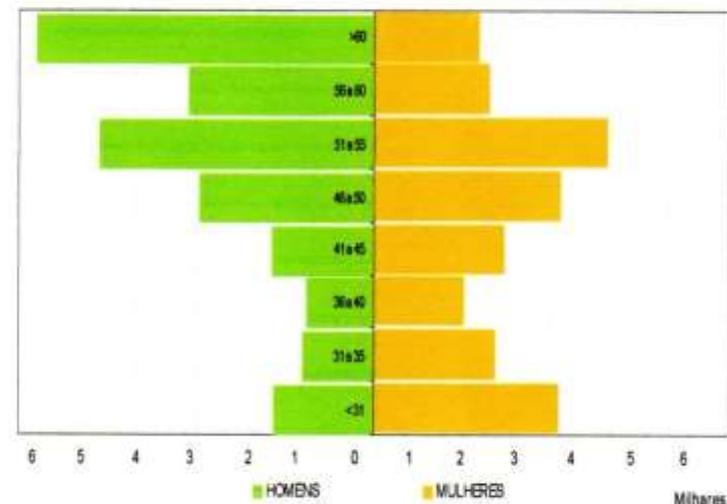
As RESPOSTAS da INOVAÇÃO CIENTÍFICA

NOVO PENSAMENTO

- " ... *os três pilares fundamentais onde deve assentar o pensamento científico: Criatividade, Veracidade, e Sentido da Dignidade Humana ... " (sic.)* (Jacob Bronowski)
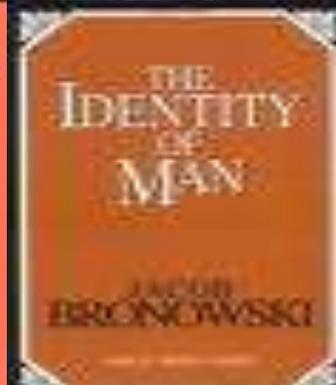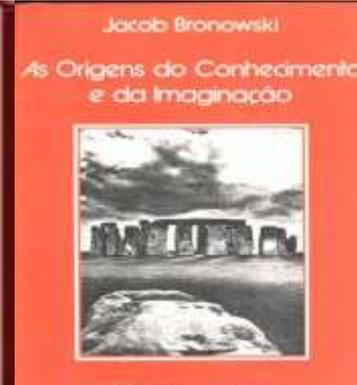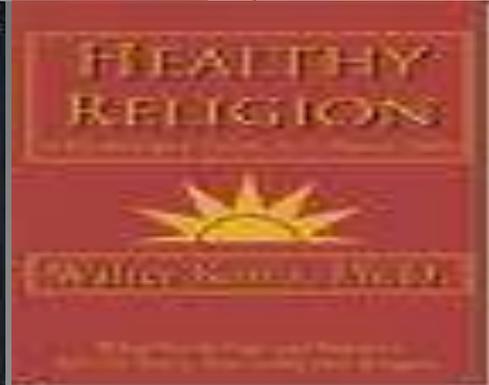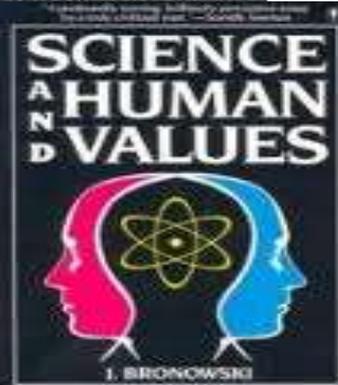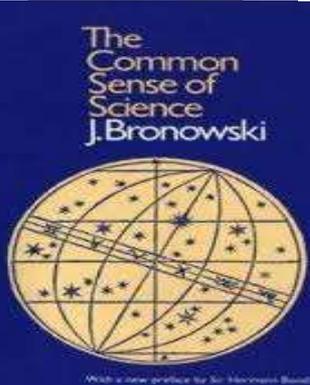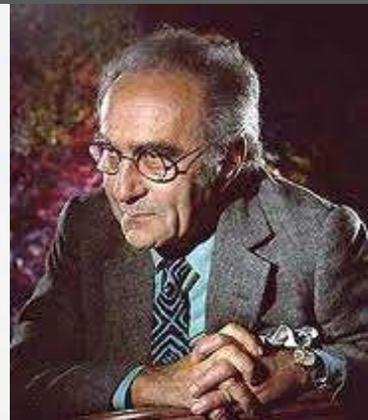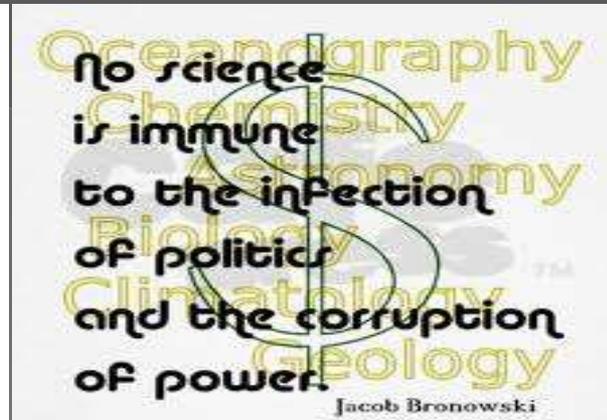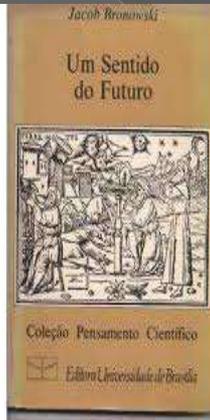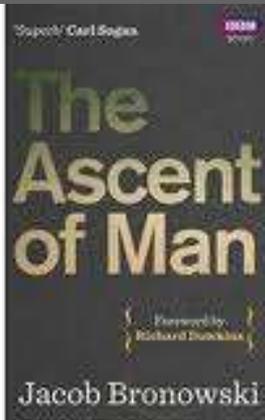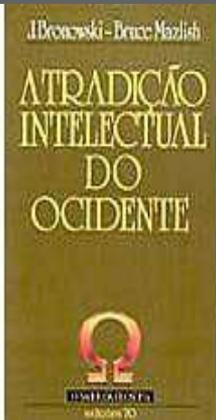
Virtual Medical Imaging

Alternative-Doctor.com

virtualmedical centre

Virtual Medical School

2nd Edition
VIRTUAL MEDICINE
By Prof Keith Scott-Mumby
MB, CHB, MD, PHD

The Birth of Energy Medicine in the West
Ancient Healing Arts Combine with
Western Scientific Technology to
Create a Medical Revolution

IMHOTEP VIRTUAL MEDICAL SCHOOL

The Future of Medicine Begins Here...

INTERACTIVE SOFTWARE
Virtual Medical Office

vhl
virtual health library

Surveillance Sans Frontières: Internet-based Emerging Infectious Disease Intelligence and the HealthMap Project

John S. Brownstein; Clark C. Freifeld; Ben Y. Reis; Kenneth D. Mandl
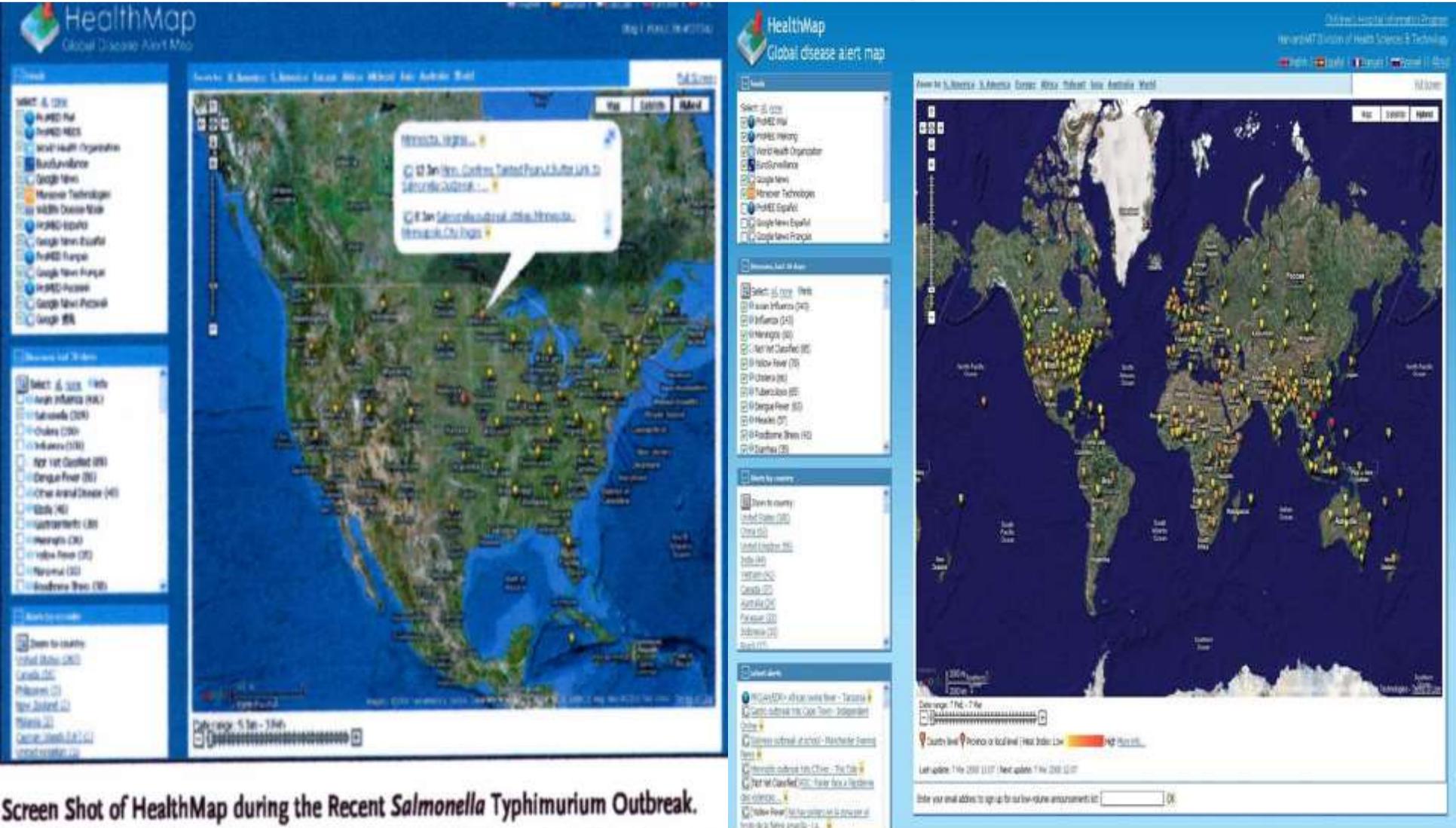
Screen Shot of HealthMap during the Recent *Salmonella* Typhimurium Outbreak.

HealthMap displays 319 articles about the outbreak that has affected 38 U.S. states.

IF SMALLPOX STRIKES PORTLAND...

BY CHRIS L. BARRETT, STEPHEN G. EUBANK AND JAMES P. SMITH

"EPISIMS" UNLEASHES VIRTUAL PLAGUES IN REAL CITIES TO SEE HOW SOCIAL NETWORKS SPREAD DISEASE; THAT KNOWLEDGE MIGHT HELP STOP EPIDEMICS

CREATING THE EPISIMS

The original EpiSims model was based on Portland, Ore., but gathering sufficiently detailed information about 1.6 million real people and their activities would have been difficult and intrusive. A synthetic population, statistically indistinguishable from the real one, could nonetheless be constructed and given realistic daily lives using publicly available data.

SYNTHETIC HOUSEHOLDS

The U.S. Census Bureau provided demographic information, such as age, household composition and income, for the entire city as well as 5 percent of its complete records for smaller study areas of a few square blocks. Through a statistical technique called iterative proportional fitting, these two data sets were combined to create households and individuals with statistically correct demographics and geographic distribution.

HOUSEHOLD #2375

Age:	28	27	7	3
Income:	$37K	$28K	$0	$0
Status:	worker	worker	student	day care
Auto:			n/a	n/a

DAILY ACTIVITIES

ACTIVITIES

Most metropolitan planning offices conduct detailed traveler activity surveys for small population samples of a few thousand. These logs track the movements of each household member over the course of one or more days, noting the time of each activity. By matching the demographics of survey respondents to the entire synthetic population, realistic daily activities can be generated for every synthetic household member.

LOCATIONS

Setting the population in motion requires assigning locations to every household's activities. Land-use data for buildings, parking lots, parks and other places were associated with 180,000 locations in the model, providing estimates of the number of people performing various types of activities there. Activities were anchored for individuals' work or school locations, and then places were chosen for additional activities, such as grocery shopping or recreation, taking into account their distance and other measures of their appeal.

BUILDING SOCIAL NETWORKS

TYPICAL HOUSEHOLD'S CONTACTS

This diagram shows where the household members go and what they do all day but reveals little about how their individual contacts might be interconnected or connected to others.

LOCAL SOCIAL NETWORK

EXPANDER GRAPH

SIMULATED SMALLPOX ATTACKS

DAY 1: UNDETECTED SMALLPOX RELEASE

NO RESPONSE

INFECTED: 1,281
DEAD: 0

TARGETED VACCINATION AND QUARANTINE STARTING DAY 14

INFECTED: 1,281
QUARANTINED: 0
VACCINATED: 0
DEAD: 0
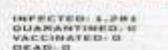
DAY 35: SMALLPOX EPIDEMIC

INFECTED: 23,916
DEAD: 331

INFECTED: 2,564
QUARANTINED: 29,910
VACCINATED: 38,566
DEAD: 312

DAY 70: EPIDEMIC UNCONTAINED OR CONTAINED

INFECTED: 390,582
DEAD: 12,499

INFECTED: 2,564
QUARANTINED: 36,725
VACCINATED: 37,207
DEAD: 495

RESPONSE EFFECTIVENESS

Didier Raoult · Michel Drancourt *Editors*

Paleomicrobiology
Past Human Infections

② Springer

The **Black Death**

2.5 Million People Dead

The Plague and Medicine
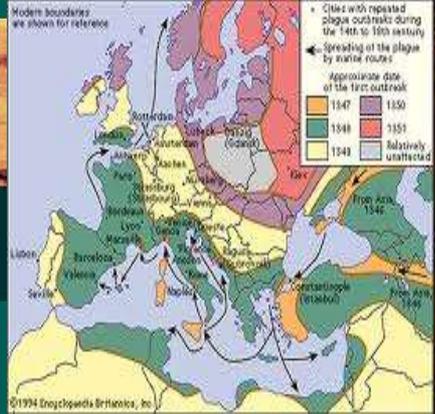
8. TIBIA DROIT D'UNE JEUNE FEMME (Gloucester, XIII°-XIV° siècles) présentant une lésion liée à une tréponématose. Antérieur à 1493, ce cas, avec d'autres observations réalisées en Grande-Bretagne, conforte la thèse d'une origine européenne de la syphilis.
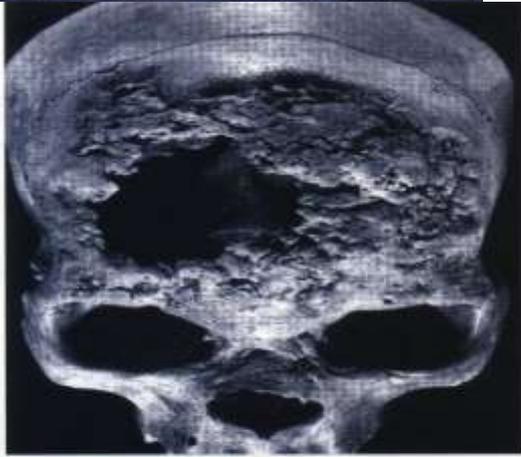
9. CRÂNE FÉMININ présentant des lésions typiques de tréponématose (Mexique, époque pré-colombienne). Les lésions du tréponème, prennent, sur le crâne, un aspect de nodules, nommés gommes.

10. FŒTUS d'environ sept mois présent dans la cavité pelvienne de sa mère (Costebelle, IV° siècle). Les lésions résultent d'une syphilis congénitale et infirment l'origine américaine de la maladie.

AÑO 2006

Breuiffima relación de la conftrupción de la Infectología de Badajoz: colegida por el doct°r Só Aguftin Muñoz Sanz/del Serviçio Extremeño de Salud y de la Uniuerfidad de Extremadura
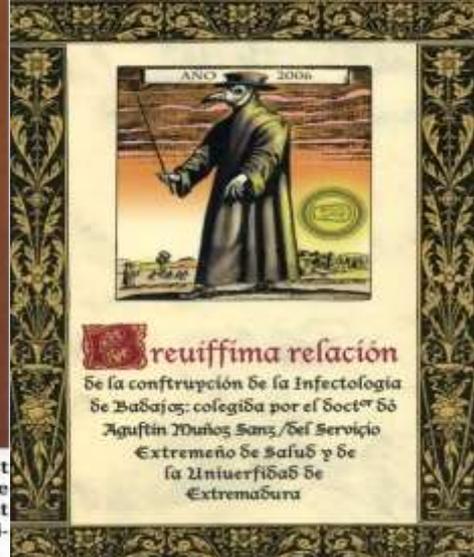
INFLUENZA

- **Séc. XVIII**
  - Três grandes epidemias
- **Séc. IX**
  - Quatro grandes epidemias
- **1918 – 1920**
  - "Gripe Espanhola"
    - 20.000.000 – 40.000.000 mortos
- **1957**
  - "Gripe Asiática"
    - 4.000.000 mortos
- **1968**
  - "Gripe de Hong Kong"
    - 1.000.000 mortos

Seattle Police Officers Wearing Protective Face Masks during the Influenza Epidemic of 1918.

Tissue samples and a list of child victims of the 1918 influenza pandemic

REVIEW ARTICLE

CURRENT CONCEPTS

Historical Perspective — Emergence of Influenza A (H1N1) Viruses

Shanta M. Zimmer, M.D., and Donald S. Burke, M.D.

PERSPECTIVE

THE PERSISTENT LEGACY OF THE 1918 INFLUENZA VIRUS
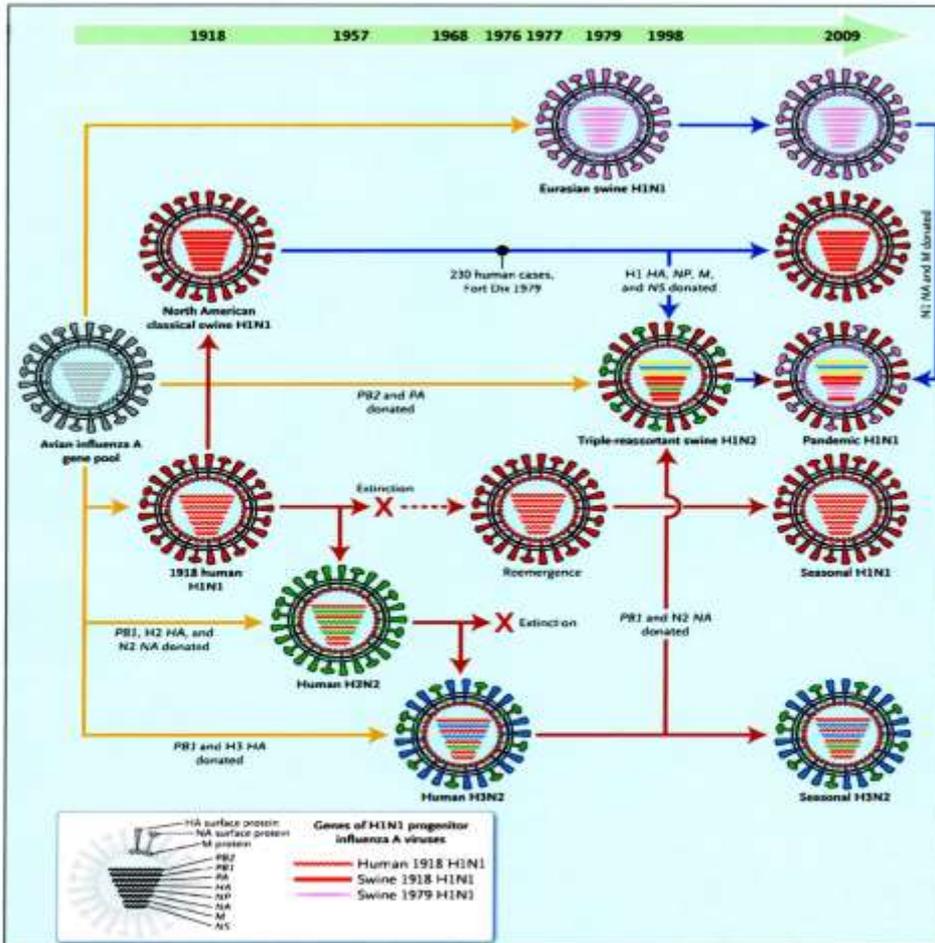
Genetic Relationships among Human and Relevant Swine Influenza Viruses, 1918–2009.

Yellow arrows reflect exportation of one or more genes from the avian influenza A virus gene pool. The dashed red arrow indicates a period without circulation. Solid red arrows indicate the evolutionary paths of human influenza virus lineages; solid blue arrows, of swine influenza virus lineages; and the blue-to-red arrow, of a swine-origin human influenza virus. All influenza A viruses contain eight genes that encode the following proteins (shown from top to bottom within each virus): polymerase PB2, polymerase PB1, polymerase PA, hemagglutinin (HA), nuclear protein (NP), neuraminidase (NA), matrix proteins (M), and nonstructural proteins (NS). The genes of the 1918 human and swine H1N1 and the 1979 H1N1 influenza A viruses were all recently descended from avian influenza A genes, and some have been "donated" to the pandemic human H1N1 strain.
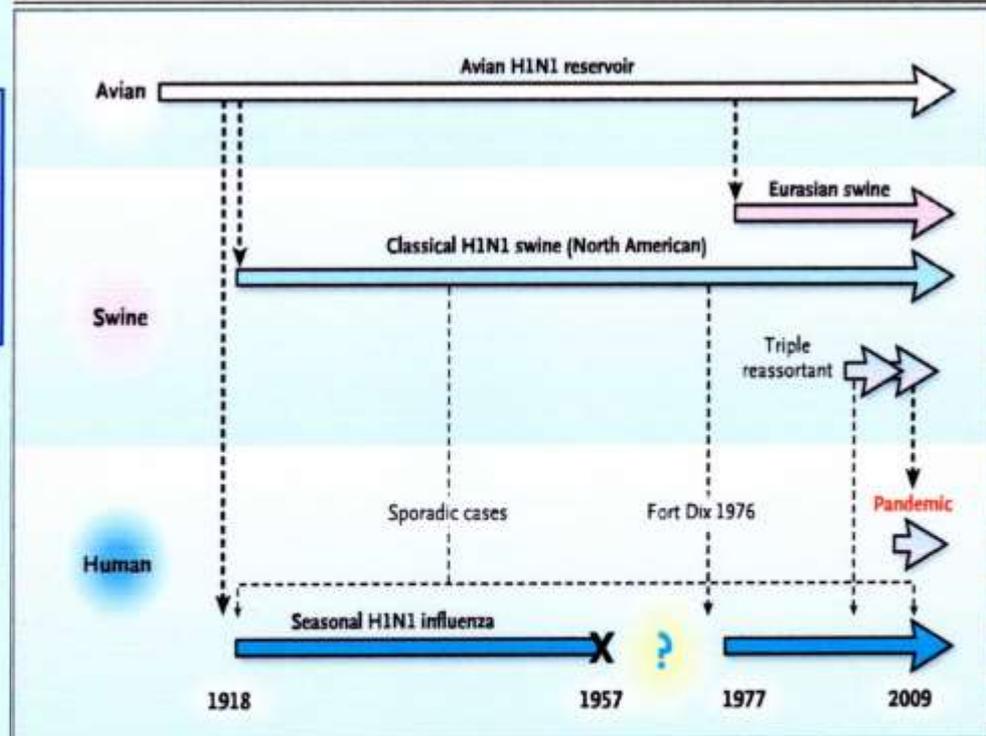
Figure 1. Emergence of Influenza A (H1N1) Viruses from Birds and Swine into Humans.

The diagram shows the important events and processes in the emergence of influenza A (H1N1) viruses during the past 91 years. Avian, swine, and human populations are represented in the top, middle, and bottom of the diagram, respectively. Epidemic or zoonotic viruses are shown as wide horizontal arrows (white for avian viruses, light blue or pink for swine viruses, and dark blue for human viruses). Cross-species transmissions are shown as vertical dashed lines, with thick lines for transfers that gave rise to sustained transmission in the new host and thin lines for those that were transient and resulted in a self-limited number of cases. Principal dates are shown along the bottom of the diagram. The disappearance of H1N1 in 1957 most likely represents competition by the emerging pandemic H2N2 strain in the face of population immunity to H1N1. The reemergence in 1977 is unexplained and probably represents reintroduction to humans from a laboratory source.

