

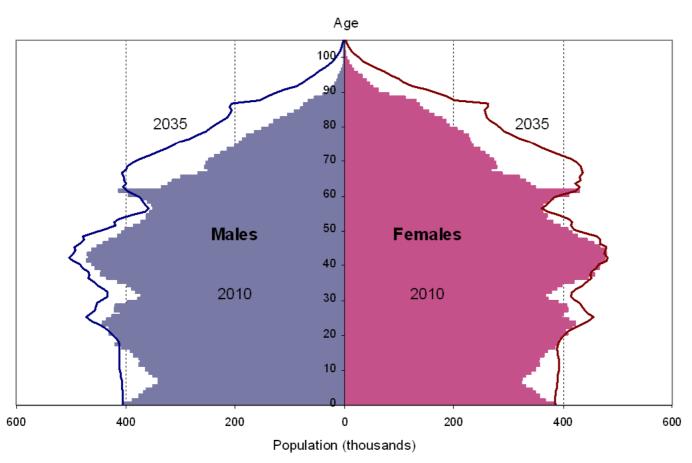
making financial sense of the future

Momentum conference 2011: for Actuaries of Today and Tomorrow Daniel Ryan

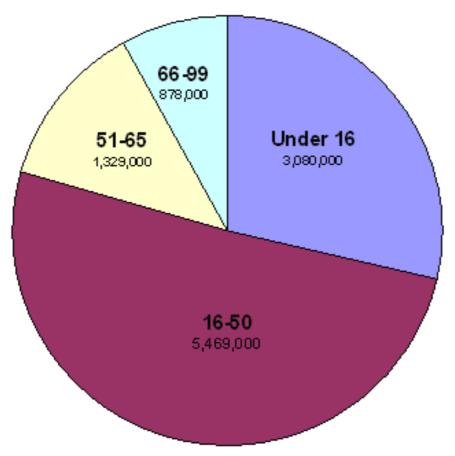


### The future shape of our population

Estimated and projected age structure of the United Kingdom population, mid-2010 and mid-2035



#### Who are our future centenarians?



Source: Office for National Statistics, 2008-based Population Projections (UK)

#### Why you will live to 100 (or not)

- Genes
- Behaviours
- Environment
- Societal pressures
- Accidents
- Medical interventions
- Stochastic variation
- Historical trends
  - New England Centenarian study suggested dominant impact of genes in extreme longevity

# Tips for healthy ageing National Institute of Ageing

- Eat a balanced diet.
- Exercise regularly.
- Get check-ups on a regular basis.
- Do not smoke.
- Practice safety habits at home to prevent falls and bone fractures.
   Wear a seatbelt in the car.
- Stay in contact with family and friends.
- Avoid too much exposure to the sun or cold weather.
- Drink alcohol only in moderation.
- Keep personal and financial records in good order.
- Keep a positive attitude toward life. Do the things that bring happiness.

#### Relative importance of genes and environment

Genome Environment

Genotype Non - Genetic

Phenotypes

Genotype Thrombosis Non - Genetic

Schizophrenia Alzheimer

Fam. breast cancer Diabetes

Colon cancer syndrome Asthma

Cystic Fibrosis Lung cancer

Huntington's Spectrum of Disease Car accident

#### Genes associated with increased risk of disease

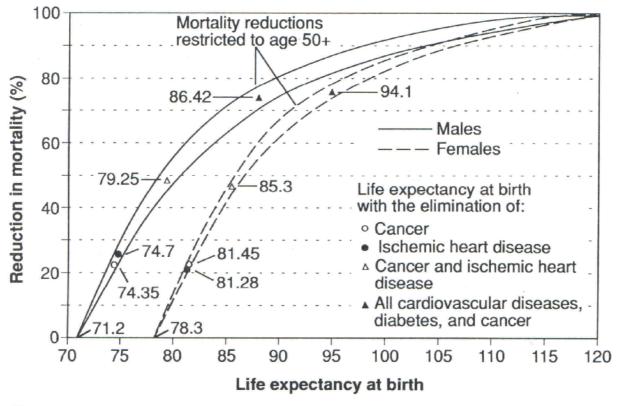
Cancer site	Relative Risk ≥5.0 Family studies	Relative Risk ≥1.5 and >5.0 Resequencing	Relative Risk ≥1.01 and >1.5 Genome-wide association studies
Lung	RB1, TP53		rs1051730, rs8034191 (CHRNA3, CHRNB4, CHRNA5)
Breast	BRCA1, BRCA2, TP53, PTEN, SK11, CDH1	CHEK2, ATM, PALB2, BRIP1	CASP8, FGFR2, MAP3K1, 8q24, 5p, TOX3, 2q, 6q22, LSP1
Colon and rectum	APC, MLH1, MSH2, MSH6, PMS2	APC (I1307K), BLM	MUTYH, CASP8, 8q24, 8q23 (EIF3H), 10p14, 11q23, CRAC1, SMAD7
Prostate	BRCA2	8q24	rs6501455, rs721048, NBS1, EHBP1, TCF2, CTBP2, JAZF1, MSMB, LMTK2, KLK3, SLC22A3
Pancreas	BRCA2, CDKN2A, STK11, TP53, PRSS1, SPINK1	BRCA1, MSH2, MLH1	

