# SOME SUGGESTIONS FOR ALLOCATION OF WORLD NEUROSURGERY ASSETS (SUPPLY, COST AND CURE)

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### I. Introduction

- As the world grows closer together and as scientific and medical advances become more evident, major question arises as for the best use for world healthcare resources.
- The practicalities of geopolitical arrangement make easy reallocation difficult and indeed rarely considered.
- We endeavor to explore the subject



### I.a. Outline of the presentation

- INTRODUCTION
- THE VIEW FROM MT. OLYMPUS AT PRESENT
  - World population, income and healthcare dynamics
  - Breakdown of health expenditures in the world
  - Neurosurgery expenditures
- THE VIEW FROM MT. OLYMPUS RESOURCE REALLOCATION (COMPARATIVE ANALYSIS)
  - Glioblastomas
  - Ruptured intracranial aneurysms
- WHY NOT WORLDWIDE ACCESS TO STANDARD OF CARE/ CLINICAL GUIDELINES/ TREATMENT OF GLIOBLASTOMAS AND RUPTURED ANEURYSMS?
- THE VIEW FROM MT. OLYMPUS HOW CAN WORLDWIDE HEALTHCARE RESOURCE DYSFUNCTION BE IMPROVED?
  - Altruistic coefficient graph
  - Collaborative arrangement between the developed and developing world
  - The developed world's efforts
  - What can developing countries do?
  - A third solution

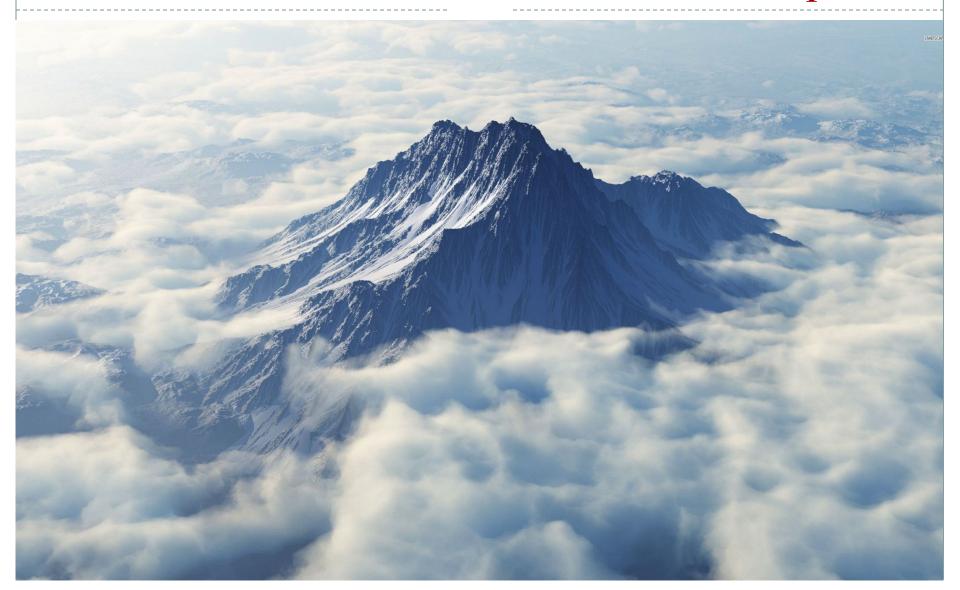
### I.b. Limitations of presented data

- Fluctuations in currency rates
- Insufficiency or inaccuracy of the available international data
- Errors in population numbers
- Errors in international statistical data
- Methodological difficulties and data summation