Sistema P450 na Metabolização dos Fármacos: I

SUBSTRATES

FDA preferred[1] and acceptable[2] **substrates** for in vitro experiments.*

1A2	2B6	2C8	2C19	2C9	2D6	2E1	3A4,5,7
amitriptyline	bupropion[1]	paclitaxel	**Proton Pump Inhibitors:** lansoprazole omeprazole[2] pantoprazole rabeprazole S-3810	**NSAIDs:** diclofenac[1] ibuprofen lornoxicam meloxicam S-naproxen=>Nor piroxicam suprofen	**Beta Blockers:** carvedilol S-metoprolol propafenone timolol	**Anesthetics:** enflurane halothane isoflurane methoxyflurane sevoflurane	**Macrolide antibiotics:** clarithromycin erythromycin (not 3A5) NOT azithromycin telithromycin
caffeine[2] clomipramine clozapine cyclobenzaprine estradiol fluvoxamine haloperidol imipramine N-DeMe	cyclophosphamide efavirenz[1] ifosfamide methadone	amodiaquine[2] cerivastatin[2] repaglinide	**Anti-epileptics:** diazepam=>Nor phenytoin(O) S-mephenytoin[1] phenobarbitone		**Antidepressants:** amitriptyline clomipramine desipramine imipramine paroxetine	acetaminophen =>NAPQI aniline[2] benzene chlorzoxazone[1] ethanol N,N-dimethyl formamide theophylline =>8-OH	**Anti-arrhythmics:** quinidine=>3-OH (not 3A5)
mexiletine naproxen olanzapine ondansetron phenacetin[1]== acetaminophen=>NAPQI propranolol riluzole ropivacaine tacrine[2] theophylline[2] tizanidine verapamil (R)warfarin zileuton zolmitriptan			amitriptyline carisoprodol citalopram chloramphenicol clomipramine clopidogrel cyclophosphamide hexobarbital imipramine N-DeMe R-mephobarbital moclobamide nelfinavir nilutamide primidone progesterone proguanil propranolol teniposide R-warfarin=>8-OH	**Oral Hypoglycemic Agents:** tolbutamide[1] glipizide **Angiotensin II Blockers:** losartan irbesartan **Sulfonylureas:** glyburide/ glibenclamide glipizide glimepiride tolbutamide amitriptyline celecoxib fluoxetine fluvastatin glyburide nateglinide phenytoin-4-OH[2] rosiglitazone tamoxifen torsemide S-warfarin[1]	**Antipsychotics:** haloperidol perphenazine risperidone=>9OH thioridazine zuclopenthixol alprenolol amphetamine aripiprazole atomoxetine bufuralol[1] chlorpheniramine chlorpromazine codeine (=>O-desMe) debrisoquine[2] dexfenfluramine dextromethorphan[1] duloxetine encainide flecainide fluoxetine fluvoxamine lidocaine metoclopramide methoxyamphetamine mexilletine minaprine nebivolol nortriptyline ondansetron oxycodone perhexiline phenacetin phenformin promethazine propranolol sparteine tamoxifen tramadol venlafaxine		**Immune Modulators:** cyclosporine tacrolimus (FK506) **HIV Antivirals:** indinavir nelfinavir ritonavir saquinavir **Prokinetic:** cisapride **Antihistamines:** astemizole chlorpheniramine terfenadine[2] **Calcium Channel Blockers:** amlodipine diltiazem felodipine lercanidipine nifedipine nisoldipine nitrendipine verapamil **HMG CoA Reductase Inhibitors:** atorvastatin cerivastatin lovastatin NOT pravastatin simvastatin **Steroid 6beta-OH:** estradiol hydrocortisone progesterone testosterone[1] **Miscellaneous:** alfentanyl aprepitant aripiprazole buspirone cafergot caffeine=>TMU cilostazol cinacalcet cocaine codeine- N-demethylation dapsone dexamethasone dextromethorphan[2] docetaxel domperidone eplerenone fentanyl finasteride gleevec haloperidol irinotecan LAAM lapatinib lidocaine methadone nateglinide ondansetron pimozide propranolol quetiapine quinine risperidone NOT rosuvastatin salmeterol sildenafil sirolimus tamoxifen taxol terfenadine trazodone vincristine zaleplon ziprasidone zolpidem
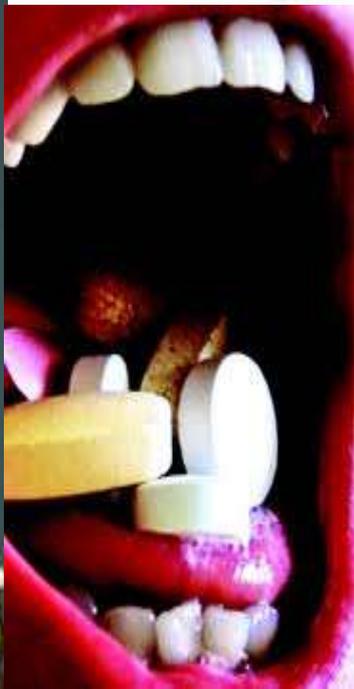
Sistema P450 na Metabolização dos Fármacos: II

INHIBITORS

Inhibitors compete with other drugs for a particular enzyme thus affecting the optimal level of metabolism of the substrate drug which in many cases affect the individual's response to that particular medication, e.g. making it ineffective.

strong

A **Strong inhibitor** is one that cause a > 5-fold increase in the plasma AUC values or more than 80% decrease in clearance.

moderate

A **Moderate inhibitor** is one that cause a > 2-fold increase in the plasma AUC values or 50-80% decrease in clearance.

weak

A **Weak inhibitor** is one that cause a > 1.25-fold but < 2-fold increase in the plasma AUC values or 20-50% decrease in clearance.

others

All other inhibitors.

FDA preferred[1] and acceptable[2] **inhibitors** for in vitro experiments.*

1A2	2B6	2C8	2C19	2C9	2D6	2E1	3A4,5,7
fluvoxamine	thiotepa	gemfibrozil[2]	**PPIs:** lansoprazole omeprazole[2] pantoprazole rabeprazole	fluconazole[2]	bupropion fluoxetine paroxetine quinidine[1]	diethyl-dithiocarbamate[2] disulfiram	**HIV Antivirals:** indinavir nelfinavir ritonavir
ciprofloxacin	ticlopidine[2]			amiodarone			
cimetidine		trimethoprim[2]		fenofibrate fluvastatin	duloxetine terbinafine		clarithromycin itraconazole[1]
amiodarone		glitazones		fluvoxamine[2]	amiodarone		ketoconazole[1]
fluoroquinolones		montelukast[1]	chloramphenicol cimetidine	isoniazid lovastatin	cimetidine		nefazodone
furafylline[1]		quercetin[1]	felbamate		sertraline		saquinavir
interferon			fluoxetine	phenylbutazone			telithromycin
methoxsalen			fluvoxamine	probenecid	celecoxib		aprepitant
mibefradil			indomethacin	sertraline	chlorpheniramine		erythromycin
			ketoconazole		chlorpromazine		fluconazole
			modafinil	sulfamethoxazole	cinacalcet		grapefruit juice
			oxcarbazepine		citalopram		verapamil[2]
			probenicid	sulfaphenazole[1]	clemastine		diltiazem
			ticlopidine[2]	teniposide	clomipramine		
			topiramate	voriconazole	cocaine		cimetidine
				zafirlukast	diphenhydramine		
					doxepin		amiodarone
					doxorubicin		NOT
					escitalopram		azithromycin
					goldenseal		chloramphenicol
					halofantrine		delaviridine
					histamine H1 receptor antagonists		
					hydroxyzine		diethyl-dithiocarbamate
					levomepromazine		fluvoxamine
					methadone		gestodene
					metoclopramide		imatinib
					mibefradil		mibefradil
					midodrine		mifepristone
					moclobemide		norfloxacin
					perphenazine		norfluoxetine
					ranitidine		star fruit
					red-haloperidol		voriconazole
					ritonavir		
					ticlopidine		
					tripelennamine		

INDUCERS

Inducers stimulate the production of the enzyme thus increasing the rate of metabolism causing the substrate drug to clear out of the system faster. This will also affect the individual's response to the medication, i.e. making he drug ineffective because it has not been in the system long enough to have an effect.

FDA preferred[1] and acceptable[2] **inducers** for in vitro experiments.*

1A2	2B6	2C8	2C19	2C9	2D6	2E1	3A,4,5,7
broccoli brussel sprouts char-grilled meat insulin methylcholanthrene[1] modafinil nafcillin beta-naphthoflavone[1] omeprazole[1] tobacco	phenobarbital rifampin	rifampin[1]	carbamazepine norethindrone NOT pentobarbital prednisone rifampin[1]	rifampin secobarbital	dexamethasone rifampin	ethanol isoniazid	**HIV Antivirals:** efavirenz nevirapine barbiturates carbamazepine efavirenz glucocorticoids modafinil nevirapine oxcarbazepine phenobarbital[2] phenytoin[2] pioglitazone rifabutin rifampin[1] St. John's wort troglitazone[1]

GENETICS

1A2	2B6	2C8	2C19	2C9	2D6	2E1	3A4,5,7
Chr15	Chr19	Chr10	Chr10	Chr10	Chr22	Chr10	Chr7

HELIX HEALTH®

PREDICTION PREVENTION PRIVACY

GenomeMedicine

Medicine in the post-genomic era

CURIOSITY GUIDES

THE HUMAN GENOME

The Book of Essential Knowledge

John Quackenbush, Ph.D.

Proteins Peptides

MAPPING OUR GENES

The Genome Project and the Future of Medicine

LOIS WINGERSON

GENOMIC MEDICINE: TURNING PROMISE INTO PRACTICE

Genomic Medicine

AGCTTACCGTAAT

S.B. Primrose and R.M. Twyman

PRINCIPLES OF GENE MANIPULATION AND GENOMICS

EIGHTH EDITION

GENOME RESEARCH

Genomic achievements since the Human Genome Project

Genoma Humano

- 23 (22+1) pares de Cromosomas
- 23.000 Genes
- 3.000.000.000 de pares de Nucleotidos
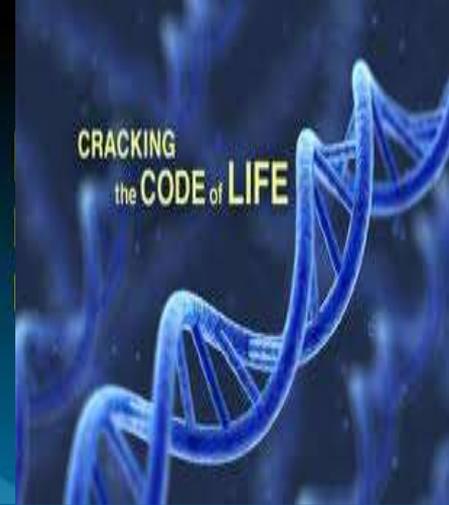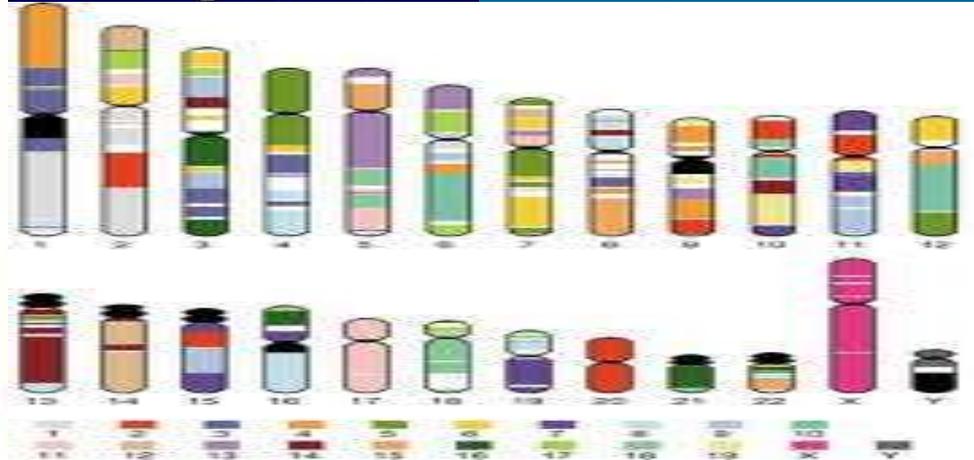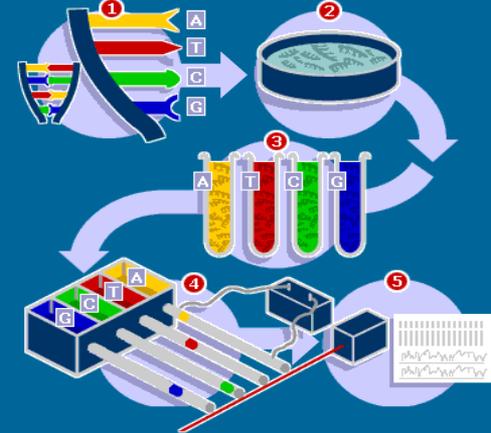
A Questão dos Endoretrovírus

- ⊚ **ERVs (Endo-Retrovírus)** *(Singh,S et al, Brit. J. Dermat., 2009, 161, 6, 1225-1231)*
  - › **Estudos remontam à década de 70**
  - › **8% do total do Genoma Humano**
  - › **Já foram identificados cerca de 20 estirpes diferentes**
  - › **Tiveram origem numa integração que remonta há 25.000.000 anos**
  - › **Estão presentes em muitos Símios**
  - › **Poderão ter uma relação etiopatogénica c/ várias D. ditas ideopáticas e crónicas (Neoplasias, D. auto-imunes, etc)**
    - · **Esclerose Múltipla**
    - · **Artrite Reumatóide**
    - · **Esquizofrenia**
    - · **Lupus Eritematoso Sistémico**
    - · **Psoríase**
    - · **Sindromas Linfoproliferativos**
    - · **Melanoma**
    - · **Tumores das Células Germinais**
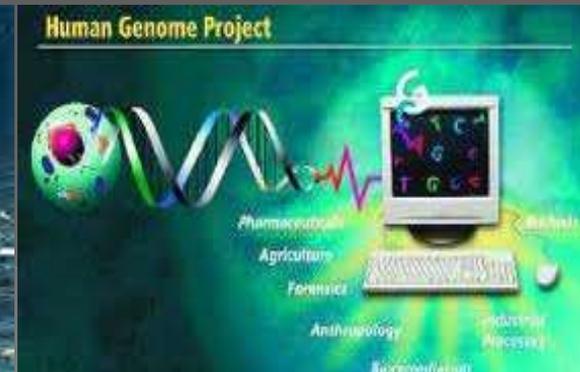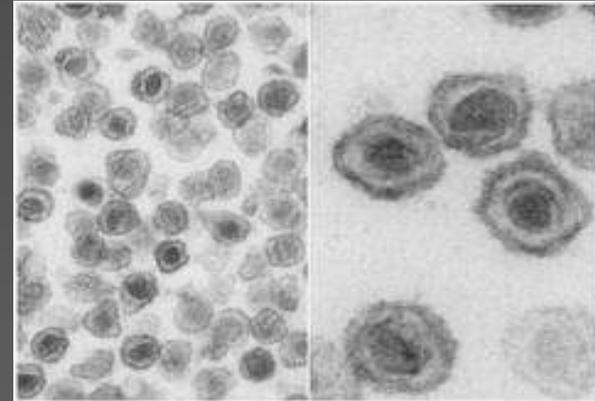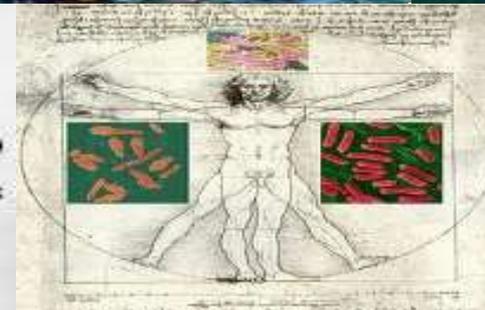
Discovery and analysis of the first endogenous lentivirus

Aris Katzourakis*, Michael Tristem[†], Oliver G. Pybus*, and Robert J. Gifford[‡§]

*Department of Zoology, University of Oxford, South Parks Road, Oxford OX1 3PS, United Kingdom; [†]Division of Biology, Imperial College London, Silwood Park, Ascot, Berkshire SL5 7PY, United Kingdom; and [‡]Division of Infectious Diseases, Stanford University, Stanford, CA 94305
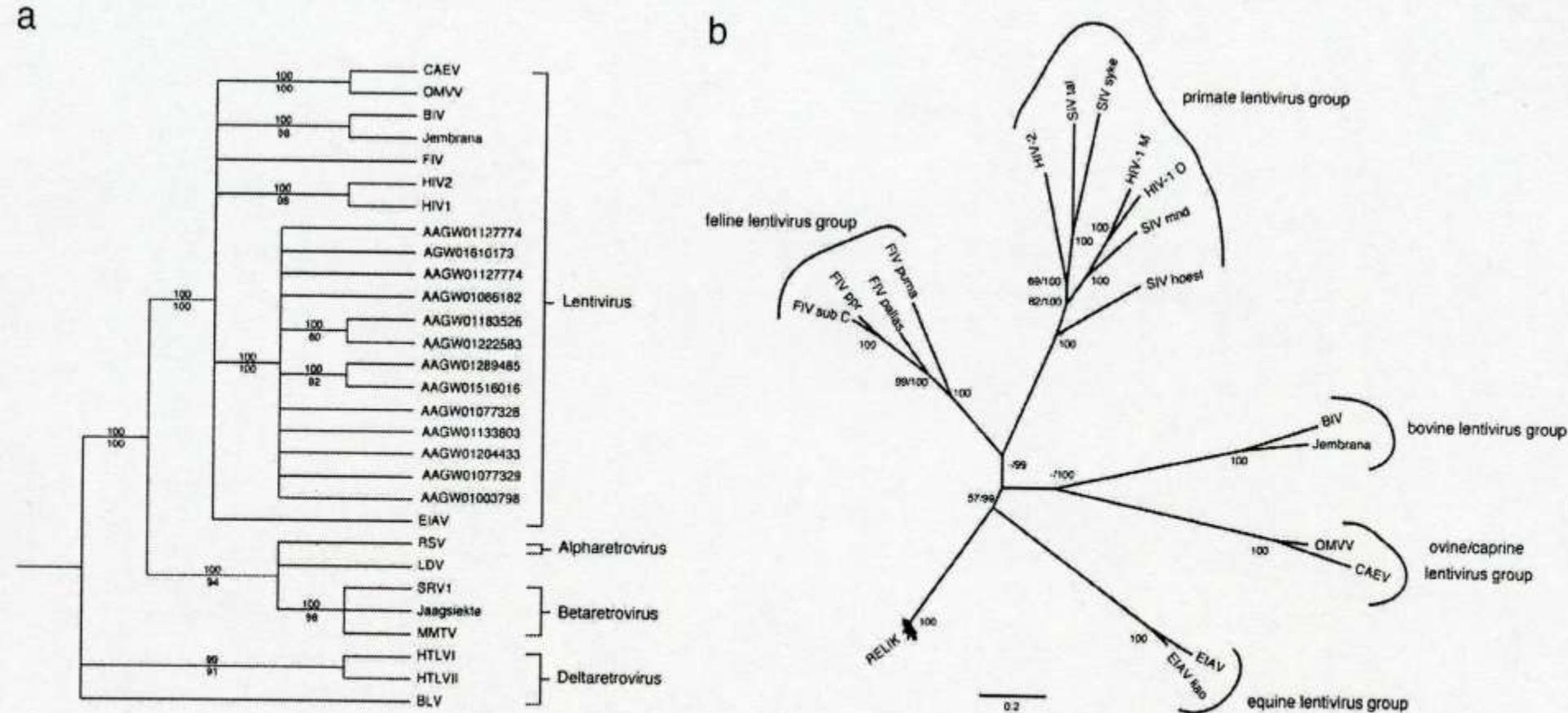
Fig. 3. Phylogenetic relationships of RELIK to other retroviruses. (a) Phylogeny of RELIK and other lentiviruses together with a sample of nonlentiviral exogenous retroviruses, rooted on BLV, HTLV1, and HTLV2. Support for the ML trees was assessed via 1,000 nonparametric bootstrap replicates, and posterior clade probabilities were assessed for the Bayesian phylogeny. Both support indices are indicated as values out of 100, with posterior clade probabilities indicated above the branches and bootstrap scores below. Branches with posterior probabilities <95% were collapsed. The RELIK sequences are indicated by their accession numbers. (b) Unrooted phylogeny showing the relationships of the lentiviruses. Bootstrap scores and Bayesian posterior probabilities are indicated to the left and right of the forward slash, respectively, and nodes with only 100 indicated showed maximal support under both measures.

Emerging Infectious Determinants of Chronic Diseases

Siobhán M. O'Connor,* Christopher E. Taylor,† and James M. Hughes‡

Timeline	Infectious agent*	Chronic disease
1950	Schistosoma spp	Bladder cancer
	Epstein-Barr virus	Burkitt lymphoma
	Hepatitis B virus	Chronic liver disease, hepatocellular carcinoma
	Human papillomavirus	Cervical cancer
	Borrelia burgdorferi	Chronic Lyme arthritis
	Human T-lymphotropic virus type 1	Acute T-cell leukemia/lymphoma
	Helicobacter pylori	Chronic gastritis, peptic ulcer disease
	Campylobacter jejuni	Guillain-Barré syndrome (persistent sequelae)
	Human T-lymphotropic virus type 1	Tropical spastic paraparesis, chronic arthropathy
	B. burgdorferi	Neuroborreliosis
	Hepatitis C virus	Chronic liver disease, hepatocellular carcinoma
	Bartonella henselae	Bacillary angiomatosis
	Hepatitis C virus	Mixed cryoglobulinemia
	Tropheryma whipplei	Whipple disease
	KSHV/human herpesvirus 8	Kaposi sarcoma
	New variant CJD agent	Variant CJD
2000	West Nile virus	Paralysis (persistent postpoliomyelitis)

*KSHV, Kaposi sarcoma herpesvirus; CJD, Creutzfeldt-Jakob disease

Figure 1. Emergence timeline for infectious determinants of chronic diseases. For references to support this figure, see online version (available from http://www.cdc.gov/ncidod/EID/vol12no07/06-0037-G1.htm)

Pathogens	Syndromes and organ systems	Triggers and outcomes	Duration of infection	Timing of infection
Bacteria	Cardiovascular	1 microbe → multiple syndromes	Acute	Prenatal
Fungi	Endocrine		Persistent active	Infancy
Parasites	Gastrointestinal	Several microbes → 1 outcome	Persistent non-replicating	Childhood
Prions	Immune		Cleared	Adolescence
Viruses	Musculoskeletal		Enduring normal flora	Adulthood
	Neurologic			Recurrent or coinfection
	Neuropsychiatric			
	Ocular			
	Pulmonary			
	Renal			
	Respiratory			
	Skin			

Figure 2. Infectious determinants of chronic diseases.

One Health
initiative

THE LANCET Infectious Diseases

LE NOUVEAU
NEWKASTLE,
OU
NOUVEAU TRAITÉ
DE
CAVALERIE.