Source: Foulkes W; N Engl J Med; 2008;359:2143-2153

## Different approaches to considering the future Converging or Diverging?

- "Projectionists" e.g. Vaupel no current evidence of restrictions to improvements in life expectancy leading to expectations that medical advances will deliver – up to 0.25 years per calendar year
- "Realists" e.g. Olshansky treatment of disease without affecting ageing process has limited potential to expand life expectancy, and not clear how "less healthy" cohorts will develop
- No current treatments affect ageing process and no biomarkers to determine effectiveness of treatments
- Acceptance of the possibility of future treatments that could slow ageing process

# Understanding the potential for further improvements from disease-elimination models



**Fig. 2.** Percentage of reduction in the conditional probability of death for the United States (from 1985 levels) required to produce a life expectancy at birth from 80 to 120 years.

#### The impact of the ageing process

- Hand grip strength reduces by 45% by age 75
- Blood flow to brain reduces by 15-20% by age 70
- Sense of smell reduces to 50% of peak by age 80
- Maximum heart rate reduces by 15-20% by age 70
- Blood pressure of 50% population at age 65 is mild or worse hypertension
- Maximum breath capacity reduces by 40% by age 80
- Dementia affects 10% of those over age 65; 20% of those over age 85

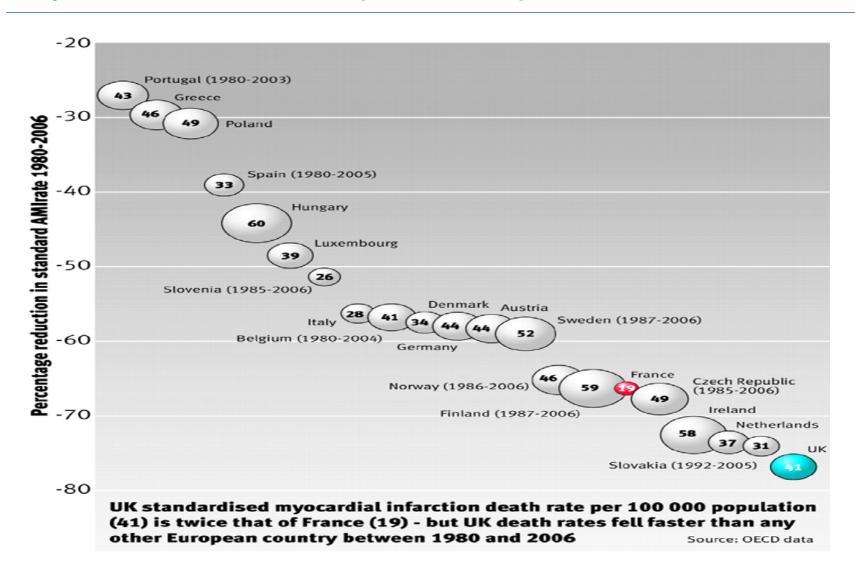
# Potential benefits from disease elimination and slowing of ageing process

TABLE 2

U.S. Social Security Administration, U.S. Census Bureau, and the Network Observed (2000) and Projected (2030, 2050) Life
Expectancy at Birth and at Ages 65 and 85 for U.S. Males and Females

	2000	2030				2050			
		SSA	СВ	Network A (lower disease mortality)	Network B (slow aging)	SSA	СВ	Network A (lower disease mortality)	Network B (slow aging)
					Life Expectancy a	t Birth			
Males	74.0	78.0	78.4	79.4	80.4	80.0	80.9	83.2	85.9
Females	79.4	81.8	83.1	85.1	86.3	83.4	85.3	89.2	93.3
Total	76.7	79.9	80.8	82.3	83.4	81.7	83.1	86.2	89.6
				I	Life Expectancy at	Age 65			
Males	15.9	18.2	19.0	20.2	21.8	19.3	20.6	23.4	27.1
Females	19.0	20.3	21.7	23.7	25.5	21.4	23.2	27.4	32.4
Total	17.4	19.2	20.4	22.0	23.7	20.4	21.9	25.4	29.8
				I	Life Expectancy at	Age 85			
Males	5.2	6.0	6.8	7.6	9.2	6.5	7.6	9.7	13.6
Females	6.4	7.1	8.0	9.5	11.5	7.6	8.9	12.3	17.8
Total	5.8	6.6	7.4	8.6	10.4	7.1	8.3	11.0	15.7