### II. THE VIEW FROM MT. OLYMPUS

- at present



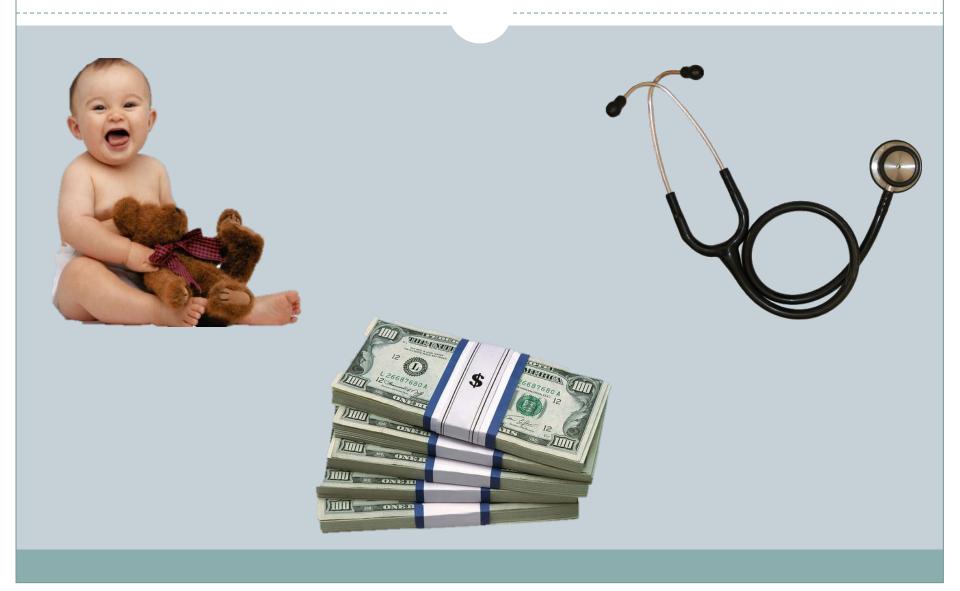


Total Value - \$256 trillion (Credit Suisse 2016)

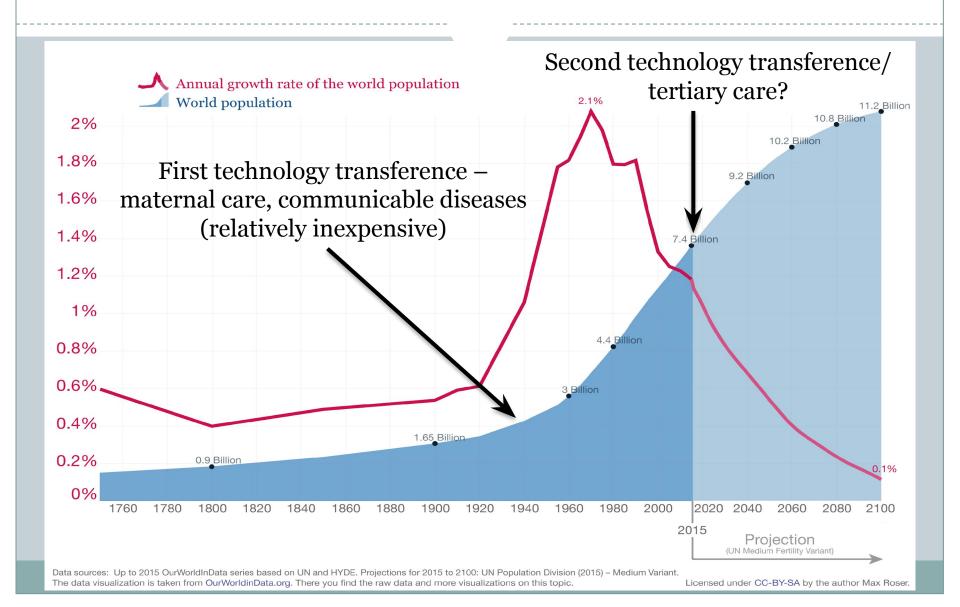
GWP – \$77.9 trillion (CIA World Factbook 2014)

Health expenditure – \$9.5 trillion (World Bank/WHO 2015)