A LYON,
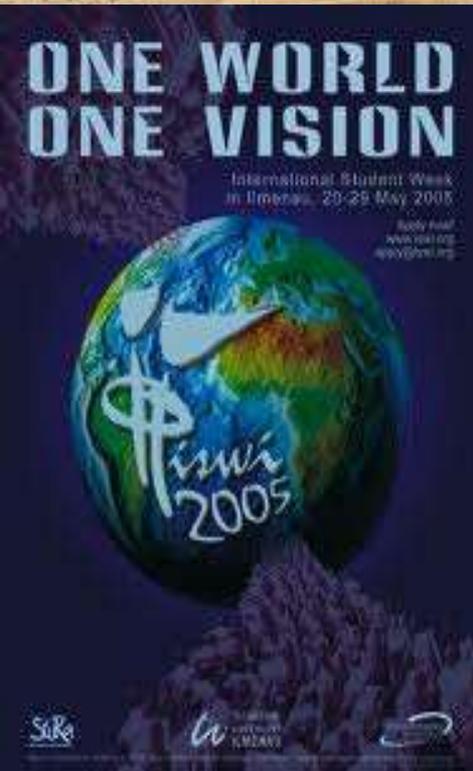
Towards a conceptual framework to support one-health research for policy on emerging zoonoses

Richard Coker, Jonathan Rushton, Sandra Mounier-Jack, Esron Karimuribo, Pascal Lutumba, Dominic Kambarage, Dirk U Pfeiffer, Katharina Stärk, Mark Rweyemamu

In the past two decades there has been a growing realisation that the livestock sector was in a process of change, resulting from an expansion of intensive animal production systems and trade to meet a globalised world's increasing demand for livestock products. One unintended consequence has been the emergence and spread of transboundary animal diseases and, more specifically, the resurgence and emergence of zoonotic diseases. Concurrent with changes in the livestock sector, contact with wildlife has increased. This development has increased the risk of transmission of infections from wildlife to human beings and livestock. Two overarching questions arise with respect to the real and perceived threat from emerging infectious diseases: why are these problems arising with increasing frequency, and how should we manage and control them? A clear conceptual research framework can provide a guide to ensure a research strategy that coherently links to the overarching goals of policy makers. We propose such a new framework in support of a research and policy-generation strategy to help to address the challenges posed by emerging zoonoses.
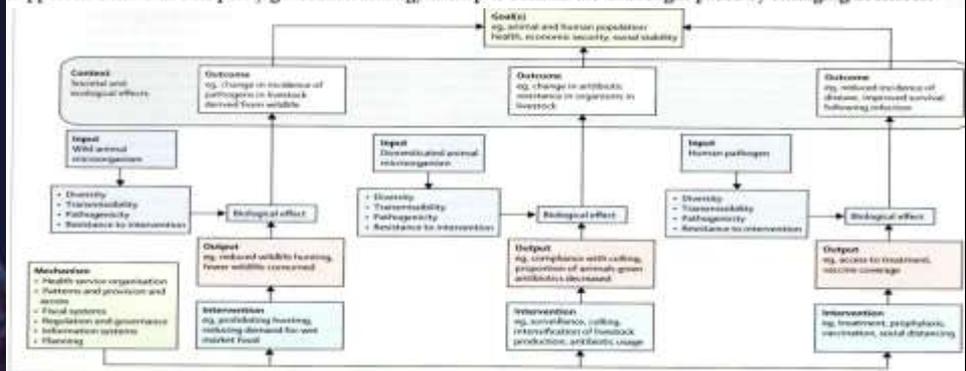
Figure: Schematic representation of a framework for research to inform one-health policy

ONE WORLD
ONE VISION

International Student Week
in Ilmenau, 20-29 May 2005

iswi
2005

STEPHEN MICHAEL APATOW

ONE WORLD
ONE HEALTH

One Health Initiative

SM AM EDIA COMMUNICATIONS: INTERNATIONAL

24 X 7

HUMANITARIAN RESOURCE INSTITUTE:
GLOBAL ARTS INTEGRATION INTO EDUCATION
WWW.UNARTS.ORG
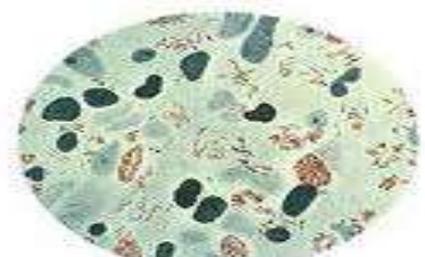
M. leprae bacteria, the cause of leprosy. Source: U.S. Centers for Disease Control and Prevention

Leprosy bacteria transmission from armadillo to man

INDIAN CAMP PLANTATION
The plantation home, built in the 1850's, became the site of the Louisiana State Leprosarium in 1894. The U.S. Public Health Service acquired it in 1921. It is now known as the National Hansen's Disease Center.
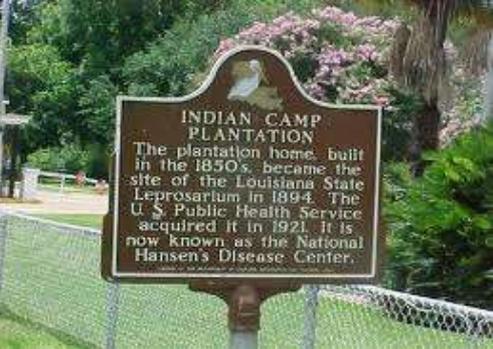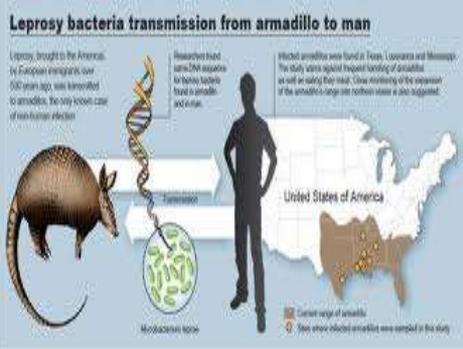
ORIGINAL ARTICLE

Probable Zoonotic Leprosy in the Southern United States

Richard W. Truman, Ph.D., Pushpendra Singh, Ph.D., Rahul Sharma, Ph.D., Philippe Busso, Jacques Rougemont, Ph.D., Alberto Paniz-Mondolfi, M.D., Adamandia Kapopoulou, M.S., Sylvain Brisse, Ph.D., David M. Scollard, M.D., Ph.D., Thomas P. Gillis, Ph.D., and Stewart T. Cole, Ph.D.

ABSTRACT

BACKGROUND

In the southern region of the United States, such as in Louisiana and Texas, there are autochthonous cases of leprosy among native-born Americans with no history of foreign exposure. In the same region, as well as in Mexico, wild armadillos are infected with *Mycobacterium leprae*.

METHODS

Whole-genome resequencing of *M. leprae* from one wild armadillo and three U.S. patients with leprosy revealed that the infective strains were essentially identical. Comparative genomic analysis of these strains and *M. leprae* strains from Asia and Brazil identified 51 single-nucleotide polymorphisms and an 11-bp insertion–deletion. We genotyped these polymorphic sites, in combination with 10 variable-number tandem repeats, in *M. leprae* strains obtained from 33 wild armadillos from five southern states, 50 U.S. outpatients seen at a clinic in Louisiana, and 64 Venezuelan patients, as well as in four foreign reference strains.

RESULTS

The *M. leprae* genotype of patients with foreign exposure generally reflected their country of origin or travel history. However, a unique *M. leprae* genotype (3I-2-v1) was found in 28 of the 33 wild armadillos and 25 of the 39 U.S. patients who resided in areas where exposure to armadillo-borne *M. leprae* was possible. This genotype has not been reported elsewhere in the world.

CONCLUSIONS

Wild armadillos and many patients with leprosy in the southern United States are infected with the same strain of *M. leprae*. Armadillos are a large natural reservoir for *M. leprae*, and leprosy may be a zoonosis in the region. (Funded by the National Institute of Allergy and Infectious Diseases and others.)

Figure 1. Genotyping of Mycobacterium leprae Strains.
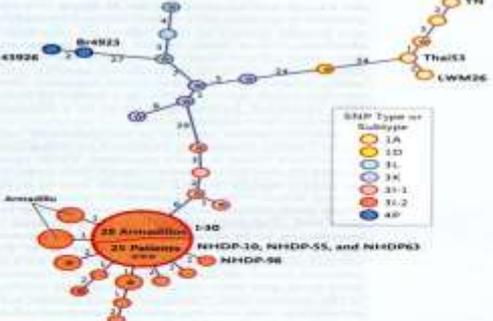
Figure 3. Minimum-Spanning Phylogenetic Tree of Mycobacterium leprae Genotypes Based on Analysis of Single-Nucleotide Polymorphisms (SNPs) and Variable-Number Tandem Repeats (VNTRs).

Des cancers
traités par des virus

J. ROMMELAERE, C. DINSART, A. RÉGNIER-VIGOUROUX, N. SALOMÉ, J. SCHLEHOFER

Les traitements conventionnels contre les cancers ont une efficacité limitée dans de nombreux cas. La thérapie oncolytique, fondée sur l'utilisation de virus tueurs de tumeurs, offre de nouvelles perspectives dans la lutte contre ce fléau mondial.


1. CETTE BULLE STERILE protégeait David, atteint d'un déficit immunitaire combiné sévère (DICS), maladie héréditaire. Un essai de thérapie génique, réalisé en 1999 à l'Hôpital Necker de Paris, a permis de soigner huit enfants. Par la suite, des améliorations des vecteurs rétroviraux ont été proposées.

3. RÉGRESSION D'UNE TUMEUR CÉRÉBRALE. Des essais cliniques ont montré l'efficacité de la thérapie oncolytique sur des cellules gliales cancéreuses du cerveau. L'injection de virus de la maladie de Newcastle a provoqué la régression de la tumeur (de a à f), comme le montrent les images par résonance magnétique prises lors du test, en 2006.

Les virus vecteurs
de molécules thérapeutiques

Jean-Christophe PAGÈS et Éric PIVER

En désamorçant leur pouvoir pathogène, on convertit des virus en vecteurs d'instructions génétiques. Ceux-ci deviennent des outils thérapeutiques, capables de livrer leur message à des cellules spécifiques.

1. POUR FABRIQUER UN VACCIN génétique, on isole un ou plusieurs gènes (a) d'un agent pathogène que l'on insère dans des plasmides (b), des boucles d'ADN. On introduit alors ces plasmides dans des cellules, souvent par injection intramusculaire (c) ou par un pistolet à gènes (d). Les gènes sélectionnés codent des antigènes — des substances qui entraînent la réaction immunitaire — synthétisés normalement par l'agent pathogène.


3. LE CYCLE D'UN VECTEUR VIRAL est tronqué par rapport à celui d'un virus : ne sont conservés que l'entrée et le transfert nucléaire du génome, qui permettront l'expression du gène thérapeutique. Comme les gènes structuraux ne sont plus présents dans le génome du vecteur, celui-ci est incapable de se multiplier.

imatinib
STI-571
Gleevec
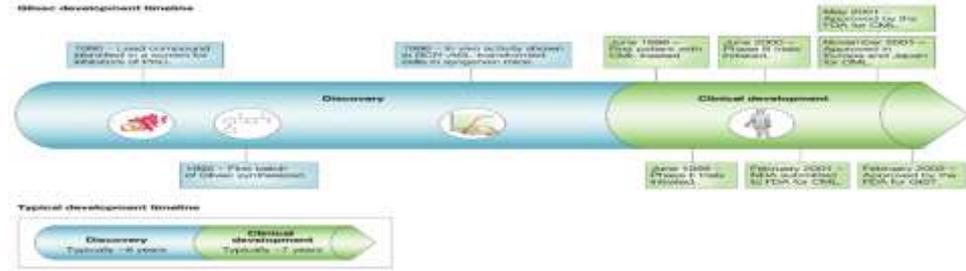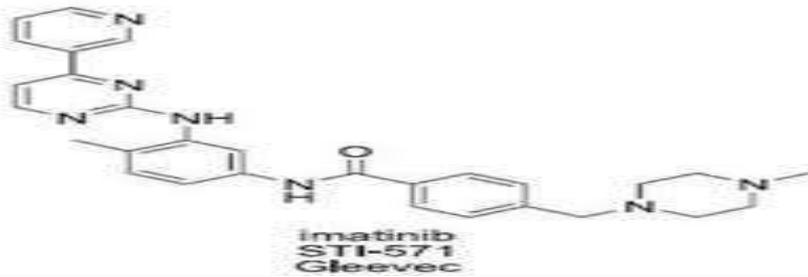
Source: Nature Reviews | Drug Discovery

Diagnosis

Glivec therapy

Adequate response — Inadequate response

No suitable donor | HLA-matched donor | HLA-matched donor | No suitable donor

Loss of response

Continued treatment with Glivec | ? | Allogeneic transplantation with conventional or reduced-intensity regimen | Increased dose of Glivec combined with conventional therapy or novel agents, including dasatinib

IMATINIB GREATLY IMPROVED SURVIVAL IN GIST

Overall survival (% pts)

Glivec®

Doxorubin-based

O	N					Treatment
273	946	850	786	638	393	—
79	86	57	31	19	14	
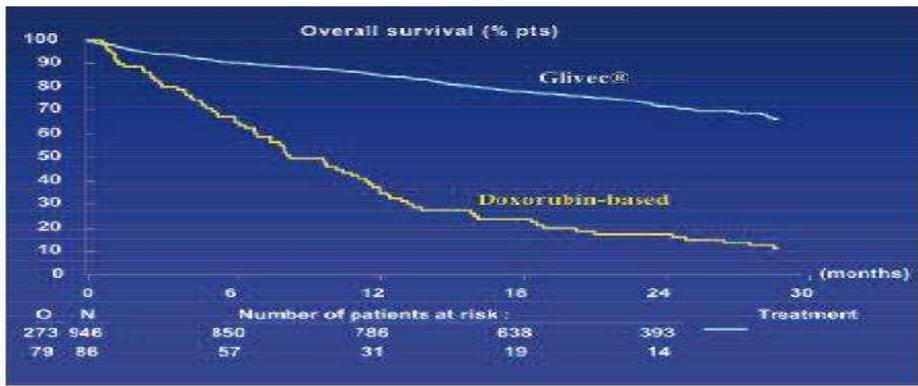
Results from the Conticanet series of GIST patients demonstrated the huge survival benefit conferred by the new therapy

Source: Adapted from J Verweij et al. *The Lancet* 2004, 364:1127–1134

Medscape

a

PML H₂N — R B B CC — COOH
                       394   552

RARα NH₂ — A B C D E F — COOH

t(15;17)

PML–RARα H₂N — R B B CC B C D E F — COOH

b

HDAC Co-repressors HDAC

PML-RARα PML-RARα

RA 10⁻⁶ M   Co-activators HAT

RARα targets are activated

PML-induced dimerization: enhanced co-repressor binding and repression through histone deacetylation

Clinical remission ← Differentiation
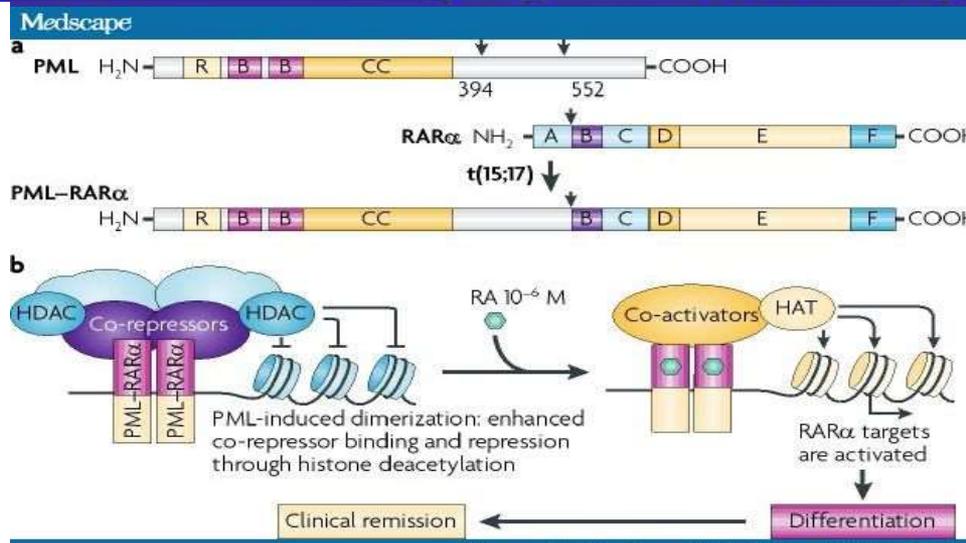
Source: Nat Rev Cancer © 2010 Nature Publishing Group

ZIAGEN
(abacavir sulfate)
TABLETS
300 mg

GX 623

Abacavir Patch Test
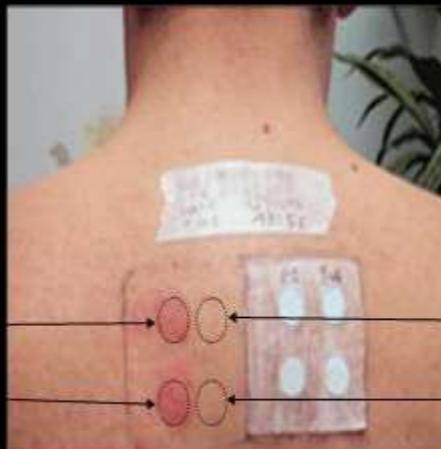
1% abacavir → → Petroleum control
10% abacavir → → Excipient control

24 h   48 h

S. Mallal, 3rd IAS Conference, Sydney, July 2007

www.aids-images.ch

Drug Rash with eosinophilia and systemic syndrome DRESS, here due to abacavir

www.aids-images.ch

Abacavir Hypersensitivity: Effects of HLA-B*5701 testing
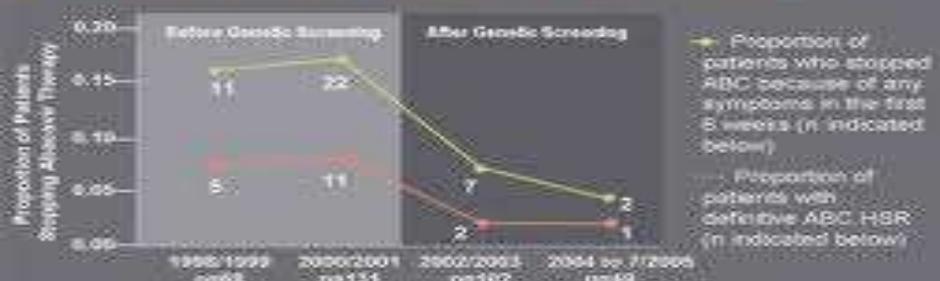
Lower incidence of ABC HSR during screening period than before screening: 2% vs 8% ($P=0.01$)

Life Science Technologies

Proteomics
INTERACTING INSTRUMENTS
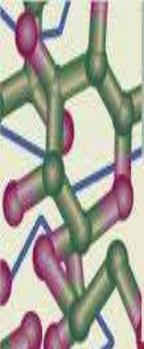
prom!se
ADVANCED PROTEOMICS

applied
proteomics
inc.

PROTEOMICS
*Journal of* PROTEOMICS
Register to receive information
on our new journal

Principles of Proteomics

ADVANCED TEXT

Genome, Transcriptome, Proteome
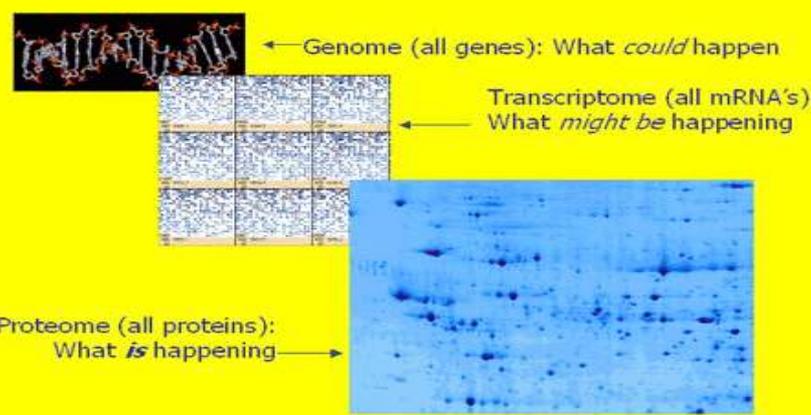
Genome (all genes): What *could* happen

Transcriptome (all mRNA's):
What *might be* happening

Proteome (all proteins):
What *is* happening

STRUCTURAL
PROTEOMICS
and its impact on the
LIFE SCIENCES

Joel L. Sussman
Israel Silman

World Scientific
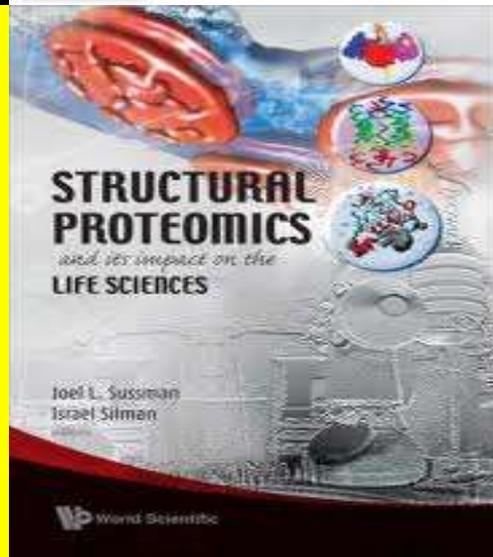
Hubert Rehm Thomas Letzel

Der Experimentator
**Proteinbiochemie/
Proteomics**

Der Experimentator

Spektrum

MCP MOLECULAR & CELLULAR
PROTEOMICS

Clinical
Proteomics
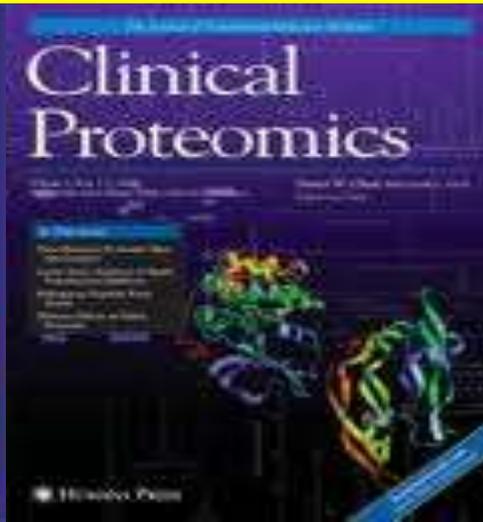
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www.proteomics-journal.de
4'04

Technology
Bioinformatics
Cell Biology
Microbiology
Plant Proteomics
Clinical Proteomics

First
Impact Factor
4.007

WILEY-VCH
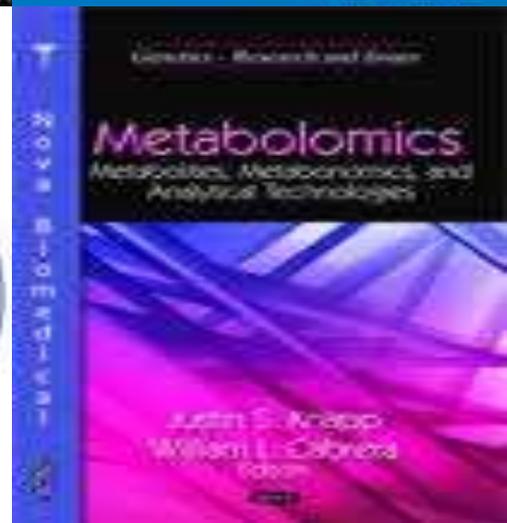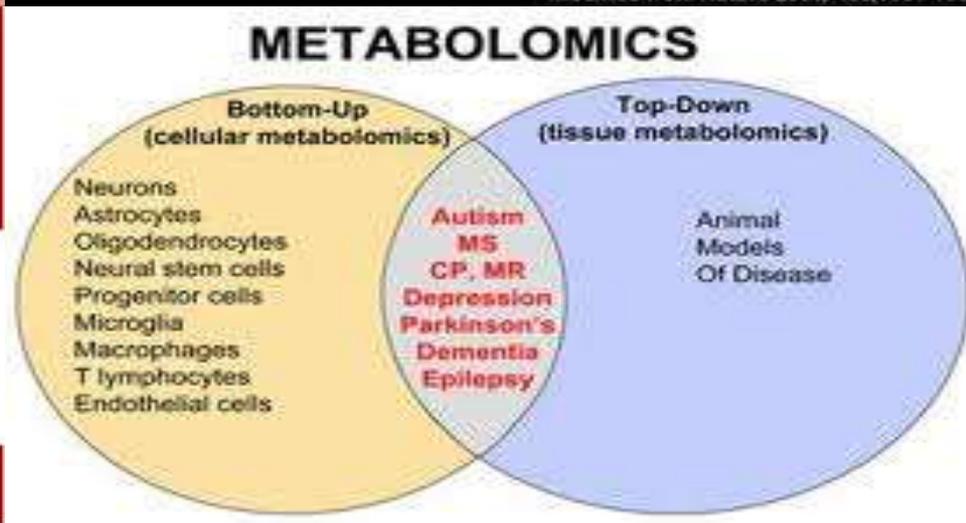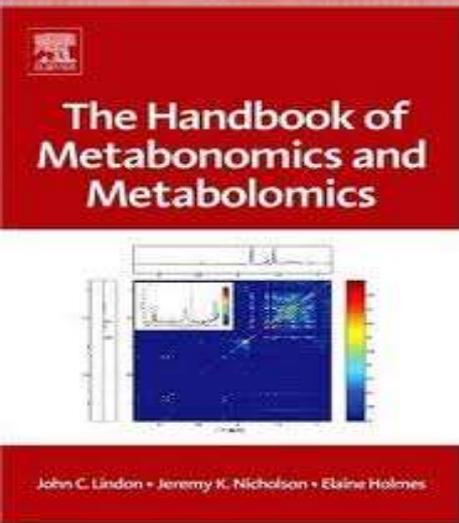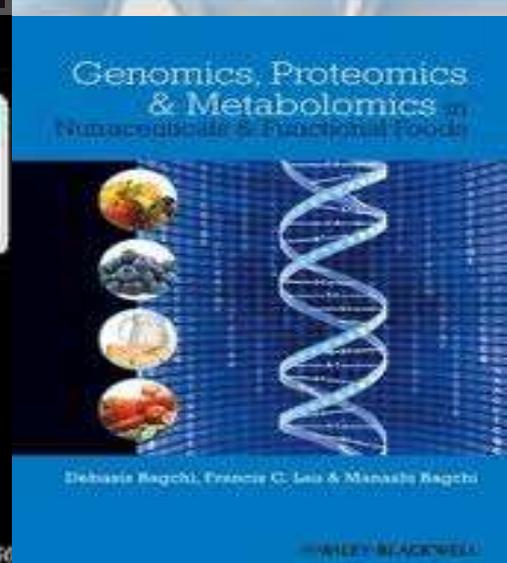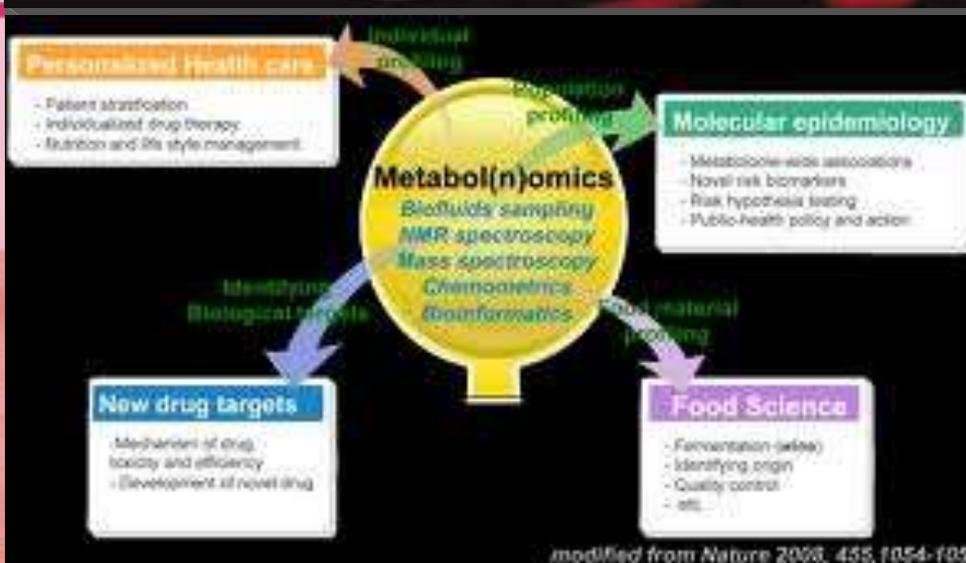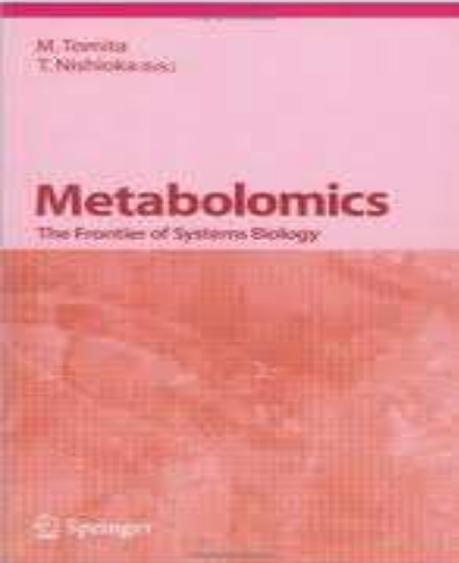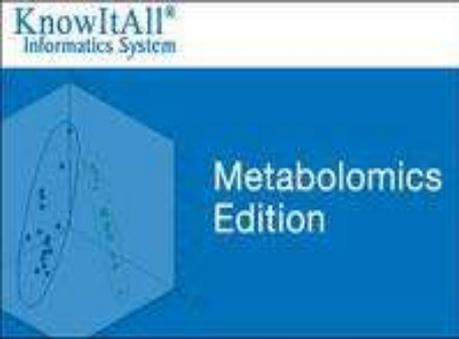
REVIEW ARTICLE