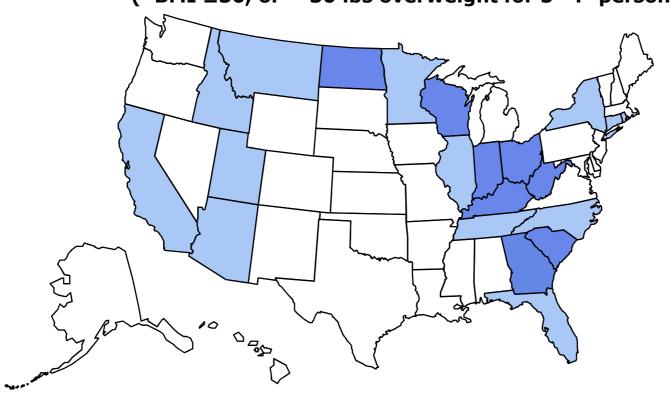
# International trends in cause-specific mortality Myocardial infarction (1980-2006)



### Relative importance of risk factors and treatment

Risk Factor	Effect	Treatments	Effect
Obesity	+3%	AMI	-8%
Diabetes	+5%	Secondary prevention	-11%
Blood Pressure	-10%	Heart failure	-13%
Smoking	-48%	Angina: CABG/PTCA	-7%
Cholesterol	-9%	Hypertension therapy	-3%
Physical activity	+4%		
Deprivation	-3%		
Total	-58%		-42%

(\*BMI ≥30, or ~ 30 lbs overweight for 5' 4" person)



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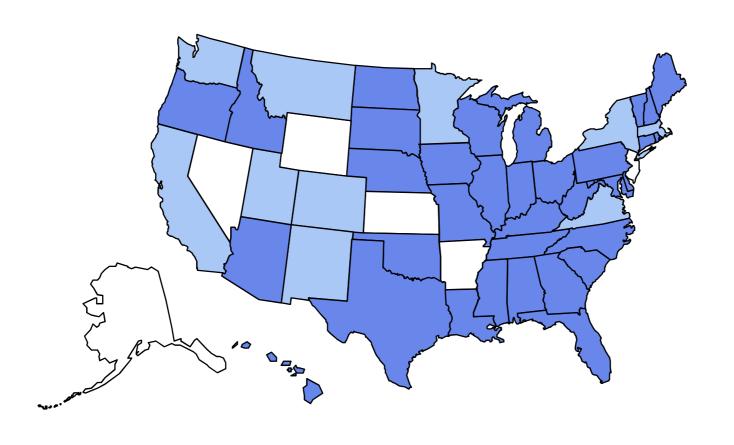
<10%