### II. World population, income and healthcare dynamics

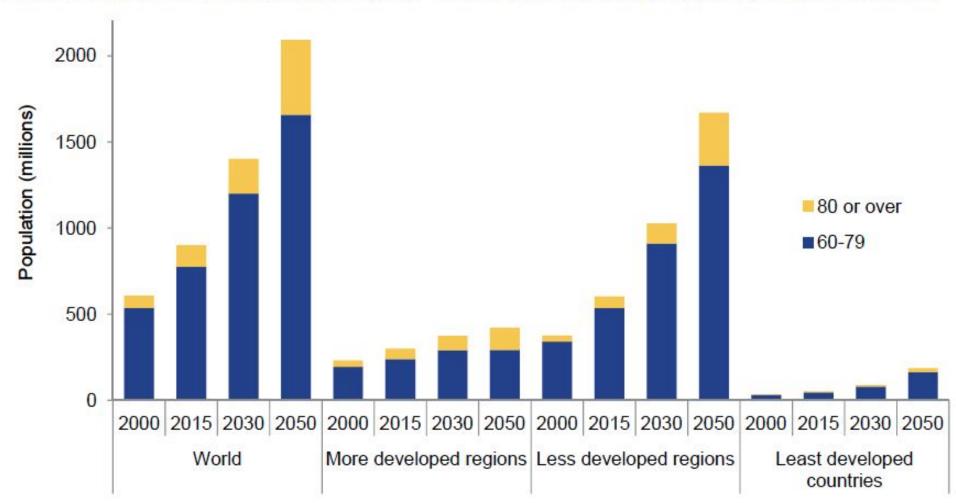


### i. World population growth and growth rate

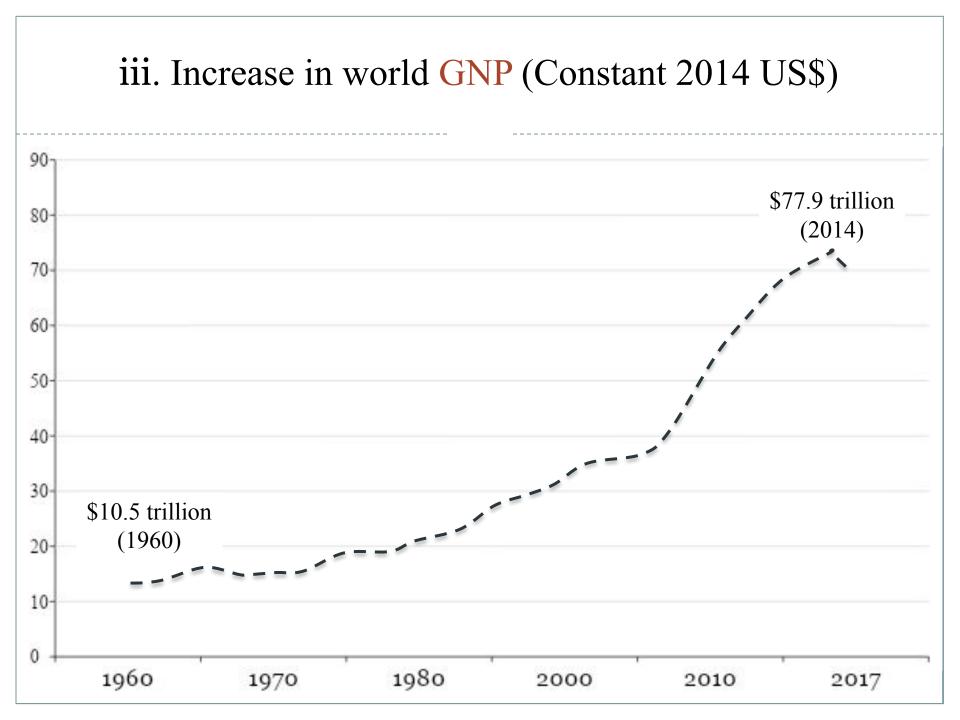


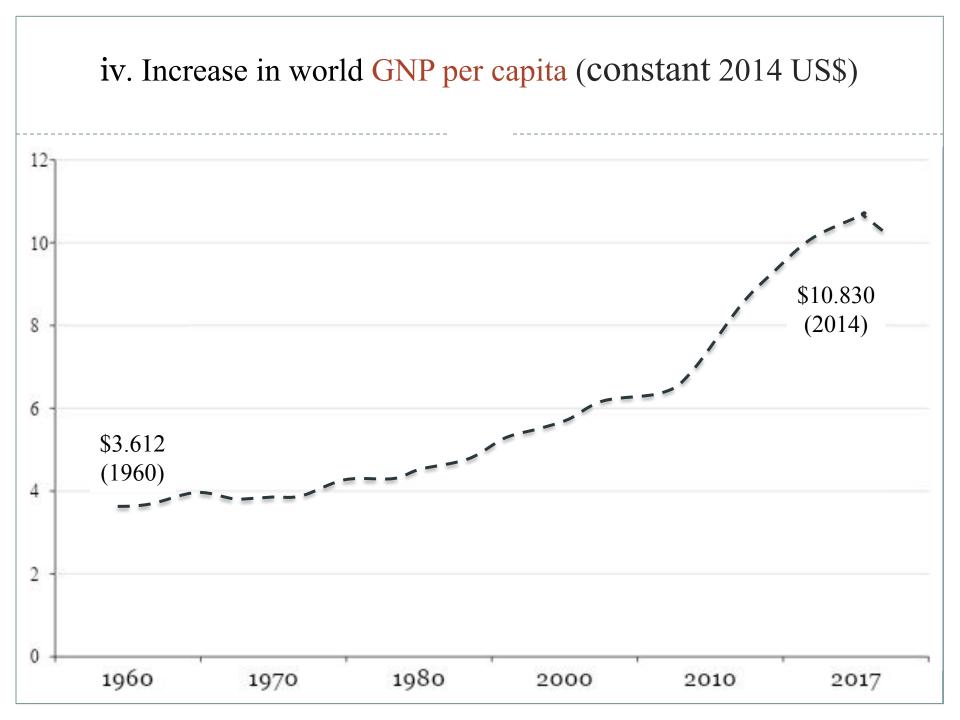