CURRENT CONCEPTS

Nanomedicine

Betty Y.S. Kim, M.D., Ph.D., James T. Rutka, M.D., Ph.D., and Warren C.W. Chan, Ph.D.

nanotechnology SCIENCE

CLICK CHEMISTRY
GENE ONTOLOGY
METABOLOMICS
NANOMEDICINE
OMICS
PROBE TOMOGRAPHY
QUANTITATIVE PROTEOMICS
SYNTHETIC BIOLOGY
SYSTEMS BIOLOGY
TRACTOGRAPHY

Table 1. Examples of Nanomaterials in Clinical Use.*

| Nanomaterial	Trade Name	Application	Target	Adverse Effects	Manufacturer	Current Status
Metallic						
Iron oxide	Feridex	MRI contrast	Liver	Back pain, vaso-dilatation	Bayer Schering	FDA approved
	Resovist	MRI contrast	Liver	None	Bayer Schering	FDA approved
	Combidex	MRI contrast	Lymph nodes	None	Advanced Magnetics	In phase 3 clinical trials
	NanoTherm	Cancer therapy	Various forms	Acute urinary retention	MagForce	In phase 3 clinical trials
Gold	Verigene	In vitro diagnostics	Genetic	Not applicable	Nanosphere	FDA approved
	Aurimmune	Cancer therapy	Various forms	Fever	Cytimmune Sciences	In phase 2 clinical trials
Nanoshells	Auroshell	Cancer therapy	Head and neck	Under investigation	Nanospectra Biosciences	In phase 1 clinical trials
Semiconductor						
Quantum dot	Qdots, EviTags, semiconductor nanocrystals	Fluorescent contrast, in vitro diagnostics	Tumors, cells, tissues, and molecular sensing structures	Not applicable	Life Technologies, eBioscience, Nanoco, CrystalPlex, Cytodiagnostics	Research use only
Organic						
Protein	Abraxane	Cancer therapy	Breast	Cytopenia	Abraxis Bioscience	FDA approved
Liposome	Doxil/Caelyx	Cancer therapy	Various forms	Hand-foot syndrome, stomatitis	Ortho Biotech	FDA approved
Polymer	Oncaspar	Cancer therapy	Acute lymphoblastic leukemia	Urticaria, rash	Rhône-Poulenc Rorer	FDA approved
	CALAA-01	Cancer therapy	Various forms	Mild renal toxicity	Calando	In phase 2 clinical trials
Dendrimer	VivaGel	Microbicide	Cervicovaginal	Abdominal pain, dysuria	Starpharma	In phase 2 clinical trials
Micelle	Genexol-PM	Cancer therapy	Various forms	Peripheral sensory neuropathy, neutropenia	Samyang	For phase 4 clinical trials

* MRI denotes magnetic resonance imaging.
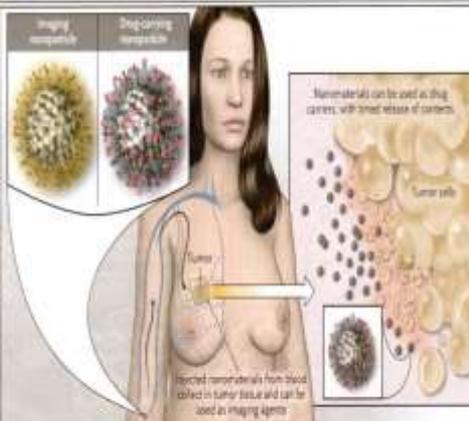
Figure 1. Nanomaterials Used as Drug Carriers or Contrast Agents for In Vivo Cancer Applications.

Tumors have poor lymphatic drainage, and their vessels are highly porous. This enables nanomaterials to diffuse and accumulate in the tumor matrix. Nanomaterials that carry chemotherapeutic agents can target and kill tumor cells, whereas nanomaterials that are magnetic or fluorescent are used as imaging agents for detecting tumors.

Nanomaterials can be used as drug carriers, with small masses of contents.

NANOTECHNOLOGY

nanotechnology

NanoLIFE

Dongfu Shi • Martin E. Yarmush

Nanoimplants

Microneedle-based vaccines

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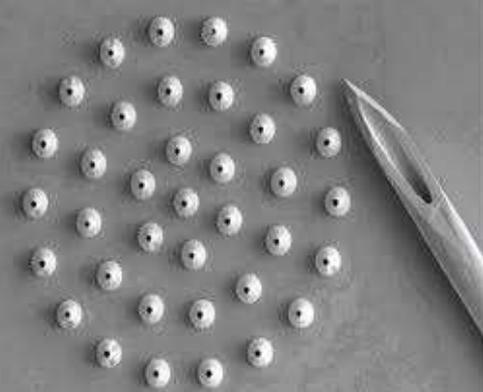
Review

Nanovaccines: recent developments in vaccination

TARALA D NANDEDKAR

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In the past 100 years, vaccination has contributed immensely to public health by preventing a number of infectious diseases. Attenuated, killed or part of the microorganism is employed to stimulate the immune system against it. Progress in biotechnology has provided protective immunity through DNA vaccines. In recent years, nanovaccine is a novel approach to the methodology of vaccination. Nanomaterials are delivered in the form of microspheres, nano-beads or micro-nanoprojections. Painless, effective and safe needle-free routes such as the intranasal or the oral route, or patches of microprojections to the skin are some of the approaches which are in the experimental stage at present but may have a great future ahead in nanovaccination.
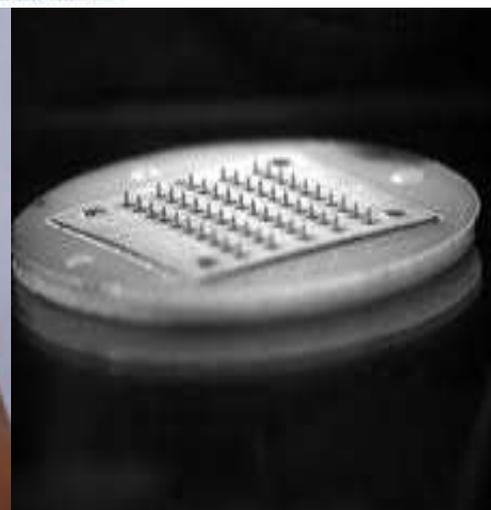
MICRONEEDLES DRUG PATCH

Syringe

Stratum corneum

Epidermis

Dermis

Chip administers drug doses

Pain receptors

150 microneedles

25 mm

SOURCE: Hewlett Packard

DNA Chips and Microarrays

LABORATORY TECHNOLOGY TRENDS

PART I

ARRAY OF OPPORTUNITIES

MICROARRAY CENTRE
University Health Network

BioTechniques BOOKS

Microarray Biochip Technology

Edited by
Mark Schena

GUIDE TO ANALYSIS OF DNA MICROARRAY DATA
SECOND EDITION
STEEN KNUDSEN

Control — Sample

Spotting — Hybridization

RNA Extraction
Labeling

RNA — cDNA

Target — Purification

Scanning

Image Analysis — Informatics

Microarray Technology and Its Applications

Springer

BIOLOGICAL AND MEDICAL PHYSICS
BIOMEDICAL ENGINEERING

Prepare cDNA Probe — Prepare Microarray

"Normal" — Tumor

RT / PCR

Label with Fluorescent Dyes

Combine Equal Amounts

Hybridize probe to Microarray — SCAN —

Microarray Technology

APPLYING GENOMIC AND PROTEOMIC MICROARRAY TECHNOLOGY IN DRUG DISCOVERY

Robert S. Matson

CRC PRESS

INNOVATIONS

Big Lab on a Tiny Chip

Squeezing a chemistry lab down to fingernail size could provide
instant medical tests at home and on the battlefield
By Charles Q. Choi
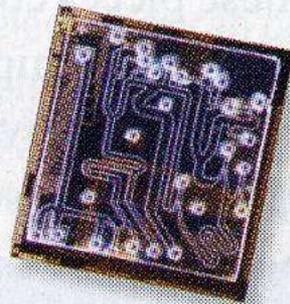
ACTUAL SIZE: The University of Michigan's influenza chip measures 1.5 by 1.6 centimeters.

Influenza Detector

A prototype from the University of
Michigan moves droplets through
a microfluidic maze. Tests on a blood
sample could show influenza or other
pathogens within 15 minutes.

Loading port

Air-injection valve

① LOAD
Blood sample (*yellow*) and
amplification reagent (*green*)
are loaded, then pushed
along by air pressure.

② AMPLIFY
Any DNA in the
sample is amplified
in a PCR chamber
(*red*), which heats
and cools 35 times.

Heaters

Underlying
electronics

③ REACT
Amplified DNA mixes
with a reagent (*gold*)
that reacts to influenza
inside a reaction
chamber (*green*).

④ ANALYZE
An electrophoresis
channel read with
ultraviolet light
indicates a telltale
signature if patho-
gens are present.
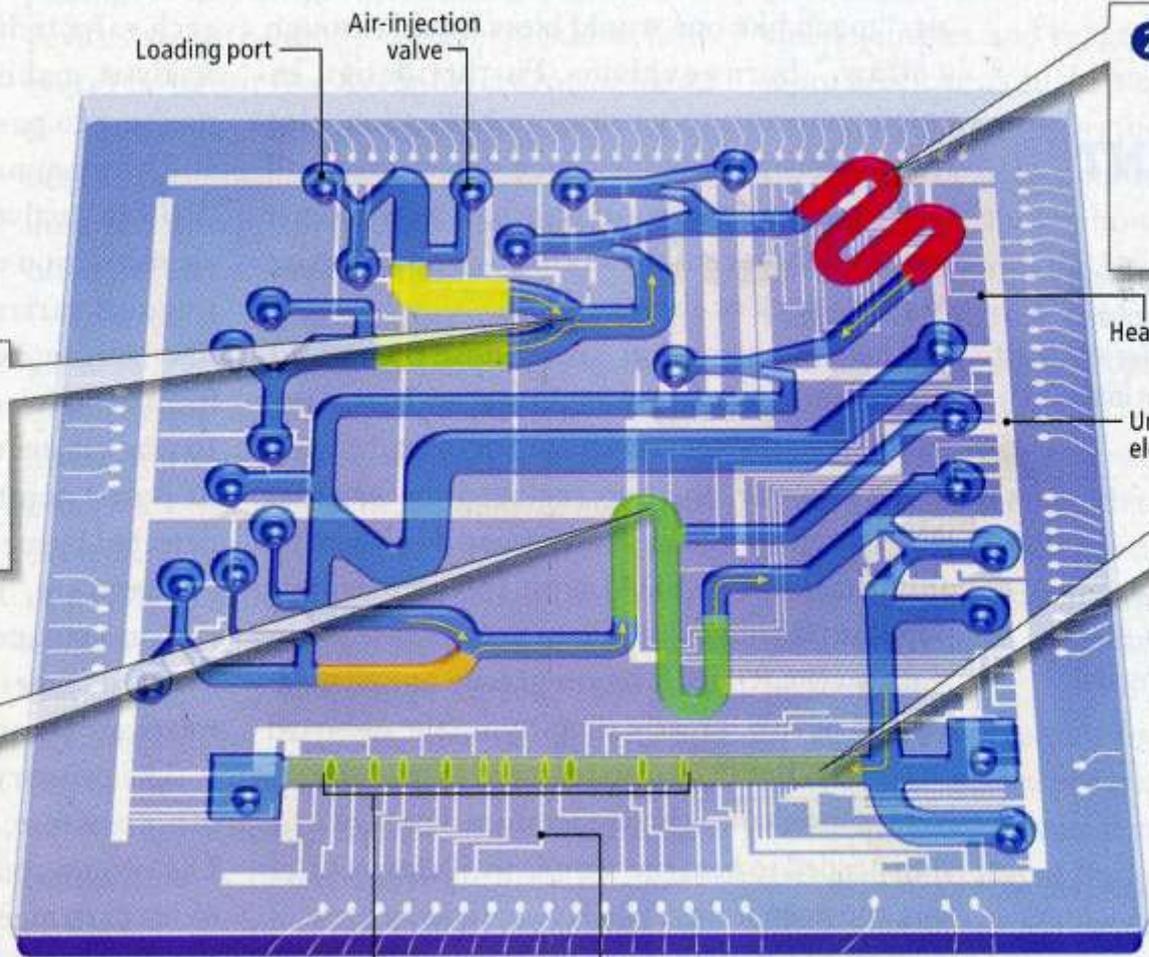
Pathogen
signature

Readout
diode

The Application of Genomics to Emerging Zoonotic Viral Diseases

Bart L. Haagmans, Arno C. Andeweg, Albert D. M. E. Osterhaus*

Department of Virology, Erasmus Medical Center, Rotterdam, The Netherlands

Technology Solutions for Global Health

Point-of-Care Diagnostics for Infectious Diseases: DxBox

Health need

While fever is a common symptom of illness, clinicians often cannot distinguish between bacterial, parasitic, viral, fungal, or noninfectious causes. In the absence of an accurate diagnostic that identifies the causative agent of a fever, treatment is based on regional prevalence of infections. In sub-Saharan Africa, fever is often treated with antimalarial drugs, which leads to over-prescription and mismanagement of the patient's care regimen. The emergence of antimalarial and antibiotic drug resistance results in an urgent need for accurate diagnosis of fever-causing agents at point-of-care in resource-limited settings.

www.medscape.com

Prototype of the DxBox instrument with a disposable device.

Authors and Disclosures

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Abstract and Introduction

Abstract

Extrapolation from recent disease history suggests that changes in the global environment, including virus, vector and human behavior, will continue to influence the spectrum of viruses to which humans are exposed. In this article, these environmental changes will be enumerated, and their potential impact on target-focused, nucleic acid-based diagnostic tests will be considered, followed by a presentation of some emerging technological responses.

Table 1. The overlapping spectrum of possible signs and symptoms for some of the clinically significant flaviviruses.

Virus	Symptom							
	Fever	Headache	Nausea/vomiting	Dizziness/disorientation	Encephalitis	Myalgia	Rash	Hemorrhagic fever
MVEV	√	√	√	√	√	√	√	
YFV	√	√	√	√	√	√	Jaundice	√
DENV	√	√	√	√	√	√	Jaundice	√
WNV	√	√	√	√	√	√	√	
JEV	√	√	√	√	√			
KUNV	√	√	√	√	Rare	√		
USUV	√	√	√	√		√		
TBEV	√	√	√	√	√	√		
SLEV	√	√	√	√	√	√		
KFDV	√	√	√	√	√	√		√
OHFV	√	√	√	√	√	√	√	√
AHFV	√	√	√	√	√	√		√

Not all symptoms or signs occur in all cases and there are febrile or meningoencephalitic forms of some of the diseases (e.g., TBEV), which present and progress differently. For several of these viruses, a significant proportion of infections can be asymptomatic (e.g., WNV). Substantial fatality is associated with symptomatic JEV, MVEV, AKHV, KFDV, WNV, DENV and YFV in unvaccinated populations.

AHFV: Alkhurma hemorrhagic fever virus; DENV: Dengue virus; JEV: Japanese encephalitis virus; KFDV: Kyasanur Forest disease virus; KUNV: Kunjin virus; MVEV: Murray Valley encephalitis; OHFV: Omsk hemorrhagic fever virus; SLEV: St Louis encephalitis virus; TBEV: Tick-borne encephalitis virus; USUV: Usutu virus; WNV: West Nile virus; YFV: Yellow fever virus.
Data from [203,204].

Rapid Molecular Detection of Tuberculosis and Rifampin Resistance

Catharina C. Boehme, M.D., Pamela Nabeta, M.D., Doris Hillemann, Ph.D., Mark P. Nicol, Ph.D.,
Shubhada Shenai, Ph.D., Fiorella Krapp, M.D., Jenny Allen, B.Tech., Rasim Tahirli, M.D., Robert Blakemore, B.S.,
Roxana Rustomjee, M.D., Ph.D., Ana Milovic, M.S., Martin Jones, Ph.D., Sean M. O'Brien, Ph.D.,
David H. Persing, M.D., Ph.D., Sabine Ruesch-Gerdes, M.D., Eduardo Gotuzzo, M.D., Camilla Rodrigues, M.D.,
David Alland, M.D., and Mark D. Perkins, M.D.

Figure 9.

MDR Tuberculosis — Critical Steps for Prevention and Control

Eva Nathanson, M.Sc., Paul Nunn, F.R.C.P., Mukund Uplekar, M.D.,
Katherine Floyd, Ph.D., Ernesto Jaramillo, M.D., Ph.D., Knut Lönnroth, M.D., Ph.D.,
Diana Weil, M.Sc., and Mario Raviglione, M.D.

Legend:
- 0 to <3%
- 3 to <6%
- 6 to <12%
- 12 to <18%
- ≥18%
- No data available

Figure 1. Distribution of the Proportion of Cases of MDR Tuberculosis among New Cases of Tuberculosis, 1994–2009.
The following 27 countries are responsible for 85% of the world's estimated cases of MDR tuberculosis and are classified as countries with a high burden of MDR tuberculosis: China, India, Russia, Pakistan, Bangladesh, South Africa, Ukraine, Indonesia, Philippines, Nigeria, Uzbekistan, Democratic Republic of Congo, Kazakhstan, Vietnam, Ethiopia, Myanmar, Tajikistan, Azerbaijan, Moldova, Kyrgyzstan, Belarus, Georgia, Bulgaria, Lithuania, Armenia, Latvia, and Estonia. Adapted from the 2010 report on MDR and XDR tuberculosis from the WHO.[1]

1. Sputum liquefaction and inactivation with 2:1 sample reagent
2. Transfer of 2 ml material into test cartridge
3. Cartridge inserted into MTB-RIF test platform (end of hands-on work)
4. Sample automatically filtered and washed
5. Ultrasonic lysis of filter-captured organisms to release DNA
6. DNA molecules mixed with dry PCR reagents
7. Seminested real-time amplification and detection in integrated reaction tube
8. Printable test result

Assay Name MTB-RIF
Test Result MTB DETECTED LOW; RIF Resistance NOT DETECTED

Time to result, 1 hour 45 minutes

Figure 2. Assay Procedure for the MTB/RIF Test.
Two volumes of sample treatment reagent are added to each volume of sputum. The mixture is shaken, incubated at room temperature for 15 minutes, and shaken again. Next, a sample of 2 to 3 ml is transferred to the test cartridge, which is then loaded into the instrument. All subsequent steps occur automatically. The user is provided with a printable test result, such as "MTB detected; RIF resistance not detected." PCR denotes polymerase chain reaction.

The Origin of the Haitian Cholera Outbreak Strain

Chen-Shan Chin, Ph.D., Jon Sorenson, Ph.D., Jason B. Harris, M.D.,
William P. Robins, Ph.D., Richelle C. Charles, M.D., Roger R. Jean-Charles, M.D.,
James Bullard, Ph.D., Dale R. Webster, Ph.D., Andrew Kasarskis, Ph.D.,
Paul Peluso, Ph.D., Ellen E. Paxinos, Ph.D., Yoshiharu Yamaichi, Ph.D.,
Stephen B. Calderwood, M.D., John J. Mekalanos, Ph.D., Eric E. Schadt, Ph.D.,
and Matthew K. Waldor, M.D., Ph.D.

A

H1 (Haiti 2010)
H2 (Haiti 2010)
CIRS 101 Bangladesh 2002
M4 (MDC126 Bangladesh 2008)
MJ-1236 Bangladesh 1994
B33 Mozambique 2004
MO10 India 1992
RC9 Kenya 1985
N16961 Bangladesh 1971
N5 (N16961 Bangladesh 1971)
C6 (C6706 Peru 1991)
BX330286 Australia 1986
NCTC 8457 Saudi Arabia 1910
MAK757 Celebes Islands 1937
2740-80 United States 1980
O395 India 1965
V52 Sudan 1968
1587 Peru 1994
62-339 Bangladesh 2002
TMA21 Brazil 1982
AM19226 Bangladesh 2001
MZO-2 Bangladesh 2001
MZO-3 Bangladesh 2001
12129(1) Australia 1985
TM11079-80 Brazil 1980
VS1 United States 1987
VL426 UK Unknown
RC385 Chesapeake Bay 1998

B

Group VI
pre-7th Pandemic
Group I
Group II
Group III
Group IV
Group V

A

C6:Peru 1991
N5:Resequencing of N16961, Bangladesh 1971
M4:Bangladesh 2008
H1:Haiti 2010
N16961 ORFs

Position
350000 360000 370000 380000 390000 400000 410000 420000

B

C6:C6706 Peru 1991
N5:Resequencing of N16961, Bangladesh 1971
M4:MDC126 Bangladesh 2008
H1:Haiti 2010
ORFs in MJ1236

Position
2960000 2980000 3000000 3020000 3040000 3060000

C

C6 (Peru 1991)
N5 (resequencing of N16961, Bangladesh 1971)
M4 (Bangladesh 2008)   c.115T→C, p.Y39H
c.203T→C, p.I68T
CIRS101 (Bangladesh 2002)
H1 (Haiti 2010)   c.58C→A, p.H20N
ctxB

Position
1566900 1567000 1567100 1567200 1567300 1567400

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Medicine 2.0

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Optimized drug
efficacy + toxicity

Theranostics
and patient
stratification

Optimized
Nutrition

Global Systems Biology

Metabonomics

Genomics   Proteomics

Microbiome   Xeno-metabolome

Patient group

Drug toxic but
beneficial

Drug toxic but
NOT beneficial

Drug NOT toxic and
NOT beneficial

Same diagnosis,
same prescription

Drug NOT toxic
and beneficial

Textbook of
Personalized
Medicine

Protein Interaction Networks
Pathways, Networks

Physiology & Pathology, Genomics
Proteomics, Metabolism, Toxicology
Pharmacogenomics, Chemistry...