10%-14%

15%-19%

20%-24%

≥25%



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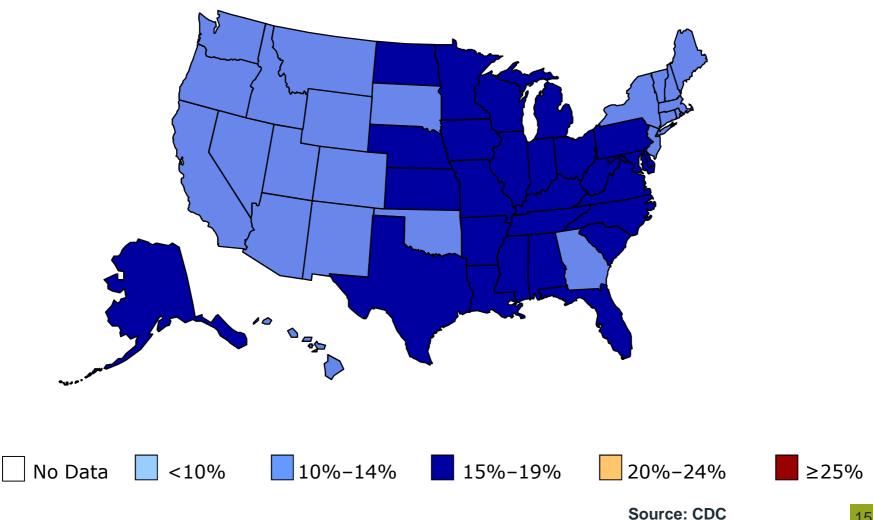
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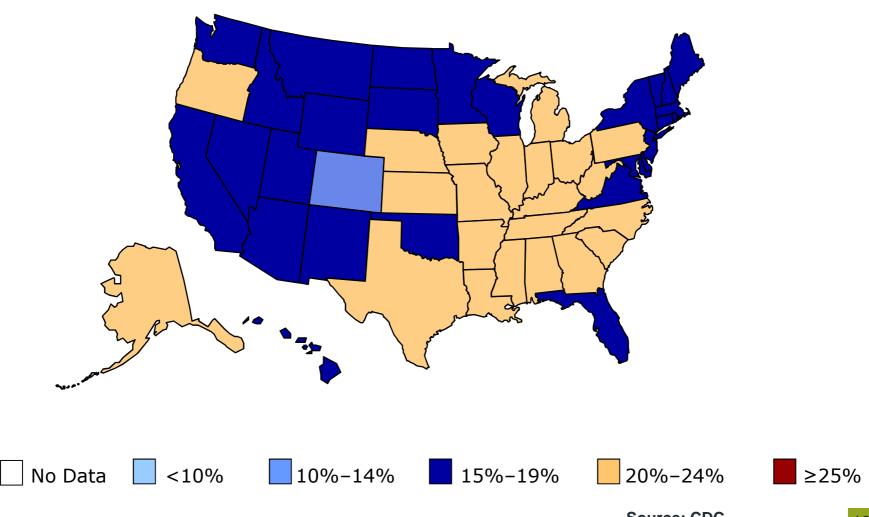
10%-14% 15%-19%

20%-24%

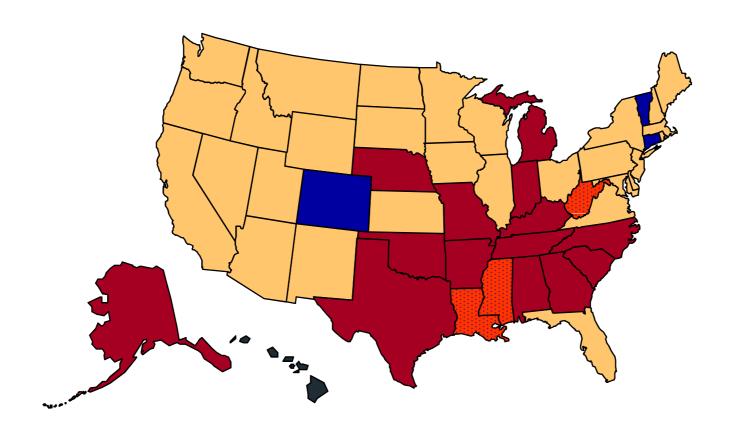
≥25%

Source: CDC





Source: CDC



No Data

<10%

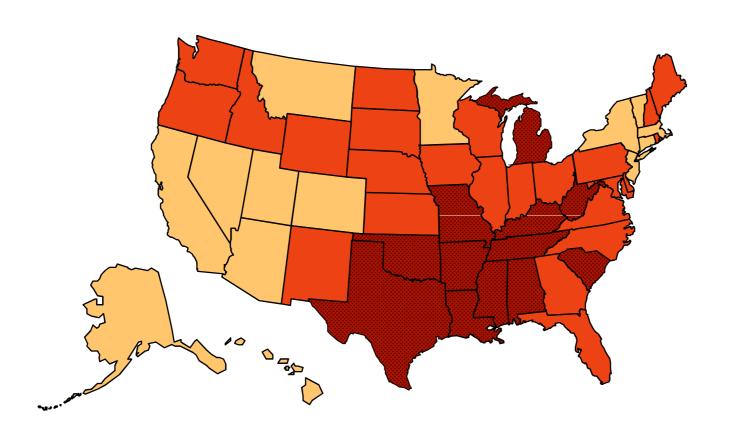
10%-14%

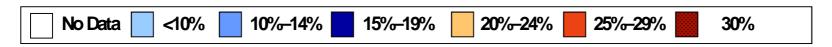
15%-19%

20%-24%

25%-29%

30+%





Source: CDC



Germany: The Melander family of Bargteheide Food expenditure for one week: 375.39 Euros or \$500.07



United States: The Revis family of North Carolina Food expenditure for one week \$341.98



Italy: The Manzo family of Sicily Food expenditure for one week: 214.36 Euros or \$260.11



Egypt: The Ahmed family of Cairo Food expenditure for one week: 387.85 Egyptian Pounds or \$68.53

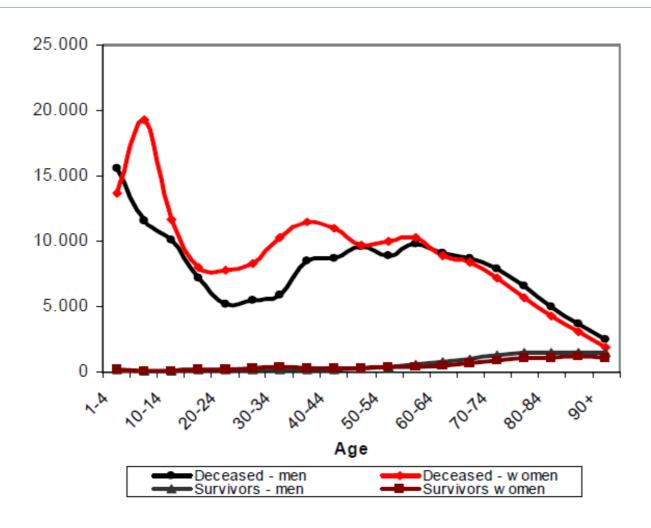


Bhutan: The Namgay family of Shingkhey Village Food expenditure for one week: 224.93 ngultrum or \$5.03

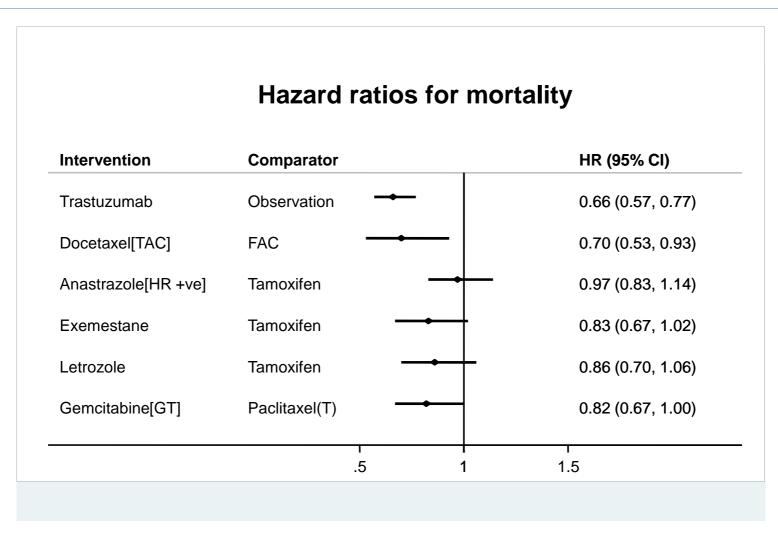
#### What do we want from our healthcare systems

- Fast access to reliable health advice
- Effective treatment delivered by trusted professionals
- Involvement in decisions and respect for preferences
- Clear, comprehensible information and support for self-care
- Attention to physical and environmental needs
- Emotional support, empathy and respect
- Involvement of, and support for, family and carers
- Continuity of care and smooth transitions