Chemist   Computer Scientist

Mathematician   Physicist

Modeller

Biologist   Engineer

Model Production

Model Evaluation
and Testing

Hypotheses
Generated?

Experimental
Data Generation

Tumors responsive to chemotherapy

Tumors resistant to chemotherapy

Frozen tissue

Microarrays

Identification of molecular markers of chemosensitivity or resistance to chemotherapy

Samples

Genes

Development and application of quantitative RT-PCR assays

Paraffin-embedded tissue sections

RNA

Initial validation and model refinement

Kaplan–Meier curve

Sensitive

Resistant

Survival probability

1.0
0.8
0.6
0.4
0.2
0.0

0   2   4   6   8   10

Overall survival (yr)

Further validation in larger patient cohorts

Mature diagnostic tool

Table 1. Human cancers caused by viruses and their potential targets for RNA interference

Virus	Cancer	Viral target
Epstein–Barr virus	Burkitt's lymphoma	EBNA-1
	Hodgkin's disease	EBNA-2
	Post-transplant lymphoproliferative disorder	LMP1
	AIDS-related central nervous system lymphoma	
Human papillomavirus	Anogenital cancers	E6 E7
Human T cell leukaemia virus-1	Adult T-cell leukaemia	Tax
Kaposi's sarcoma-associated herpes-virus	Kaposi's sarcoma	LANA
	Multicentric Castleman's disease	vcyclin
	Primary effusion lymphoma	VFLIP
		vIL-6/ORF50
Hepatitis B or C viruses	Hepatocellular carcinoma	Hepatitis B surface or core antigen
		HBx potential oncogene
		Hepatitis C virus viral proteases

AIDS	Acquired Immune Deficiency Syndrome
EBNA-1/ EBNA-2	Epstein–Barr virus Nuclear Antigen 1/2
LMP1	Latent Membrane Protein 1
LANA	Latency Associated Nuclear Antigen
ORF50	Open Reading Frame 50
Tax HTLV-1	Transactivator
vFLIP	viral FLICE inhibitory protein
vIL-6	viral interleukin 6

PHARMACIENS SANS FRONTIERES

LES JIVAROS

Apothicaires en Amazonie

Malgré une insertion de plus en plus grande dans l'économie marchande, les Jivaros n'ont jusqu'à récemment disposé d'aucun service de santé, même modeste. Par Emmanuel Thévenon

La pharmacopée
des chimpanzés

Sabrina KRIEF

L'observation des grands singes indique l'existence de comportements d'automédication. Les plantes que ces animaux sélectionnent pour « se soigner » deviendront-elles nos médicaments de demain ?

1 : LES CHIMPANZÉS CONSOMMENT DES PLANTES ayant des propriétés antiintestinales, comme les fruits de *Phytolacca dodecandra*. Il pourrait les associer à d'autres substances pour éviter de pâtir de leur toxicité.

Will you get sick?
Antibodies could foretell the future of your health
# Predicting Disease

1. Heart Line
2. Head Line
3. Life Line
4. Fate Line
5. Marriage Line
6. Travel Line
7. Fame Line
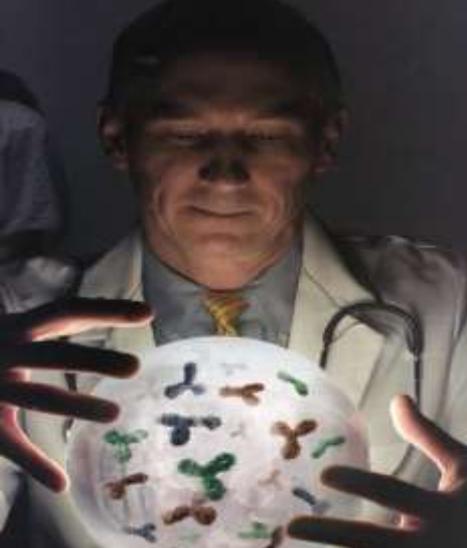
CHECKUPS OF THE FUTURE

Someday physicals could routinely include screening for autoantibodies.

Autoantigen — Autoantibody

Signal

Detail of chip segment

Blood plasma

Blood sample

1 A blood sample would be given to a laboratory, which would extract the plasma (the acellular part).

2 The plasma would be washed over a chip containing an array of autoantigens—molecules known to be capable of eliciting autoimmune reactions—at known positions. Any autoantibodies in the blood would bind to the autoantigens, triggering signals indicating the identity and quantity of the bound autoantibodies.

3 This information would be translated into a prediction of the patient's risk for becoming afflicted with specific conditions.

Chip

Patient #35622b

Disorder	Five-year risk
Type 1 diabetes	70%
Rheumatoid arthritis	No sign
Multiple sclerosis	No sign
Lupus	No sign
Thyroid disease	15%

PREDICTIVE MEDICINE
A Study in Strategy...
A Plan for Maintaining Health by Avoiding Disease

E. Cheraskin, M.D., D.M.D.
W. M. Ringsdorf, Jr., D.M.D.

Stefan Hiermaier
Editor

Predictive Modeling of Dynamic Processes
A Tribute to Klaus Thoma

Springer

Genetic variation in *IL28B* and spontaneous clearance of hepatitis C virus

David L. Thomas[1]*, Chloe L. Thio[1]*, Maureen P. Martin[2]*, Ying Qi[2], Dongliang Ge[3], Colm O'hUigin[2], Judith Kidd[4], Kenneth Kidd[4], Salim I. Khakoo[5], Graeme Alexander[6], James J. Goedert[7], Gregory D. Kirk[8], Sharyne M. Donfield[9], Hugo R. Rosen[10], Leslie H. Tobler[11], Michael P. Busch[11], John G. McHutchison[12], David B. Goldstein[3] & Mary Carrington[2,13]

Hepatitis C virus gets personal

Shawn P. Iadonato and Michael G. Katze

Many people infected with the hepatitis C virus are not cured despite gruelling therapy. A human genetic variant that predicts successful treatment has been identified. So is personalized therapy now a possibility?
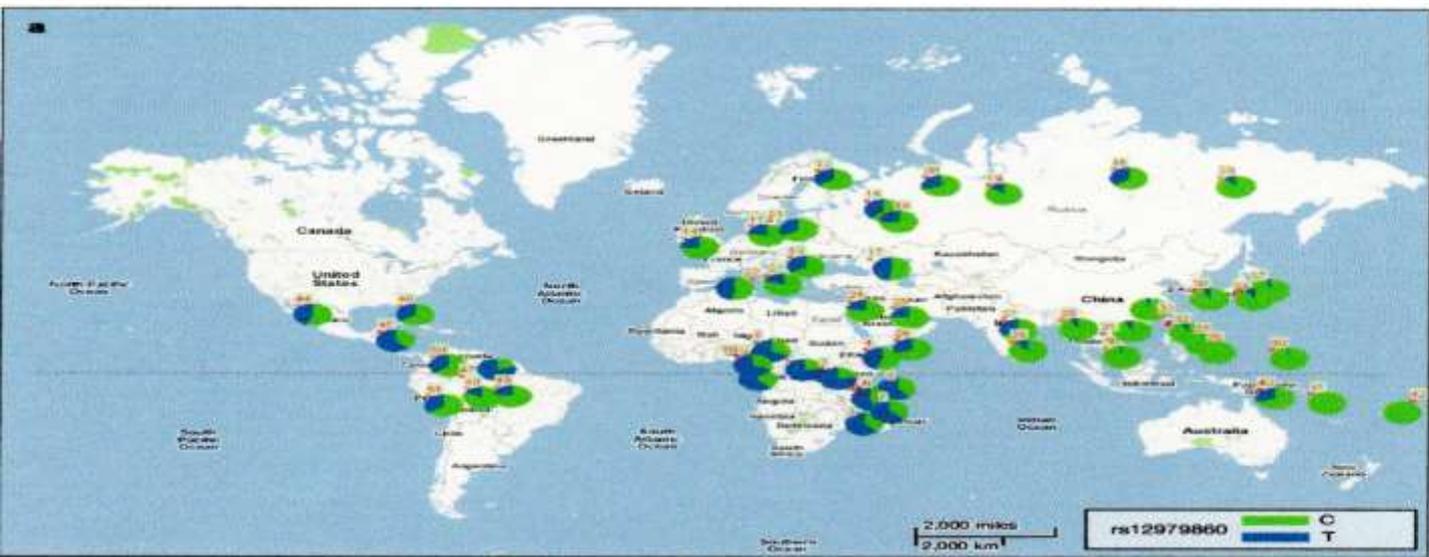
Figure 2 | Sampling locations, allele frequencies and degree of regional differentiation of the rs12979860 C allele. a, The numbers identifying populations are given in Table 3. The pie charts show the frequency of the C (green) and T (blue) alleles in each population sampled. **b,** Frequency distribution of F_{ST} values for 1,062 SNPs from 32 of the samples grouped into 6 regions (Africa, Europe, south Asia, southeast Asia, east Asia, Oceania). The red arrow indicates the position of the estimated F_{ST} for rs12979860.
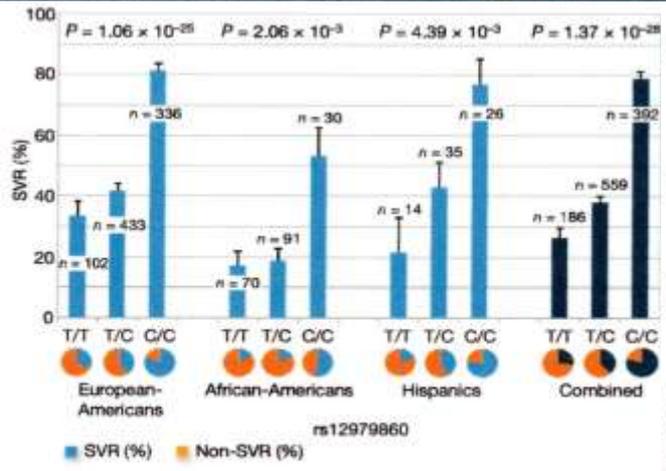

**Figure 1 | Percentage of SVR by genotypes of rs12979860.** Data are percentages + s.e.m.
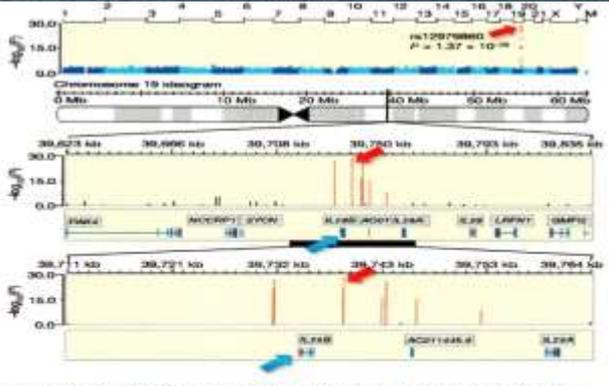
Figure 2 | Genomic overview of the region of 19q13.13 surrounding the genome-wide significant determinant of response to treatment and including the *IL28B* gene. The top panel shows a genome-wide view of the P values ($-\log_{10}(P)$). Panels below show all genotyped SNPs in the region of significance and the structures of the surrounding genes. The SNPs that show genome-wide significant association with SVR are marked in red. The polymorphism rs12979860 (red arrow) is 3 kb upstream to the gene encoding IFN-λ-3 (*IL28B*, blue arrow). Other SNPs in the same region showing genome-wide significant P values largely reflect the same signal (Supplementary Information IX). The results were annotated using the WGAViewer software[n].
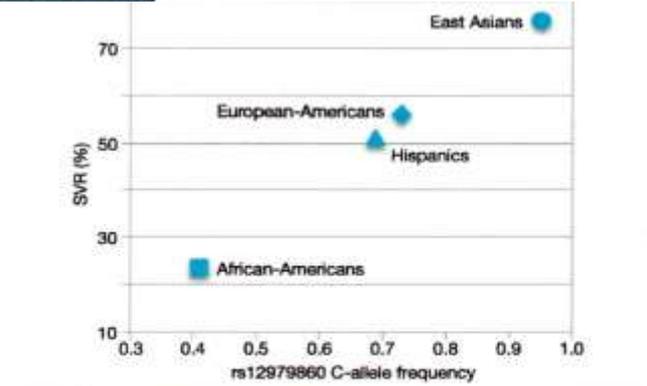

Figure 3 | Rate of SVR and rs12979860 C-allele frequency in diverse ethnic groups. The SVR rate in East Asians is adopted from Liu *et al.*[7]. Sample sizes for C-allele frequency: $n = 61$ (African-Americans); $n = 271$ (European-Americans); $n = 16$ (Hispanics); $n = 107$ (East Asians); sample sizes for SVR rate: $n = 191$ (African-Americans); $n = 871$ (European-Americans); $n = 75$ (Hispanics); $n = 154$ (East Asians).

XENOME

Xenotransplant

XENOTRANSPLANTATION
BASIC RESEARCH AND
CLINICAL APPLICATIONS

EDITED BY
JEFFREY L. PLATT, MD

Humana Press

SO HAVE YOU FILLED OUT YOUR ORGAN DONOR CARD YET??

Xeno-transplantation
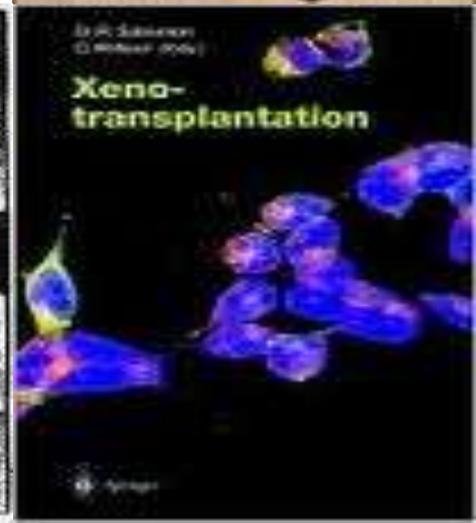
As Questões da Xenotransplantação e das Doenças Oncológicas

Detection of a Gammaretrovirus, XMRV, in the Human Population: Open Questions and Implications for Xenotransplantation

Joachim Denner

Abstract and Introduction

Abstract

XMRV (xenotropic murine leukaemia virus-related virus) is a gammaretrovirus that has been detected in human patients with prostate carcinoma, chronic fatigue syndrome (CFS) and also in a small percentage of clinically healthy individuals. It is not yet clear whether the distribution of this virus is primarily limited to the USA or whether it is causally associated with human disease. If future investigations confirm a broad distribution of XMRV and its association with disease, this would have an impact on xenotransplantation of porcine tissues and organs. Xenotransplantation is currently being developed to compensate for the increasing shortage of human material for the treatment of tissue and organ failure but could result in the transmission of porcine pathogens. Maintenance of pathogen-free donor animals will dramatically reduce this risk, but some of the porcine endogenous retroviruses (PERVs) found in the genome of all pigs, can produce infectious virus and infect cultured human cells. PERVs are closely related to XMRV so it is critical to develop tests that discriminate between them. Since recombination can occur between viruses, and recombinants can exhibit synergism, recipients should be tested for XMRV before xenotransplantation.

Questions Concerning XMRV Detection

Medscape

5´ LTR	gag	pol	env	3´ LTR
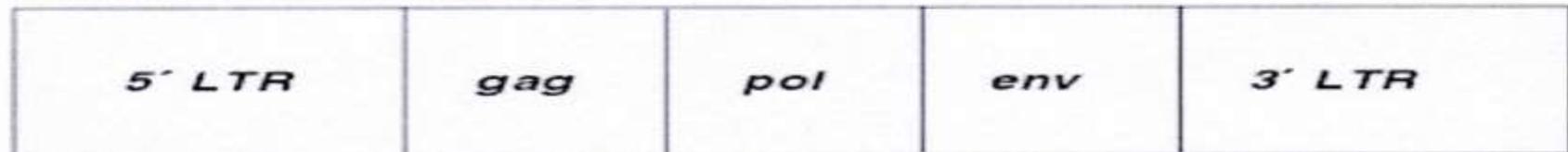
Source: Br J Dermatol © 2009 Blackwell Publishing

Figure 1. Genomic structure of a human endogenous retrovirus. LTR, long terminal repeat.
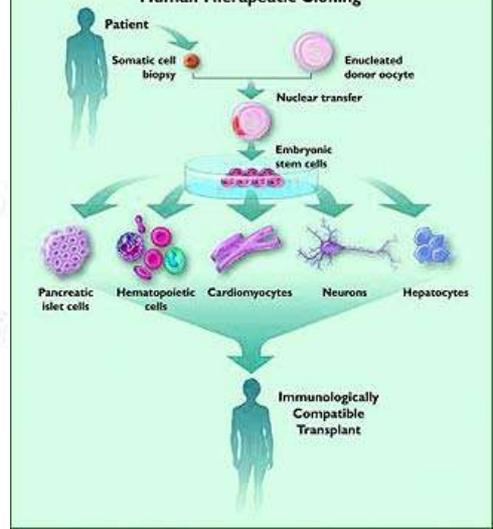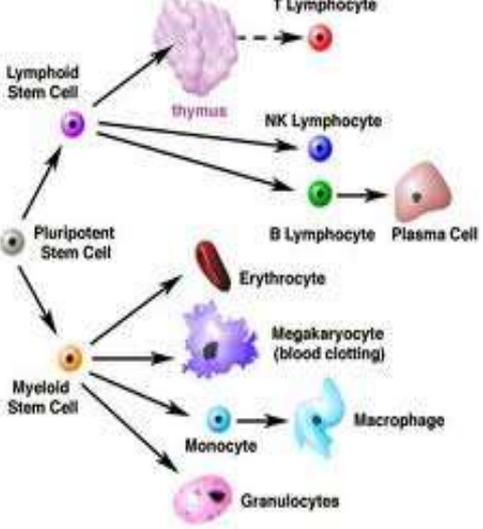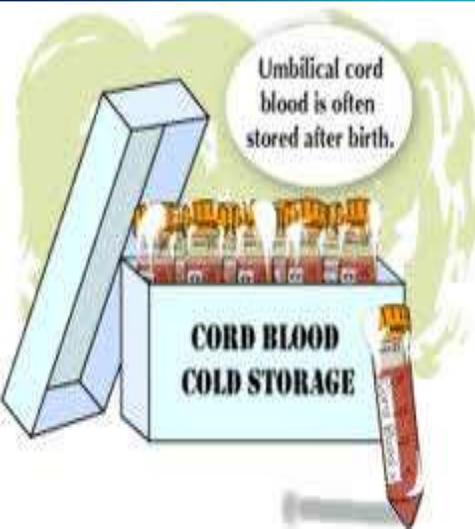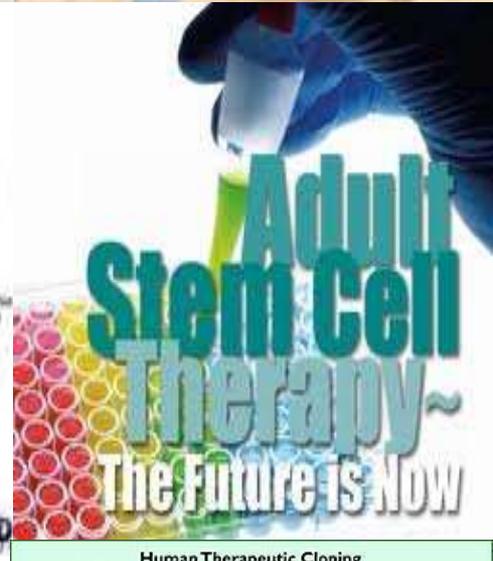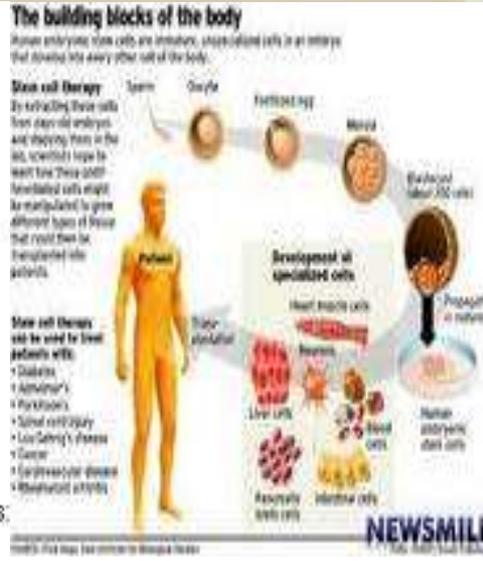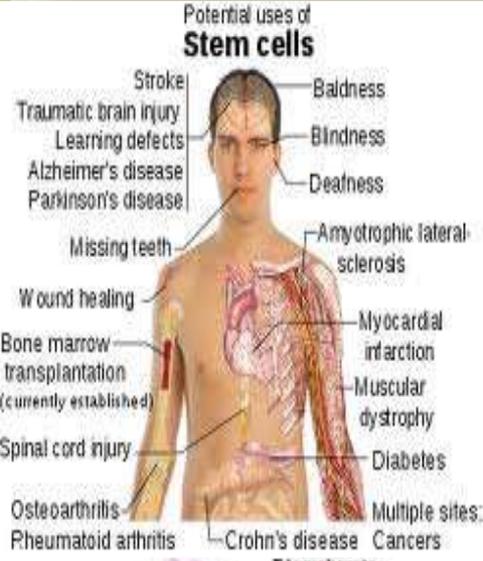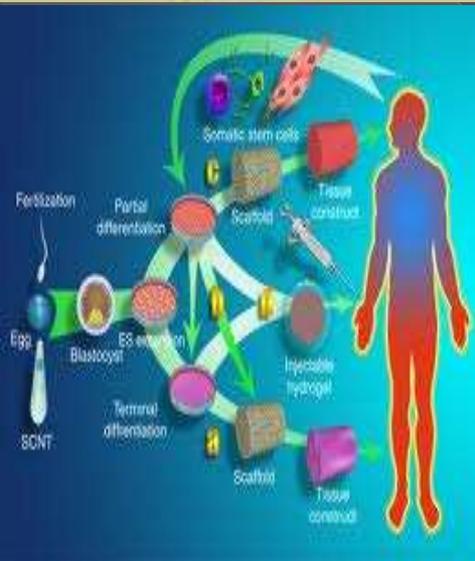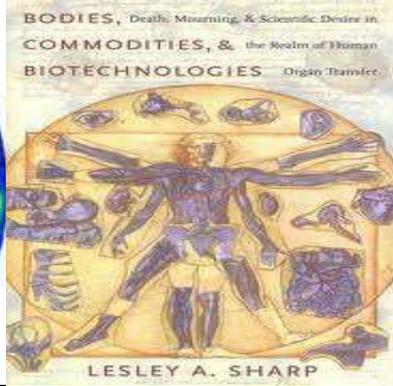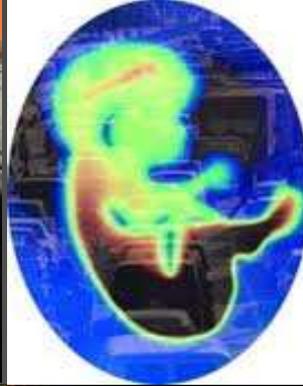
THE KEY TO ETERNAL YOUTH? FIBROBLASTS

BY LESLIE BAUMANN, MD

I WANT TO LIVE FOREVER

I DO NOT WANT TO DIE

Eternal Youth

Eternal Youth
still you, just better

Eternal Youth
beauty salon

SPACE AND ETERNAL LIFE

A Dialogue between
Chandra Wickramasinghe
and Daisaku Ikeda

Permanent youth

JOHN SHELBY SPONG
BESTSELLING AUTHOR OF
JESUS FOR THE NON-RELIGIOUS

ETERNAL LIFE:
A NEW VISION

BEYOND RELIGION,
BEYOND THEISM,
BEYOND HEAVEN AND HELL

YOUR OWN PERFECT MEDICINE

by Martha M. Christy

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Proven Natural
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INTERNAL HEALTH
The Key to Eternal
YOUTH & VITALITY
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CRITICAL LINK BETWEEN DISCOVERY & PATIENTS

Science Translational Medicine
Online issue 14 April 2010

Science Translational Medicine
Online issue 14 October 2009

SCIENTIFIC AMERICAN REPORTS
YOUR FUTURE WITH **ROBOTS**
How Smart Machines Will Change Everything

SCIENTIFIC AMERICAN
CANCER-KILLING VIRUSES · DANGEROUS MELTING IN THE ARCTIC
ARTIFICIAL MUSCLES
Shape-Shifting Plastics Replace Motors in Machines That Move

Figure 1 | The gap between genetic information and medical innovation. Although the amount of genetic and genomic information available to researchers is expanding rapidly, advances in medical innovation have not followed suit. The graph shows the size of the GenBank database (a collection of all publicly available DNA sequences) between 1982 and 2008 (purple); the number of submissions of human single nucleotide polymorphisms (SNPs) to the SNP database (dsSNP) of the National Center for Biotechnology Information (NCBI) between 2002 and 2009 (blue); the number of Investigational New Drug (IND) applications received by the Center for Drug Evaluation and Research/US Food and Drug Administration (CDER/FDA) during 1986–2008 (red); and the number of new molecular entities (drugs not previously marketed in the United States in any form) approved by the CDER/FDA between 1982 and 2008 (green). Data are compiled from the NCBI and FDA websites.