## Healthcare expenditure at end of life Survivors and deceased in regional study in Italy

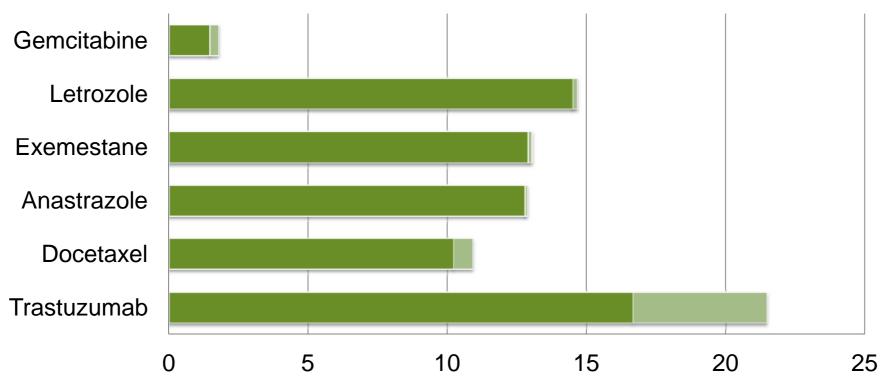


## **Comparisons of different treatments Breast cancer**



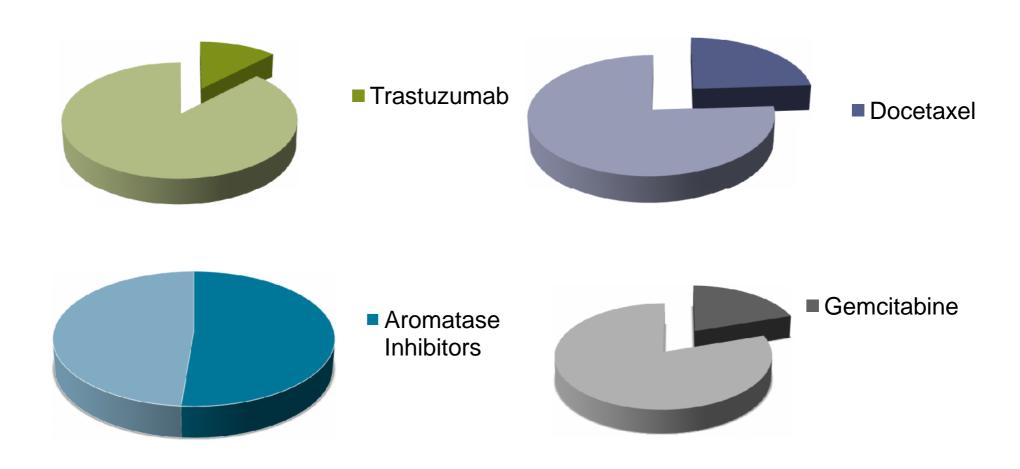
## Comparisons of different treatments Breast cancer

## Modelled life expectancy with and without interventions

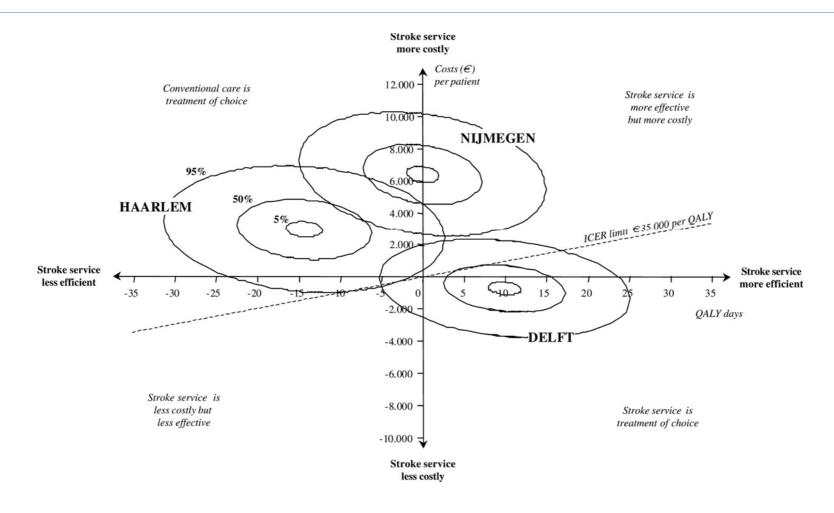


■ Mean survival without intervention
■ Additional survival with intervention

## **Guidance over use of different treatments** % of breast cancers



## **Evaluating cost-effectiveness of treatments Co-ordinated stroke care in different locations**



## Clinical trials – scope and timeframe

Time	Preclinical testing 3 - 5 years	Phase I Trials 1 year	Phase II Trials 2 years	Phase III Trials 3 years	Filing/ approval 2 years	Phase IV open
Study subject	Laboratory and animal studies	20 - 80 healthy volunteers	100 - 300 patient volunteers	1'000 – 3'000 patient volunteers	-	open, according to indication
Study aim	Assess safety & biol. activity	Determine safety & max. dose	Evaluate effective dose, side effects	Verify efficacy, monitor long term	Review process	Post marketing safety monitoring

Source: CDC

## Clinical trials – approval rates by therapeutic class

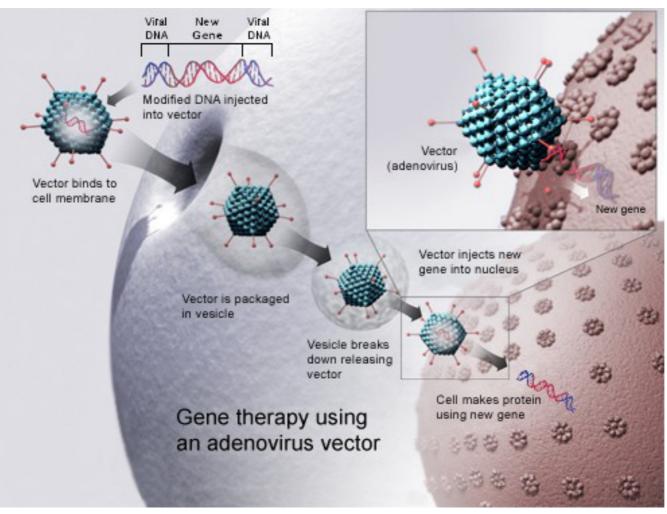
Table 3 Phase transition and clinical approval probabilities by therapeutic class for self-originated compounds first tested in humans from 1993 to 2004

Therapeutic class	Phase I–II (%)	Phase II–III (%)	Phase III–RR (%)	RR-approval (%)	Clinical approval success rate (%)
Antineoplastic/immunologic	71.8	49.0	55.3	100	19.4
Cardiovascular	62.9	32.4	64.3	66.7	8.7
CNS	59.6	33.0	46.4	90.0	8.2
GI/metabolism	67.5	34.9	50.0	80.0	9.4
Musculoskeletal	72.4	35.2	80.0	100	20.4
Respiratory	72.5	20.0	85.7	80.0	9.9
Systemic anti-infective	58.2	52.2	78.6	100	23.9
Miscellaneous	62.8	48.7	69.8	91.3	19.5

Through June 2009.

CNS, central nervous system; GI, gastrointestinal; RR, regulatory review.

# Regenerative medicine: gene therapy

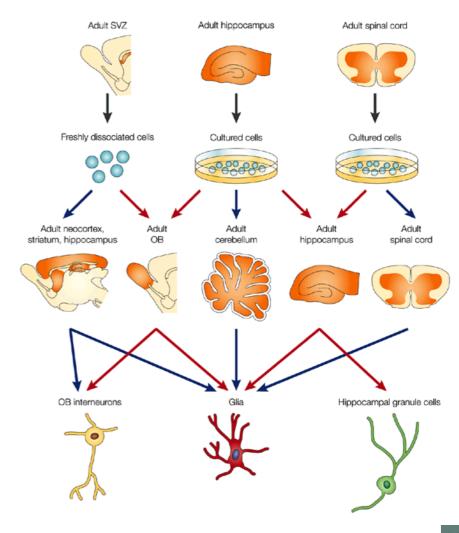


# Regenerative medicine: cell therapy

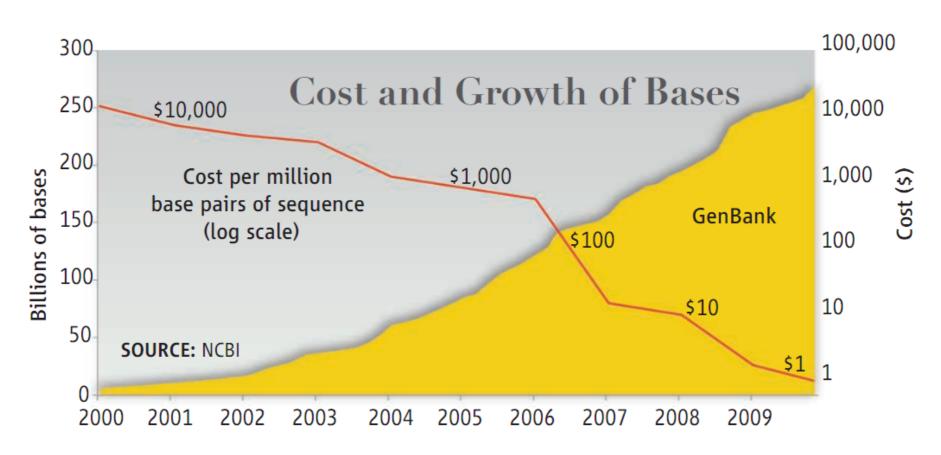




Nature Reviews Neuroscience 2002



# **Genetic information:**Revolutionary developments from DNA sequencing

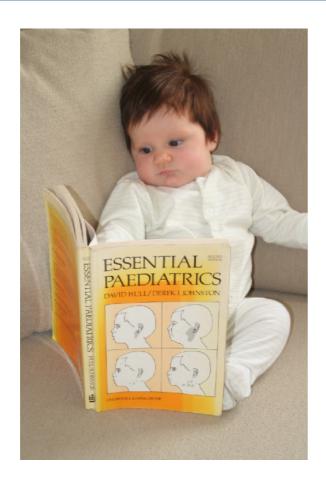


Source: E. Pennisi, Science 2011, 331, 666-8

## Combining different elements in a forwardlooking approach to assessing future mortality

Individual risk factors Healthcare funding Patient interaction Research & development General drivers to Age, gender, diet, smoking - Public vs private funding Health awareness Public vs commercial smoking considerations: Disease-based patient Trust and confidence in sponsors diagnosis and Taxes and restrictions Regulators' attitude to advocacy groups' advice given survival Current treatments influence Use of clinical guidelines developments Allocation of resources • Disease-focused approach (buproprion) to improve quality of care Future treatments (vaccines) towards cure vs vs global impact of ageing prevention Circulatory Respiratory Stroke, angina, Chronic obstructive Multiple heart attack pulmonary disease diseases Disease types and Healthy Death disease progression Cancer Neurological Lung, colorectal Dementia, prostate breast Alzheimer's. Current approaches **Risk factors** Parkinson's Targeting DCIS Family history Surgery with node follow-up Obesity Adjuvant radiotherapy • Having children later in life Herceptin, Tamoxifen **Breast cancer**  Not breast feeding Clinical trials pipeline **Early detection**  Phase II (230 trials\*) Digital mammography Factors involved in Medical innovations • Phase III (56) • MRI for high-risk Growth factor inhibition eg pertuzumab (limits cancer) assessing specific · Gail algorithm (own factors) Future of personalised growth) Klaus algorithm (family history) example disease medicine (eg tumour profiling)