bacterium

chromosome   plasmid

plant cell or animal cell

A.

plasmid isolated from bacterium

DNA isolated from donor cell

B.

the same restriction enzyme that fragmented donor DNA is also used to fragment plasmid DNA

a specific restriction enzyme fragments donor DNA

each plasmid incorporates a *different* DNA fragment

engineered plasmid (recombinant DNA) is introduced into a bacterium

C.

donor and plasmid DNA are mixed; "sticky ends" of donor DNA form hydrogen bonds with sticky ends of plasmid DNA fragment and are sealed with another specific enzyme

bacteria reproduce and clone the gene from donor cell that was spliced into plasmid

Edible Vaccines

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Scientists Dish Up Rice Vaccine to Fight Cholera

Cholera toxin implanted in rice provokes strong immunity in mice while surviving the rigors of acid digestion

By David Biello | June 11, 2007
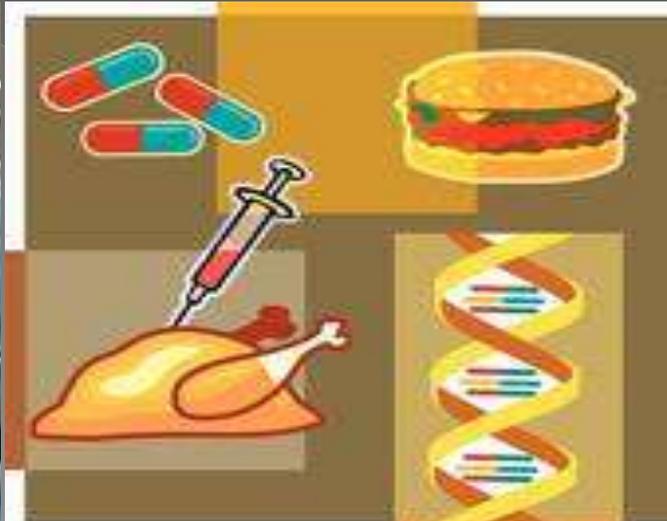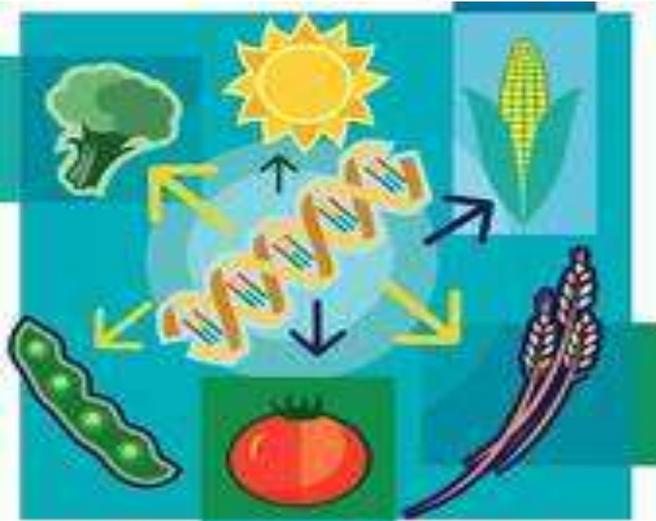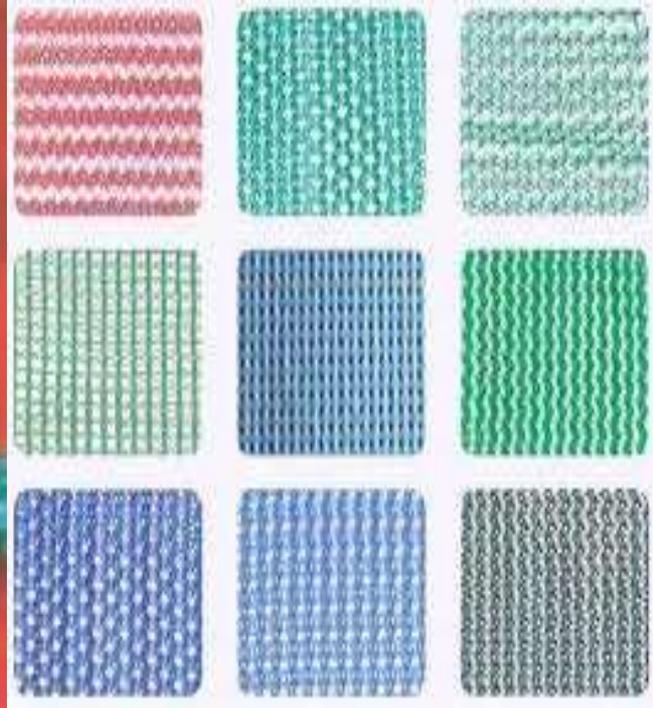
Clinical Spectrum of WNV Illness: Revised

WN Meningitis

WN Fever

WN Encephalitis

WN "Poliomyelitis"

Inflammatory Neuropathy
Radiculopathy / plexopathy

Safety concerns about CCR5 as an antiviral target
Amalio Telenti

Inhibitors of HIV-1 entry

SARS

Figure 2. Zoonotic transmission of SARS-CoV. Genomic analyses provided evidence that genetic changes in the spike gene of SARS-CoV from bats (left) and civet cats (center) are essential for the animal-to-human transmission (horizontal arrows). Species-to-species genetic variation in the (thus far unidentified) viral receptor in bats and in the *angiotensin converting enzyme 2 (ACE2)* gene, encoding the SARS-CoV receptor in civet cats and humans also affects the efficiency with which the virus can enter cells (vertical arrows). The SARS-CoV that caused the epidemic evolved a high affinity for both civet (center) and human (right) ACE2 receptors (indicated by the single diagonal and the right side vertical arrow). Image credit: Bart Haagmans, Erasmus MC. Original images (left to right) by Dodoni, Paul Hilton, and Hoang Dinh Nam.
doi:10.1371/journal.ppat.1000557.g002

Recycling of chloroquine and its hydroxyl analogue to face bacterial, fungal and viral infections in the 21st century

Jean-Marc Rolain*, Philippe Colson, Didier Raoult

Unité des Rickettsies, CNRS UMR 6020, Université de la Méditerranée, Faculté de Médecine et de Pharmacie, 27 Boulevard Jean Moulin, 13385 Marseille Cedex 5, France

Abstract

Chloroquine (CQ) and its hydroxyl analogue hydroxychloroquine (HCQ) are weak bases with a half-century long use as antimalarial agents. Apart from this antimalarial activity, CQ and HCQ have gained interest in the field of other infectious diseases. One of the most interesting mechanisms of action is that CQ leads to alkalinisation of acid vesicles that inhibit the growth of several intracellular bacteria and fungi. The proof of concept of this effect was first used to restore intracellular pH allowing antibiotic efficacy for Coxiella burnetii, the agent of Q fever, and doxycycline plus HCQ is now the reference treatment for chronic Q fever. There is also strong evidence of a similar effect on viruses against Tropheryma whipplei, the agent of Whipple's disease, and a clinical trial is in progress. Other bacteria and fungi multiply in an acidic environment and encouraging in vitro data suggest that this concept may be generated for all intracellular organisms that multiply in an acidic environment. For viruses, CQ led to inhibition of uncoating and/or alteration of post-translational modifications of newly synthesised proteins, especially inhibition of glycosylation. These effects have been well described in vitro for many viruses, with human immunodeficiency virus (HIV) being the most studied. Preliminary in vivo clinical trials suggest that CQ alone or in combination with antiretroviral drugs might represent an interesting way to treat HIV infection. In conclusion, our review re-emphasises the paradigm that activities mediated by lysosomotropic agents may offer an interesting weapon to face present and future infectious diseases worldwide.
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Keywords: Chloroquine; Hydroxychloroquine; Lysosomotropic agents; Alkalinisation; Q fever; Whipple's disease; HIV

Table 1

Bacteria, fungi and viruses inhibited by chloroquine and/or hydroxychloroquine (in vitro data)

Bacteria	Reference	Fungi	Reference	Virus
Coxiella burnetii	[5,13]	Histoplasma capsulatum	[24]	HIV
Tropheryma whipplei	[7,8]	Cryptococcus neoformans	[15,25]	SARS-CoV
Legionella pneumophila	[11]	Paracoccidioides brasiliensis	[26]	Influenza viruses
Francisella tularensis	[12]	Penicillium marneffei	[15,27]	Flavivirus, including yellow fever virus
Mycobacterium tuberculosis	[14]	Aspergillus fumigatus	[28]	Rubella virus
Mycobacterium avium	[15]			HAV
Salmonella Typhi	[16]			HBV
Escherichia coli	[17]			HCV
Bacillus anthracis	[18]			Arenavirus
Bacillus subtilis	[19]			
Borrelia burgdorferi	[20]			Lymphocytic choriomeningitis virus
Brucella abortus	[21]			Rabies virus
Staphylococcus aureus	[22]			Varicella–Zoster virus
Listeria monocytogenes	[23]			Respiratory syncytial virus
				Sindbis virus
				Herpes simplex viruses
				Epstein–Barr virus
				Polioviruses
				Newcastle disease virus
				Borna disease virus
				Vesicular stomatitis virus
				Vaccinia virus
				Murine RNA tumour virus
				FMDV
				Mayaro virus
				Feline calicivirus
				African swine fever virus
				Bovine leukaemia virus
				Canine parvovirus
				Minute Virus of Mice

Effects of chloroquine on viral infections: an old drug against today's diseases?

Andrea Savarino, Johan R Boelaert, Antonio Cassone, Giancarlo Majori, and Roberto Cauda.

A – pH increase induced by CQ/HCQ

1/ Inhibition of low-pH-dependent conformational changes that triggers fusion, penetration, and/or uncoating within the endosomal pathway

HCV, HAV, Influenza virus, lymphocytic choriomeningitis virus, Foot-and-mouth disease virus, Borna disease virus, Feline caliciviruses, Minute virus of mice, Avian leucosis virus

2/ Inhibition of post-translational modifications of viral proteins
HIV, HSV-1, Flaviviridae, Mayaro virus, Avian reticuloendotheliosis virus

B – Other possible antiviral mechanisms

1/ Immuno-modulatory and anti-inflammatory effects
HIV, SARS-CoV

2/ Iron restriction, decrease of the iron cellular accumulation
HIV
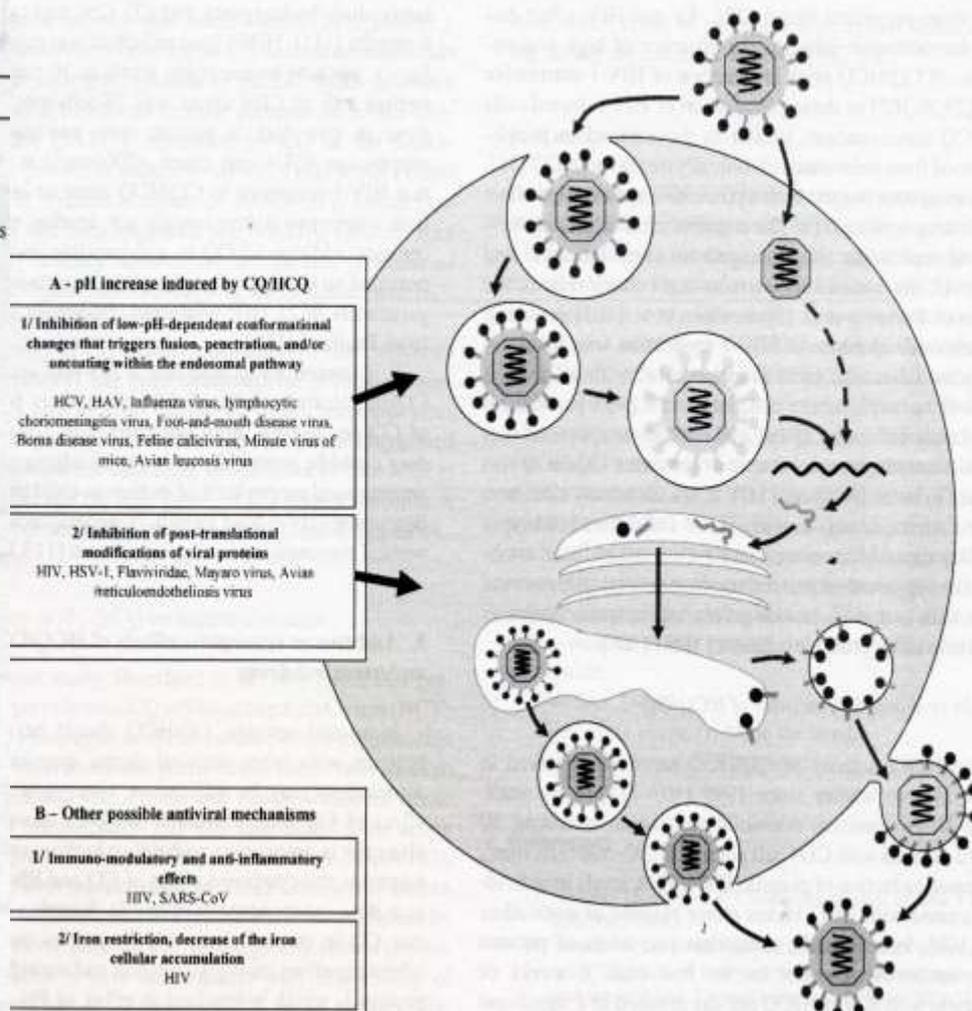
PROBLEMAS DECORRENTES da TECNOLOGIA
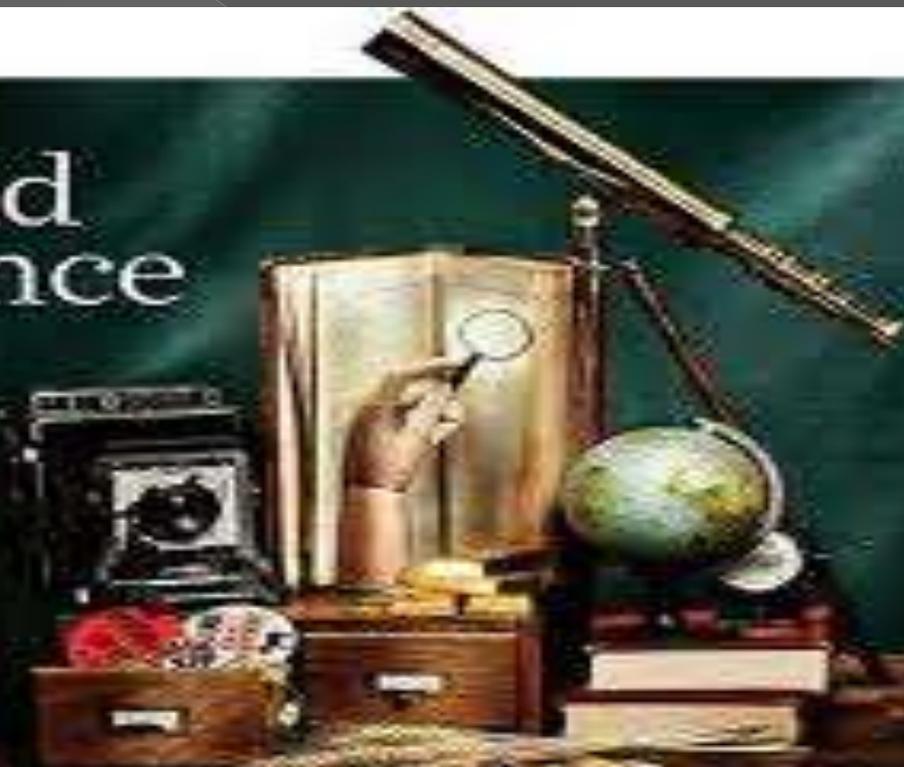
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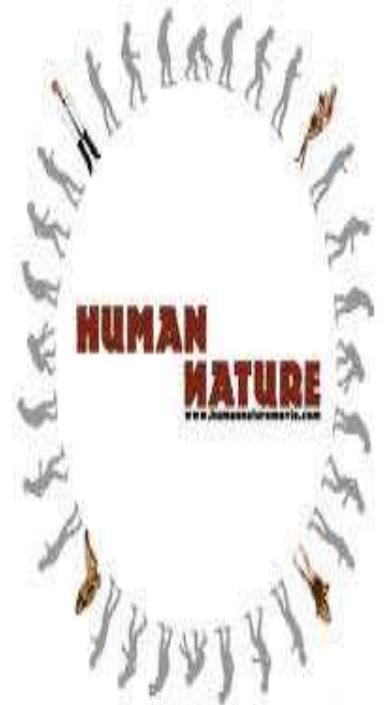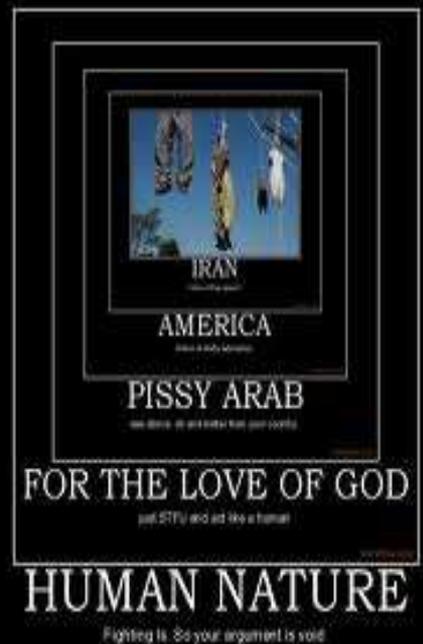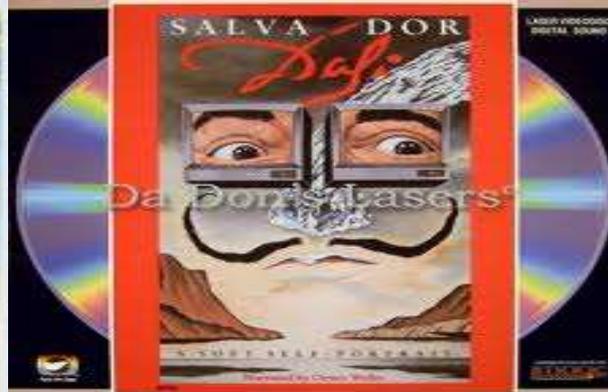
tecnologia

As consultas do dr. Google e da dra. Wikipédia

Se à mínima dor vai a correr para o computador, onde passa horas a pesquisar sintomas tentando chegar a um autodiagnóstico, o mais provável é que sofra de cibercondria, o termo que junta a clássica hipocondria à Internet. Juntas, elas podem dar-lhe cabo da saúde.

Search

| doença da gota	✕	Pesquisar

Pesquisa avançada
Ferramentas de linguagem

doença da **gota**   Sinto-me doente »
doença das **vacas loucas**
doença da **zona**
doença da **estalada**
doença de **crohn**
doença de **parkinson**
doença de **graves**
doença de **huntington**
doença de **behçet**
doença de **buerger**
doença de **bowen**
doença de **burger**
doença de **lyme**
doença de **lupus**
doença de **lou gehrig**
doença de **lobstein**
doença da **mentira**
doença da **membrana hialina**
doença da **moda**
doença do **trabalho**
doença do **tabaco**
doença do **tecido conjuntivo**
doença do **trofoblasto**

ROBOTS AND ROBOTICS
Scientific and Medical Robots
Tony Hyland
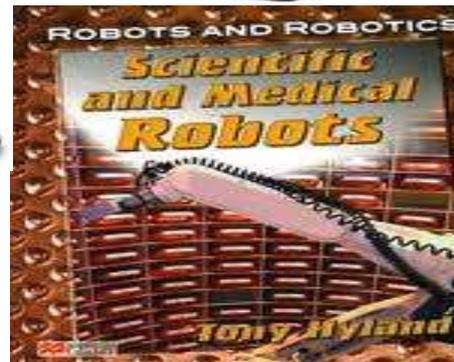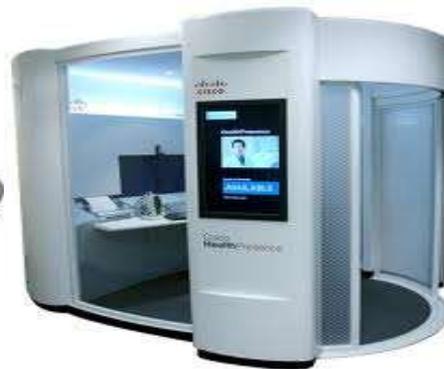
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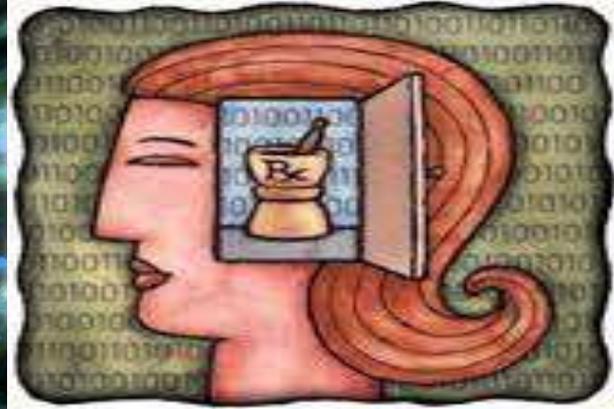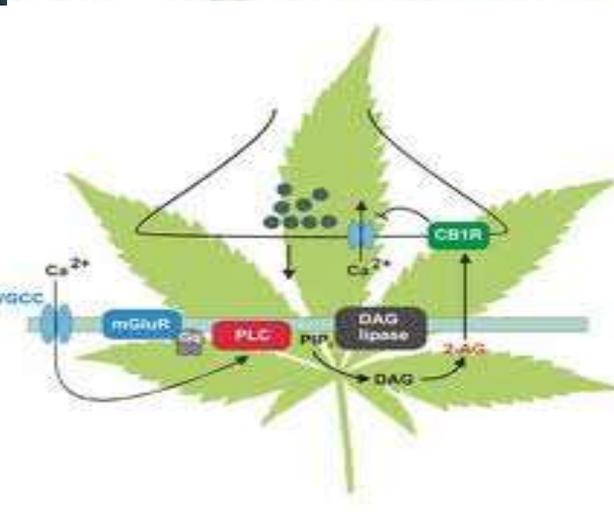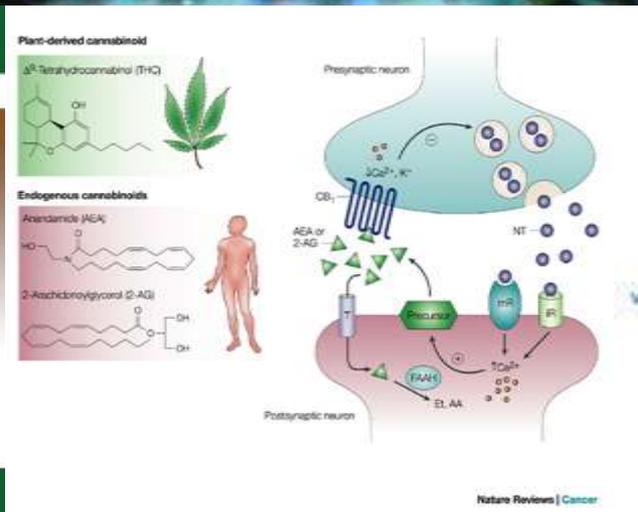
A pathogenic mechanism leading to partial lipodistrophy and prospects for pharmacological treatment of insulin resistance syndrome

Nadir M. Maraldi[1,2], Cristina Capanni[1], Elisabetta Mattioli[1], Marta Columbaro[1], Stefano Squarzoni[2], Weena K. Parnaik[4], Manfred Wehnert[5], Giovanna Lattanzi[2]

[1] Department of Human Anatomy, University of Bologna, Italy, 2 Laboratory of Cell Biology, Istituti Ortopedici Rizzoli, Bologna, Italy, [3] ITOI, CNR, Unit of Bologna, c/o IOR, Bologna, Italy, [4] Centre for Cellular and molecular Biology, Hyderabad, India, [5] Institute of Human Genetics, University of Greifswald, Germany
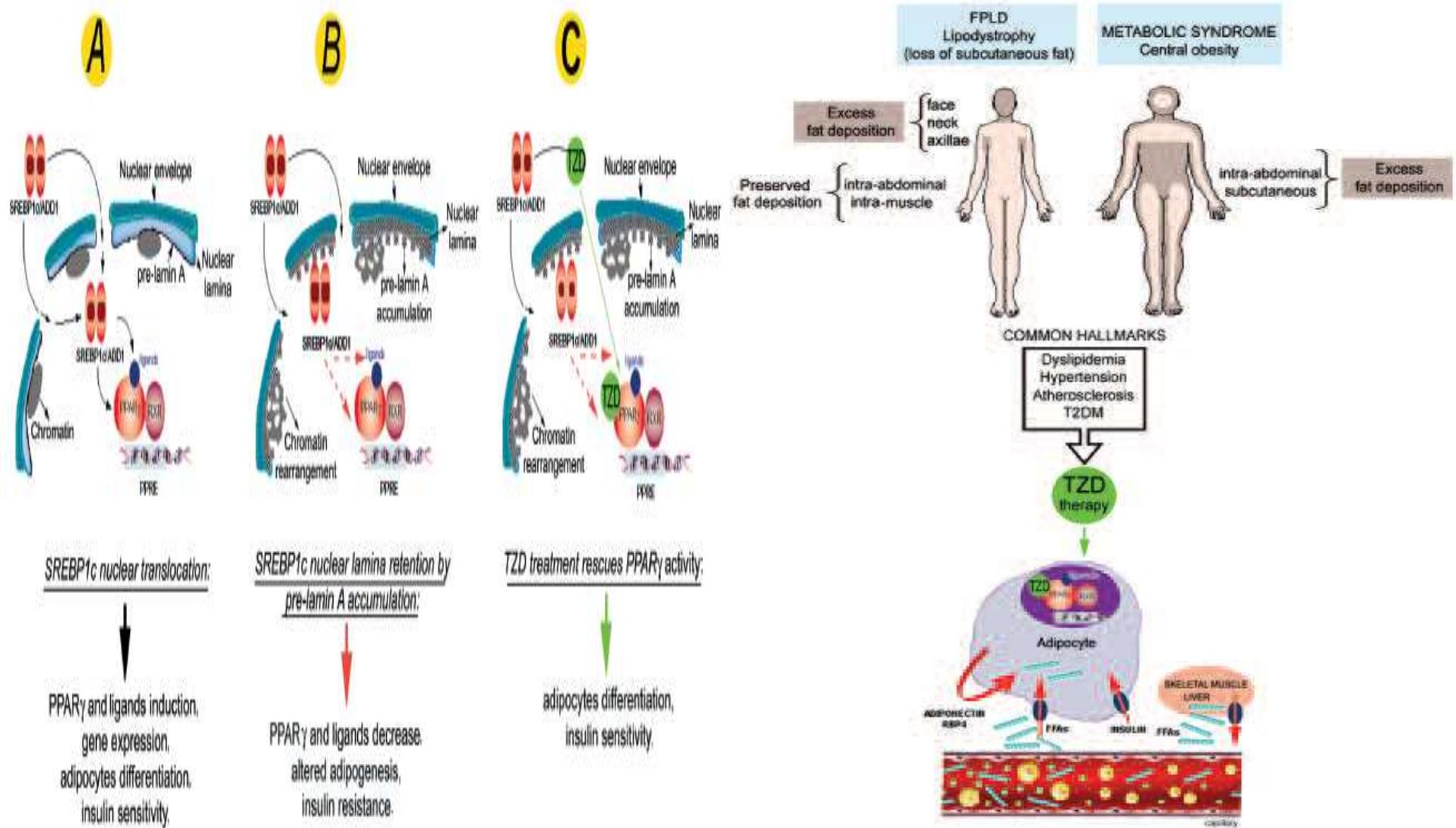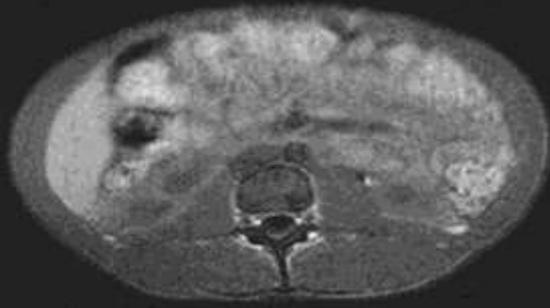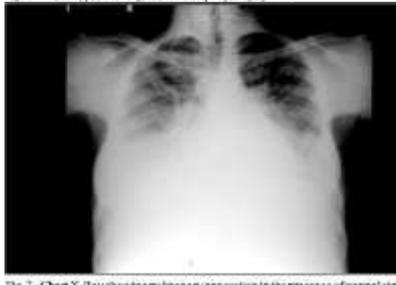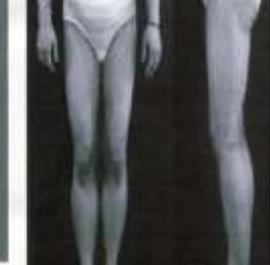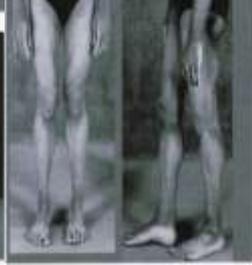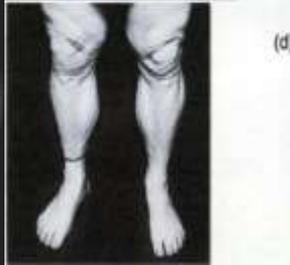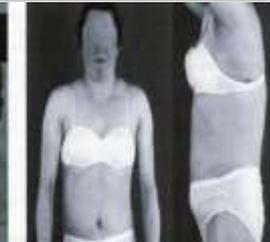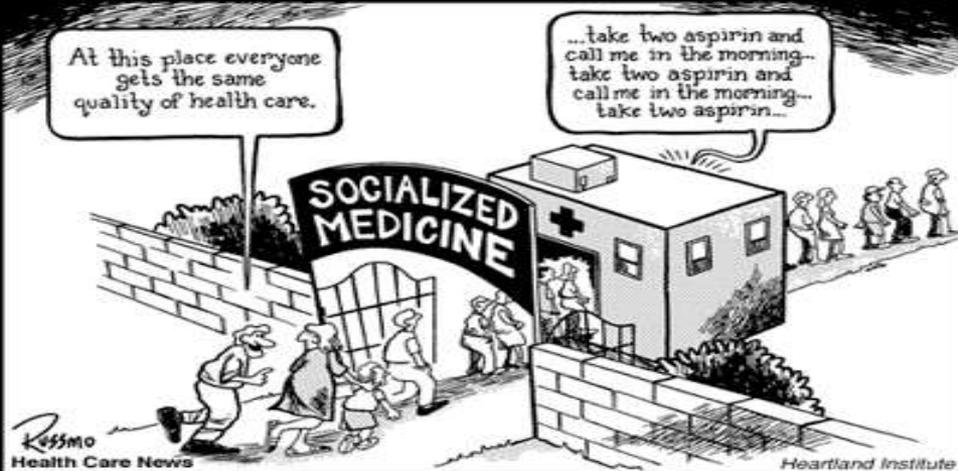
CONCLUSÕES
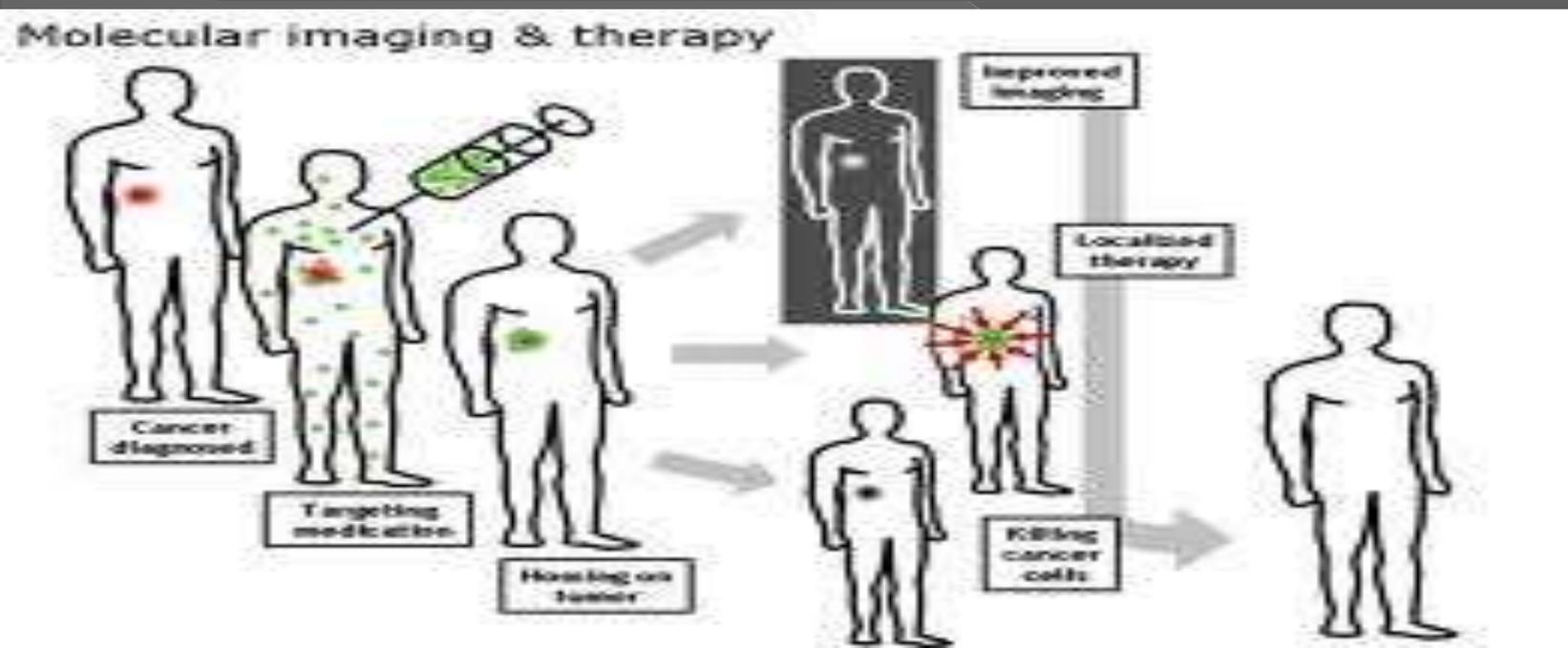
PENSAMENTO

- " ... *só podemos ver o que está mais para a frente, olhando volta e meia para trás ...*" *(sic.)* **(John G. Evans, Clinical Futures: Aging, Book of the 50 th Anniversery of the NHS)**

Editorial

The end of our National Health Service

THE LANCET

Zoom

L'UNIFICATION DES FORCES

Six théories pour fonder la physique

LE BESOIN D'UNE CAUSE FONDAMENTALE

Les dossiers de LA Recherche

La théorie du tout

Comment la physique peut expliquer l'Univers

LA THÉORIE DE TOUT

Tout l'Univers dans une équation

La théorie de tout

« Si nous découvrons une théorie complète, [...] nous connaîtrons la pensée de Dieu »
STEPHEN HAWKING
Astrophysicien

Is the world getting better or worse?

If you want to make the world a better place, it's a good idea to find out where you're starting from. When we did just that, the results came as something of a surprise. By most measures, living standards are improving across the world, and though hundreds of millions of people still live in desperate conditions, things are moving in the right direction. Less encouragingly, the environment is clearly in trouble and armed conflict remains a major problem

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Life expectancy at birth
In years

- World
- Developed countries
- Developing countries
- Sub-Saharan Africa

GDP per capita
US$ per person

Infectious diseases
Worldwide deaths from infectious diseases

80% of DEATHS from INFECTIOUS DISEASES are caused by diarrhoea, HIV/AIDS, malaria, measles, pneumonia and tuberculosis

1993 **16.4 million** deaths from infectious diseases

2002 **14.7 million** deaths from infectious diseases

Infant mortality
Deaths in first year per 1000 live births

- World
- Developed world
- Developing world
- Sub-Saharan Africa

Child survival
Beyond fifth birthday

In sub-Saharan Africa, 1 IN 7 CHILDREN DIE before their fifth birthday

Developed world · Developing world · Sub-Saharan Africa

Maternal mortality
Deaths per 100,000 live births

EVERY YEAR, 536,000 WOMEN and GIRLS die as a result of COMPLICATIONS during pregnancy or childbirth

CO₂ emissions
Increase in worldwide emissions (tonnes)

2003 25.6 bn +7%
2000 23.8 bn +12%
1990 21.3 bn

Ecological footprint
Hectares per person

An ECOLOGICAL FOOTPRINT of 2.1 HECTARES is considered to be a FAIR SHARE of the world's natural resources

US · Western Europe · China · India

Deforestation
Annual net change in forest area (millions of hectares per year)

1990-2000 · 2000-2005

DEFORESTATION continues at the RATE of about 13 MILLION HECTARES per year (roughly equivalent to the land area of Bangladesh)

Developed world · Latin America · SE Asia · Sub-Saharan Africa

Hunger
Percentage of children under 5 underweight

Developing world · South Asia · Sub-Saharan Africa

Malnutrition
Percentage of population undernourished

The DECLINING TREND in undernourishment in developing countries was REVERSED in 2008, largely due to escalating food prices

Developing world · Sub-Saharan Africa

Extreme poverty
Percentage of people living on less than $1.25 a day

As many as 1 BILLION people are likely to still be in EXTREME POVERTY by 2015

Developing world · Sub-Saharan Africa

Food supply
Kilocalories per day

World · Sub-Saharan Africa · North America · Developing world

Water
Percentage of population with access to clean water and sanitation

- World
- Sub-Saharan Africa
- Developing world

1990 · 2004

CLEAN WATER · SANITATION

Education
Percentage enrolment in primary education

World · Developed world · Developing world · Sub-Saharan Africa

War
Number of conflicts, 1945 to 2008

Number of conflicts
1945 · 1965 · 1985 · 2005

Military spending
In billions of dollars

1982 · 2007

Displaced persons
Displaced people and refugees

Internally displaced people
Refugees
2000 · 2002 · 2004 · 2006 · 2008

Natural disasters
As natural disasters rise, the number of victims is stable

1900 · 1905 · 2000 · 2005 · 2008

L'utopie du savoir total

Perspectives

The roots of ancient medicine: an historical outline

B V SUBBARAYAPPA*

Visiting Professor, National Institute of Advanced Studies, Bangalore 560 012, India

(Email, phispc@bgl.vsnl.net.in)

A Review and Integrative Analysis of Ancient Holistic Character Medicine Systems

Søren Ventegodt[1,2,3,4,5*], Suzette Thegler[2,3,5], Tove Andreasen[2,3,5], Flemming Struve[2,3,5], Susan Jacobsen[2,3,5], Margrethe Torp[5], Hans Ægedius[2,3,5], Lars Enevoldsen[2,3,5], and Joav Merrick[6,7,8]

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E-mail: ventegodt@livskvalitet.org

The ancient holistic medical systems help the patient by balancing the "elements" of the human character. This work aims to understand the nature of these elements and the process of the physician balancing them. Using the concept of poly-ray cosmology we see that the medical systems from ancient India, China and Greek basically share the same inner structure and also the same logic of the treatment processes. We analyze the double concept of yin-yang, the Ayurvedic triadic concepts of Pitta, Kapha and Vata, the four elements of Hippocratic humoral medicine, and the five elements of Chinese medicine, and find that each of these conceptual frameworks make up a "theory" or model of the world that is a perfect wholeness, allowing the physician to interpret the world and his patient in order to identify the imbalances of his or her character that need to be treated. Independently of the system this can be a palliative treatment, if energies are only balanced in present time, or a causal cure if the physician is using the similarity principle to take his patient into regression back to the events in the personal history that originally created the imbalances (the traumas). To help the patient back to the traumas he is exposed to a small dose of the original harming stimulus; this can be an internal process like visualization supported by the therapist words, or an external process provoked by his actions. If the physician is balancing the elements without such a healing of the patient's existential core this can still momentarily help the patient by alleviating the symptoms, but it will not have a permanent effect.

2 elements		Ayurveda – 3 elements			Hippocratic – 4 elements				Chinese – 5 elements				
Yang	Yin	Pitta	Kapha	Vata	Earth	Water	Air	Fire	Water	Wood	Fire	Earth	Metal
Hot	Cold	Oily	Wet	Dry	Firm	Floating	Easy Inspiration	Transformation	Salty	Rage	Joy	Worry	Sadness
Dry	Wet	Spicy	Cold	Light	Heavy	Dynamic	Light	Warm	Winter	Sour	Growth	Sweet	Spicy
Repletion	Vacuity	Hot	Heavy	Cold	Steady	Emotion	Spirit	Destructive	Cold	Spring	Heat	Change	Fall
External	Internal	Light	Dull	Rough	Dark	Persevering	Communication	Masculine	Death	Wind	Summer	Yellow	Regression
Light	Dark	Pungent	Slimy	Subtle	Hard	Feminine		Creative	Black	Birth	Red	Dampness	White
Above	Below	Mobile	Soft	Propelling	Feed				Fear	Green	Bitter	Late summer	
Up	Down	Liquid	Static		Protect				Bones			Transition	
Masculine	Feminine								Ears				

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THE OATH

ALGUMAS IDEIAS de HIPOCRATES

- " … a Prática da Arte Médica tem por base três pilares fundamentais: A Doença, o Doente e o Médico. Este último, é como um Servo dessa mesma Arte; o Paciente deve pois cooperar com ele no combate à Doença … " (sic.)

- " … existem alguns Médicos e Filósofos que afirmam que ninguém pode entender o âmago da Ciência Médica sem o prévio conhecimento da verdadeira Natureza Humana. Ninguém que se proponha iniciar-se na Arte de Cuidar e Tratar um Doente deve esquecer-se desse princípio fundamental …" (sic.)

"… o Médico que só sabe Medicina, nem de Medicina sabe … " (sic.) (Abel Salazar)

Amato Lusitano

Centúrias de Curas Medicinais

Volume I

DIALOGO DIALOGO
NEL CVAL... EN EL CVAL SE...

COSMO MEDICI

LIBRI VARY MAIORIS FORMAE

Curationum Medicinalium
AMATI LVSITANI
MEDICI PHYSICI
praestantissimi
CENTVRIAE QVATVOR.

Quibus praemittitur
Commemoratio de introitu medici ad aegrotantem,
De Crisi, & diebus Decretoriis:

INDEX rerum memorabilium copiosissimus.

VENETIIS,
Apud B. Iohanettam Constantinum, sub signo Georgii.
MDLVII

AMATI LVSITANI
Medici ac Philosophi Celeberrimi
IN
DIOSCORIDIS ANAZARBEI
de medica materia libros quinque
ENARRATIONES ERVDITISSIMAE.

Quibus etiam cum Simplicium medicamentorum nomen-
clatura Graecae, Latinae, Italicae, Hispanicae, Germani-
cae, & Gallicae proponuntur: tum errores aliorum
omnium, qui ad hanc usque diem de hac
materia scripserunt, impecabantur.

Cum Rerum ac Verum memorabilium Indice locupletissimo.

Cum priuilegio Illustrissimi Senatus Veneti.
Venetiis excudebat Iordani Zileti. 1557.

תפילת הרופא

CANADA
THE SYSTEM OF MEDICAL TRADE DE MEDICINE
SIR WILLIAM OSLER 1849-1919
6

"One of the first duties of the physician is to educate the masses not to take medicines"

sir william osler

THE PRINCIPLES AND PRACTICE OF MEDICINE

DESIGNED FOR THE USE OF PRACTITIONERS AND STUDENTS OF MEDICINE

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MICHAEL BLISS

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With other Addresses to Medical
Students, Nurses and Prac-
titioners of Medicine

By
Sir WILLIAM OSLER, Bt., M.D., F.R.S.
Late Regius Professor of Medicine, Oxford
Honorary Professor of Medicine, Johns Hopkins University

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ALGUNS PENSAMENTOS do PAI da MEDICINA INTERNA

("William Osler: Na overview of a life", JAMA, 1999, 282, 23, 2252 – 2258; "What is the Oslerian tradition?", Charles Bryan, AIM, 1994, 120, 8, 682 – 687); "Osler lagacy: The centinnial of the Princiles and Pactice of Medicine", Richard Golden, AIM, 1992, 116, 3, 255 – 260; "A legacy of Osler: Teaching clinical ethics at the bedside", Mark Siegler, JAMA, 1978, 239, 10, 951 – 956)

- " … o método natural do verdadeiro ensino médico aos estudantes de medicina, começa com o doente, continua com o doente, e termina com o doente, utilizando os livros e os artigos apenas como meros auxiliares … " (sic.)

- " … estudar o fenómeno das doenças, prescindindo dos ensinamentos dos livros, é como marear sem coordenadas, enquanto que estudar esses mesmos livros sem ouvir os pacientes, é como nem sequer iniciar a viagem … " (sic.)