# **Education of future medical professionals Implications of an ageing society**

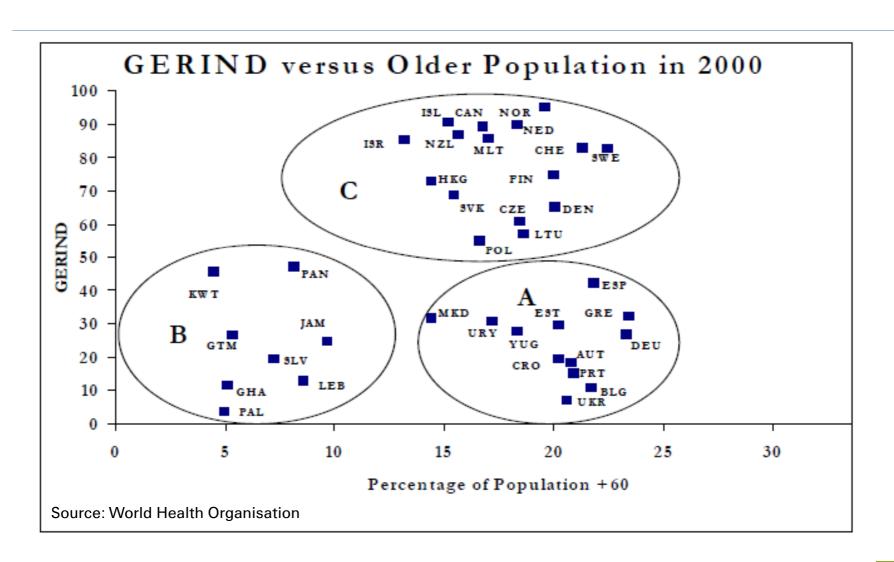




# Teaching Geriatric in Medical Education study (TeGeMe)

- Collaborative study of WHO and International Federation of Medical Students' Associations
- WHO intends healthy/active ageing and promotion of long term health to form education of all future young doctors
- Promotion of life course in graduate training and later
- 41% of medical school curricula refer explicitly to geriatrics
- GERIND index calculated by medical school and averaged across country –separation of geriatrics teaching and quality of ageing science being taught
- Central hypothesis is that countries with higher percentage of older persons are more likely to have separate high-quality teaching on geriatric medicine – not always true

#### TeGeMe – GERIND index vs. age of population



#### A new relationship between doctor & patient

- Classic asymmetrical relationship based on knowledge
- Medical and surgical specialisation driven or required because of technical information & procedures
- Doctors facing information overload
- Transforming effects of internet as clinical guidance becomes more comprehensive
- Two key roles for doctors
  - Patient advocate facilitate patient-based healthcare and act as guide to new technological breakthroughs
  - Scientist/technician maintain pace of development

A new functional divide across the profession

#### Some thoughts for holders of longevity risk

- Continuing differences in schools of thought over future longevity
- Conflicting forces between risk behaviours and treatment
- Impending revolution in genetic information
- Increasing demands from regulators for justification
- No market as yet in longevity risk
- Holders of longevity risk have several options
  - Transfer risk
  - Invest in further research and understanding
  - Wait and see