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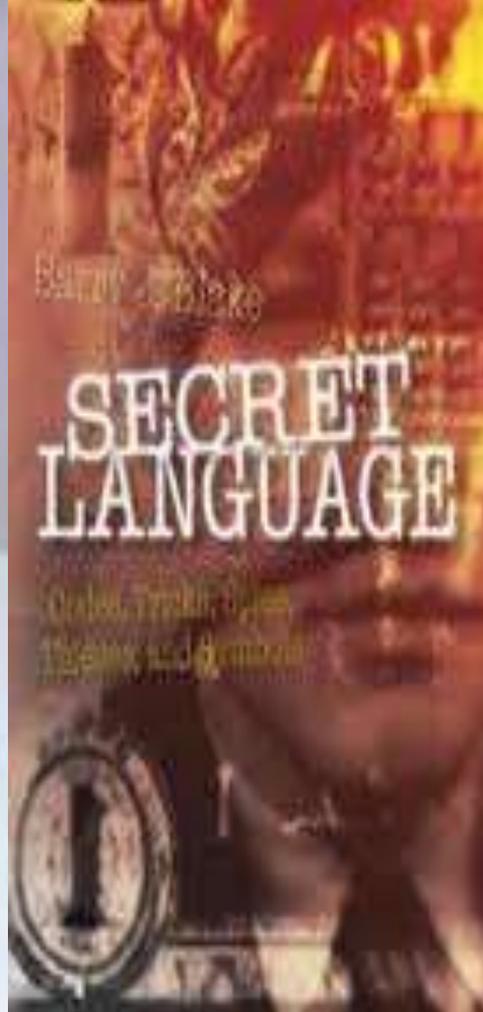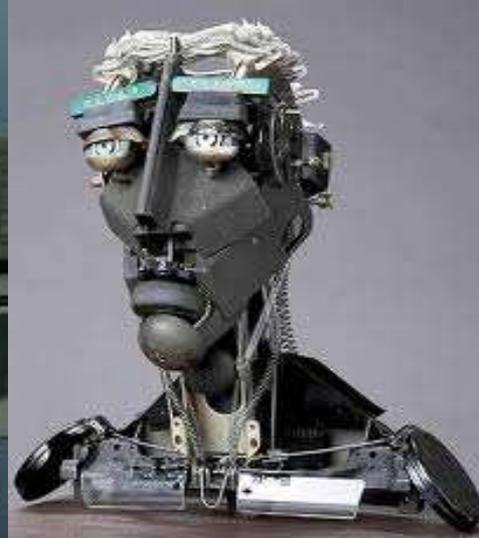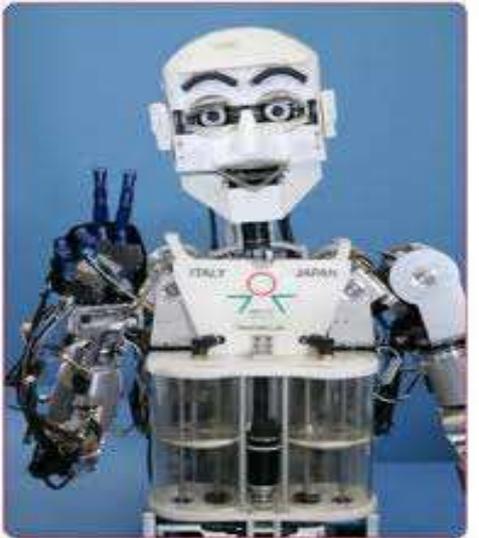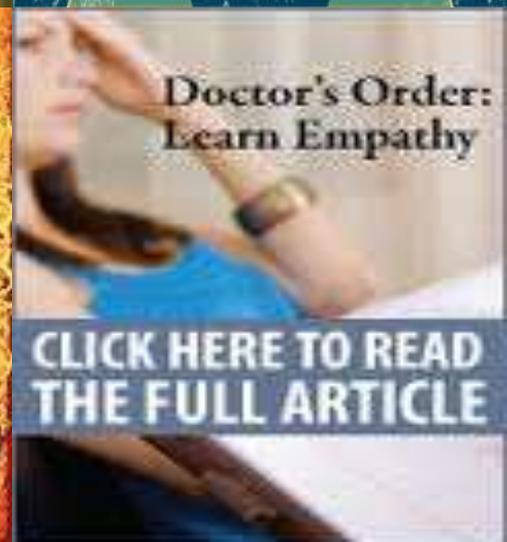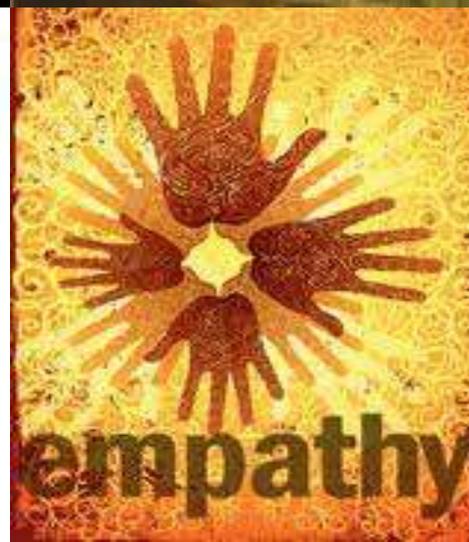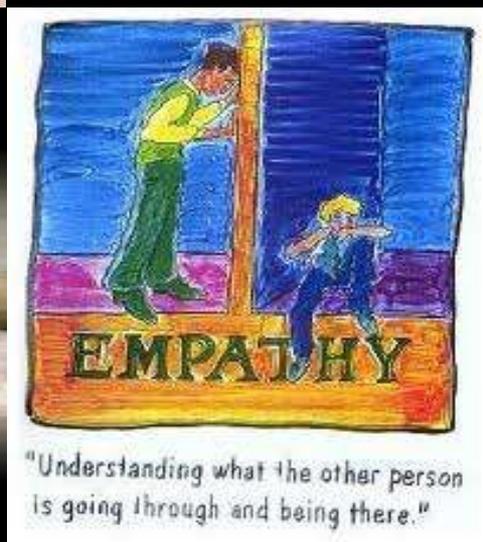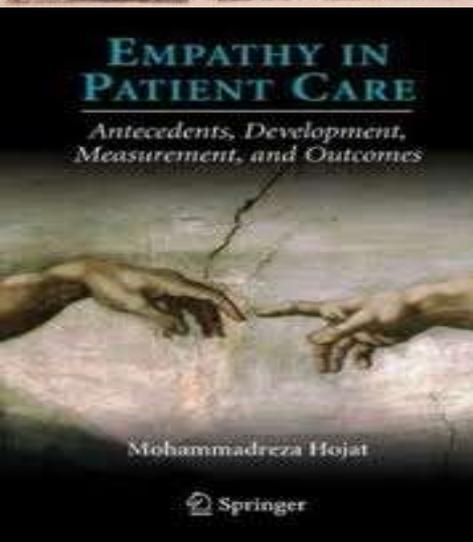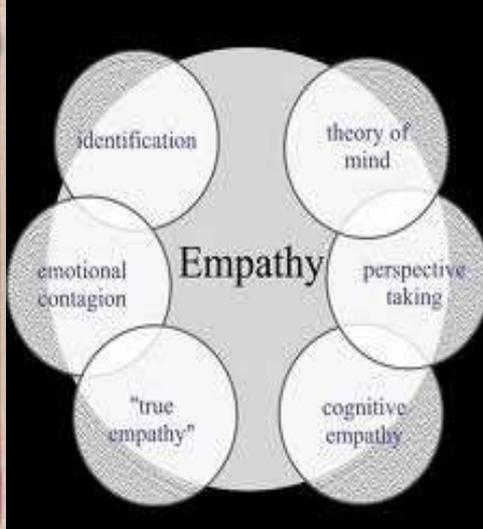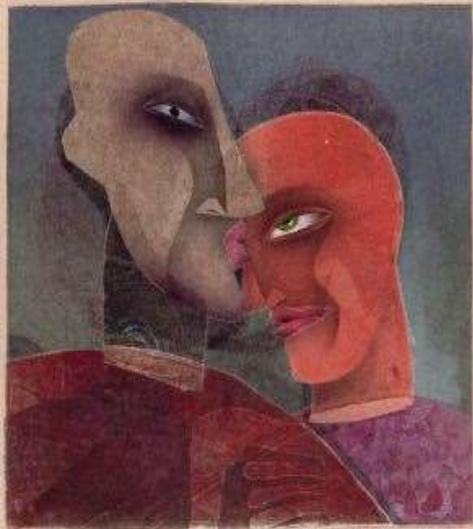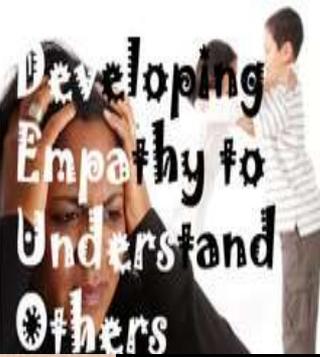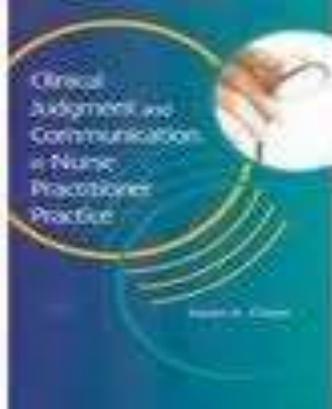
Individual Clinical Expertise

Improved Patient Outcomes

Patient's Values and Expectations

Best Available Clinical Evidence

Rosalinda Alfaro-LeFevre

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A Practical Approach to Outcome-Focused Thinking

Conversations in

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Donna D. Ignatavicius
Bette Case

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Clinical Judgment and the Practice of Medicine

KATHRYN MONTGOMERY

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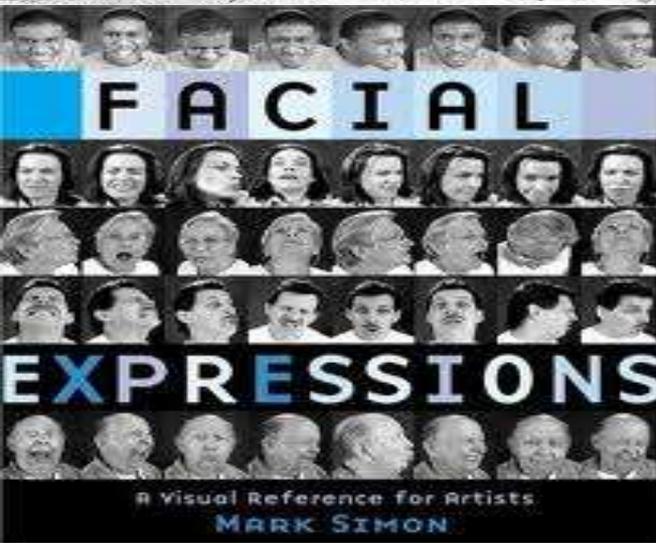
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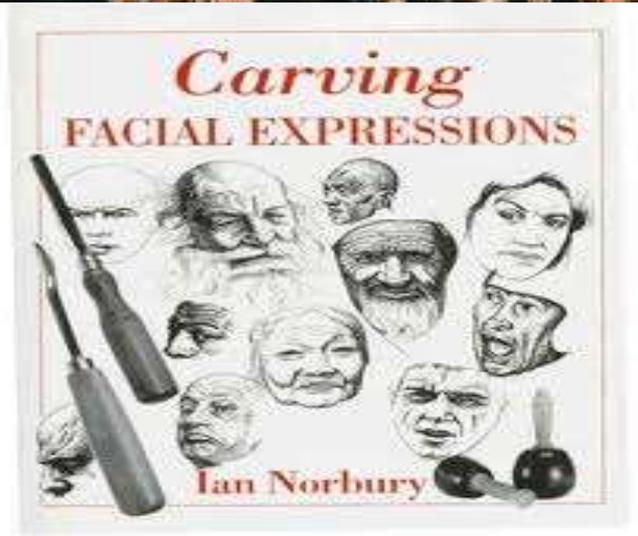
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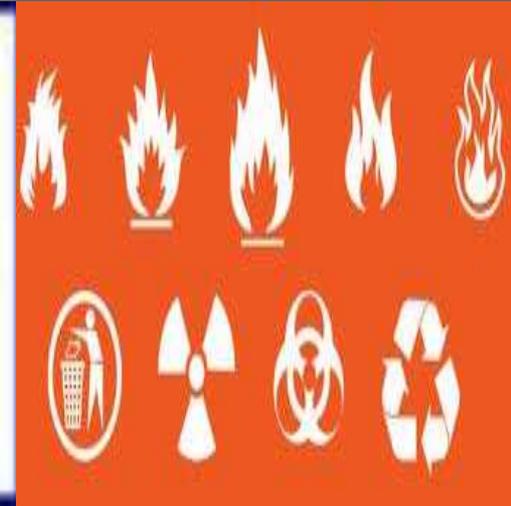
AINDA MAIS DOIS PENSAMENTOS ...

("Medicine as a dependent tradition: Historical and Etical reflections", Richard Vance, Perspective in Biology and Medicine, 1985, 28, 2, 282 – 302; "Is simulation based medicine training the future of clinical medicine?", J. Murphy, et al, European Review for Medical and Pharmacological Sciences, 2007, 11, 1 – 8)

- " ... a prática médica está presentemente numa posição algo vulnerável e indefinida, dado que é concebível que a própria Medicina se esteja a transformar em algo substancialmente diferente daquilo por que sempre clamou dever ser a sua verdadeira índole, ou seja, a missão de se dedicar essencialmente ao tratamento do Ser Humano enquanto Doente ... sendo por isso que acredito profundamente que o nosso maior desafio de natureza ética na situação presente é o do retorno a essa veneranda tradição ... " (sic.)

- " ... o desenvolvimento das qualidades humanísticas e a compaixão pelos doentes por parte dos estudantes e dos jovens internos, constituem pilares básicos fundamentais para o exercício do nosso mister, o qual irá sempre requerer um acompanhamento permanente baseado no exemplo por parte dos respectivos professores e tutores, para os quais nunca será suficientemente apenas disponibilizarem meios tecnológicos, por mais sofisticados que possam ser ... " (sic.)

GLOBALIZATION
AND HEALTH

DEBATE　　　　　　　　　　　　　　　　　　　　**Open Access**

Rethinking the 'global' in global health: a dialectic approach

Kayvan Bozorgmehr

Abstract

Background: Current definitions of 'global health' lack specificity about the term 'global'. This debate presents and discusses existing definitions of 'global health' and a common problem inherent therein. It aims to provide a way forward towards an understanding of 'global health' while avoiding redundancy. The attention is concentrated on the dialectics of different concepts of 'global' in their application to malnutrition; HIV, tuberculosis & malaria; and maternal mortality. Further attention is payed to normative objectives attached to 'global health' definitions and to paradoxes involved in attempts to define the field.

Discussion: The manuscript identifies denotations of 'global' as 'worldwide', as 'transcending national boundaries' and as 'holistic'. A fourth concept of 'global' as 'supraterritorial' is presented and defined as 'links between the social determinants of health anywhere in the world'. The rhetorical power of the denotations impacts considerably on the object of 'global health', exemplified in the context of malnutrition; HIV, tuberculosis & malaria; and maternal mortality. The 'global' as 'worldwide', as 'transcending national boundaries' and as 'holistic' house contradictions which can be overcome by the fourth concept of 'global' as 'supraterritorial'. The 'global-local-relationship' inherent in the proposed concept coheres with influential anthropological and sociological views despite the use of different terminology. At the same time, it may be assembled with other views on 'global' or amend apparently conflicting ones. The author argues for detaching normative objectives from 'global health' definitions to avoid so called 'entanglement-problems'. Instead, it is argued that the proposed concept constitutes an un-euphemistical approach to describe the inherently politicised field of 'global health'.

Summary: While global-as-worldwide and global-as-transcending-national-boundaries are misleading and produce redundancy with public and international health, global-as-supraterritorial provides 'new' objects for research, education and practice while avoiding redundancy. Linked with 'health' as a human right, this concept preserves the rhetorical power of the term 'global health' for more innovative forms of study, research and practice. The dialectic approach reveals that the contradictions involved in the different notions of the term 'global' are only of apparent nature and not exclusive, but have to be seen as complementary to each other if expected to be useful in the final step.
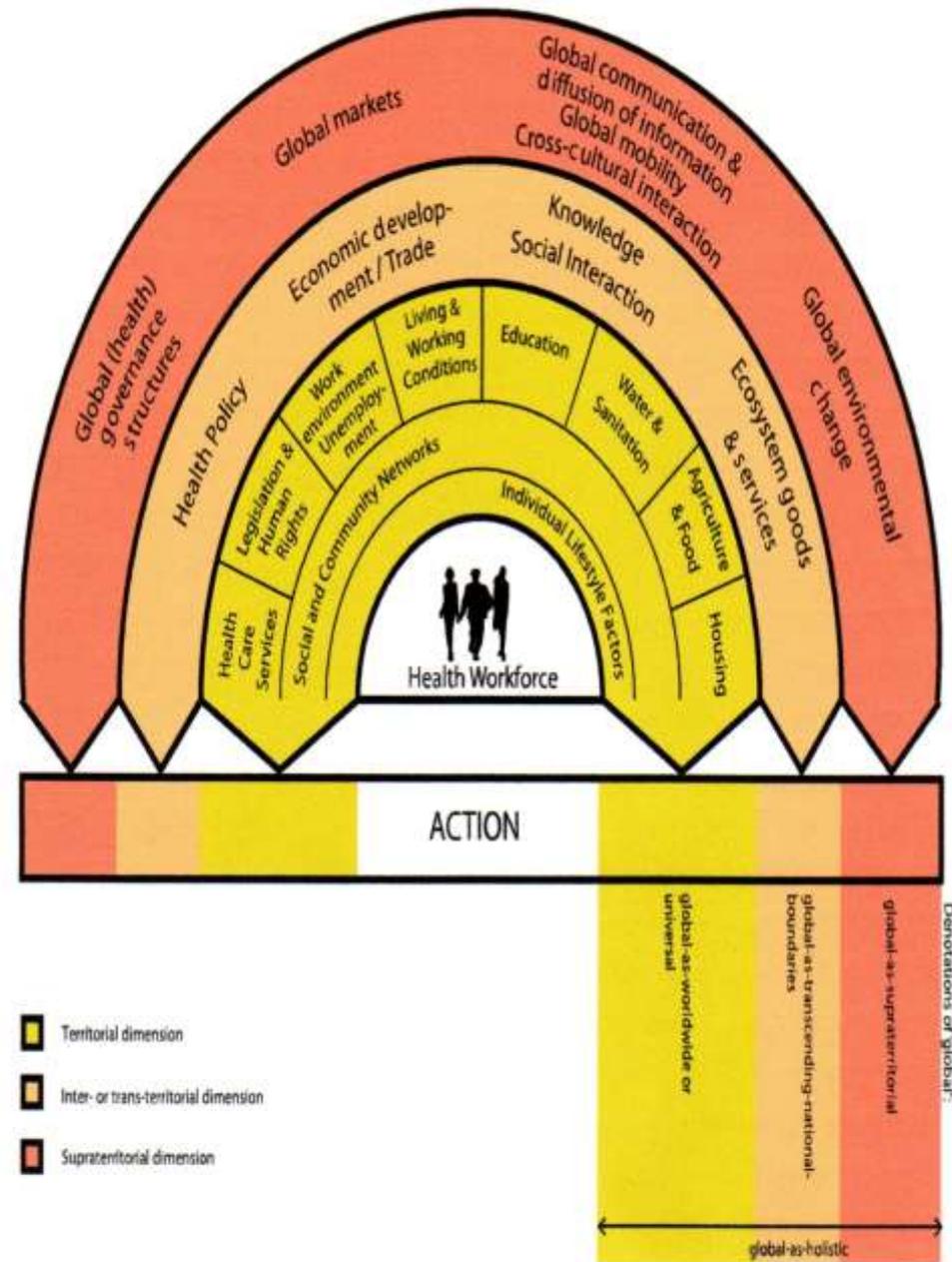
Figure 1 Concept of global health. Territorial dimension: includes for example determinants on territorial units such as community upto state or national units; Inter- or trans-territorial dimension: includes for example determinants which link and/or transcend territorial units, e.g. national borders; Supraterritorial dimension: includes social, political, economic and cultural links between determinants of health anywhere in the world regardless of territory in terms of geography.

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SUSTAINABILITY
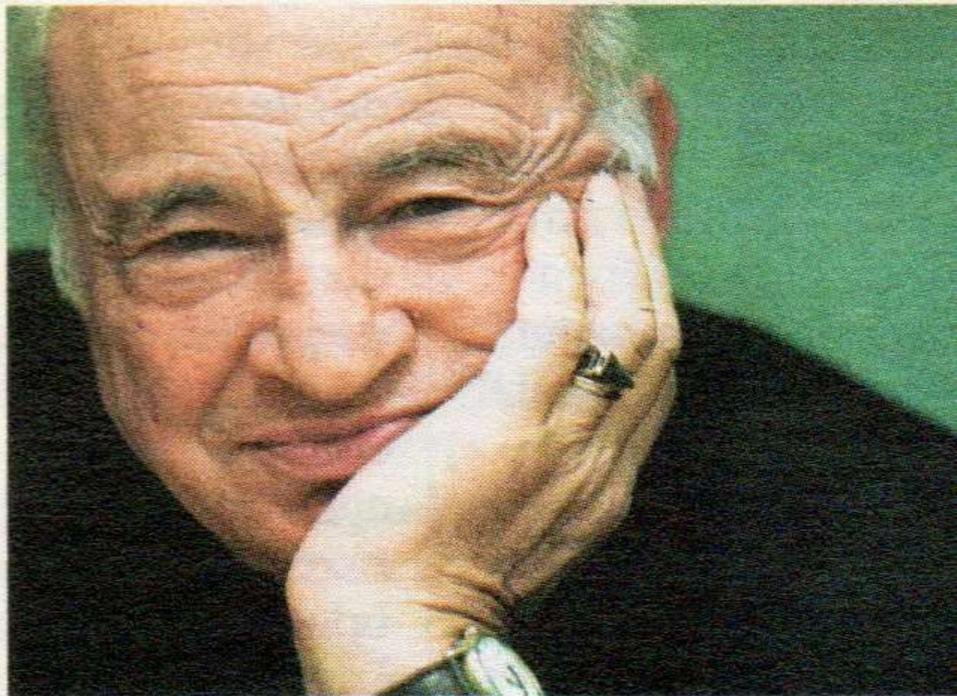
Edgar Morin

**A ideia fixa do
crescimento contínuo
e interminável não pode
continuar**

Response to population pressures
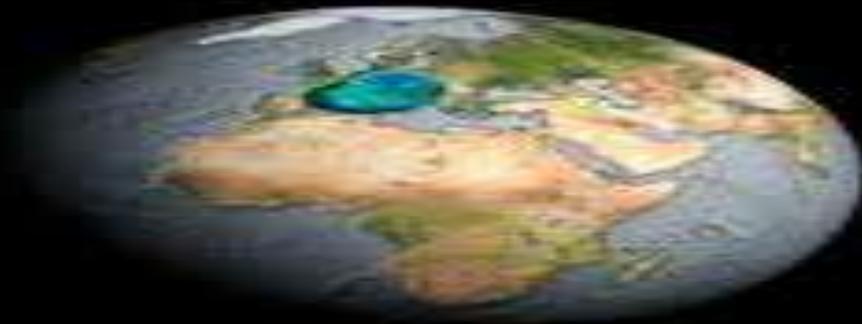
" … parece que nos habituamos a pensar nos problemas de saúde tal como o fazemos relativamente a todos os outros de natureza diversa, em que a ciência e a tecnologia nos há-de salvar, embora na realidade a resposta deva antes ser encontrada sobretudo no ser humano e na sua capacidade relacional …" (sic.) *(James Curran)*
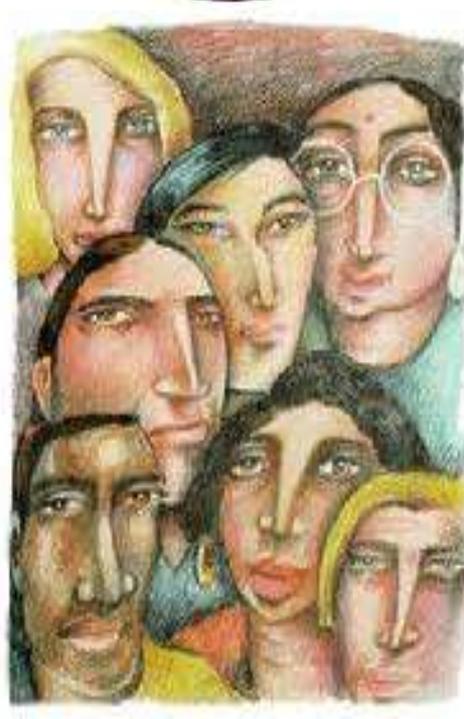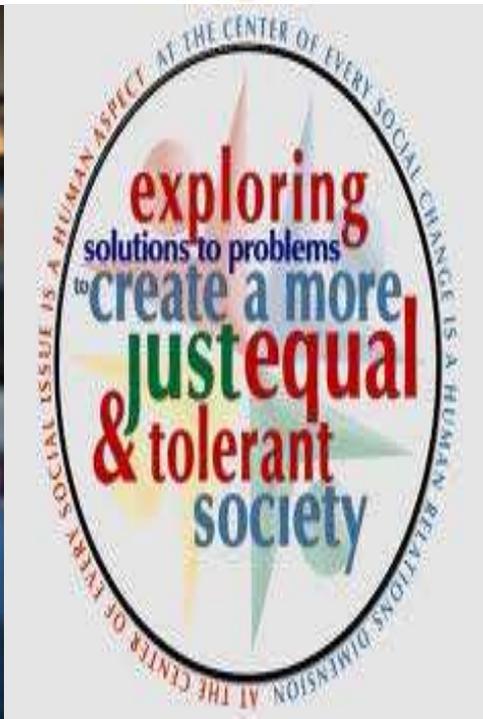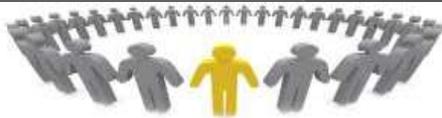
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exploring
solutions to problems
to create a more
just equal
& tolerant
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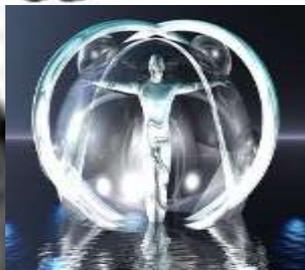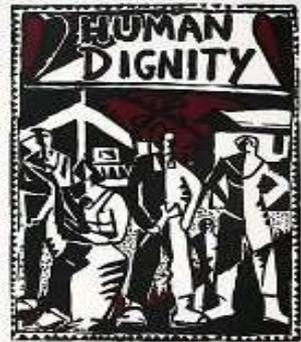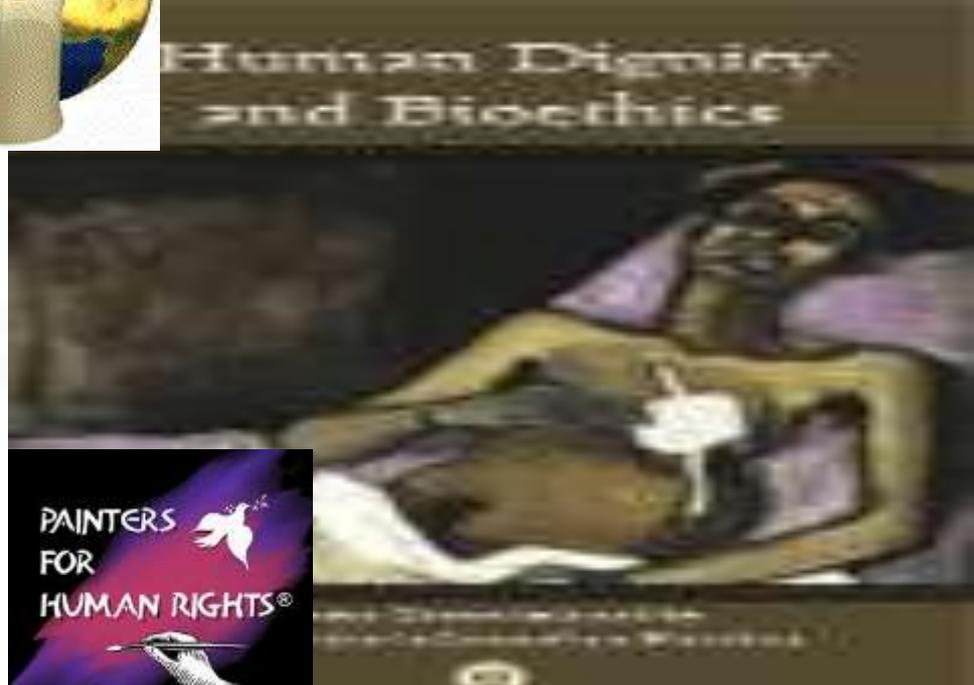
Escrito na pedra

O que mais desespera, não é o impossível. Mas o possível não alcançado.

Robert Mallet (1915-2002), escritor francês

HOPE

"LEARN FROM YESTERDAY, LIVE FOR TODAY, HOPE FOR TOMORROW."
ALBERT EINSTEIN

HOPE

YOUR EXISTENCE GIVES ME HOPE

In America No One Should Go Without HEALTH CARE

HOPE

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