### ii. Population aging

Population aged 60-79 years and aged 80 years or over by development group, 2000, 2015, 2030 and 2050

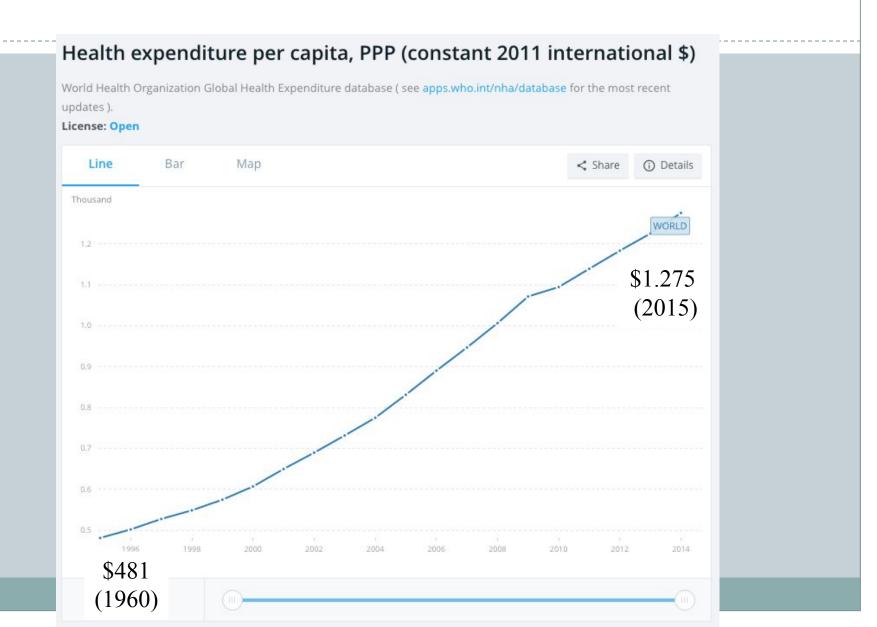


Data source: United Nations (2015). World Population Prospects: The 2015 Revision.

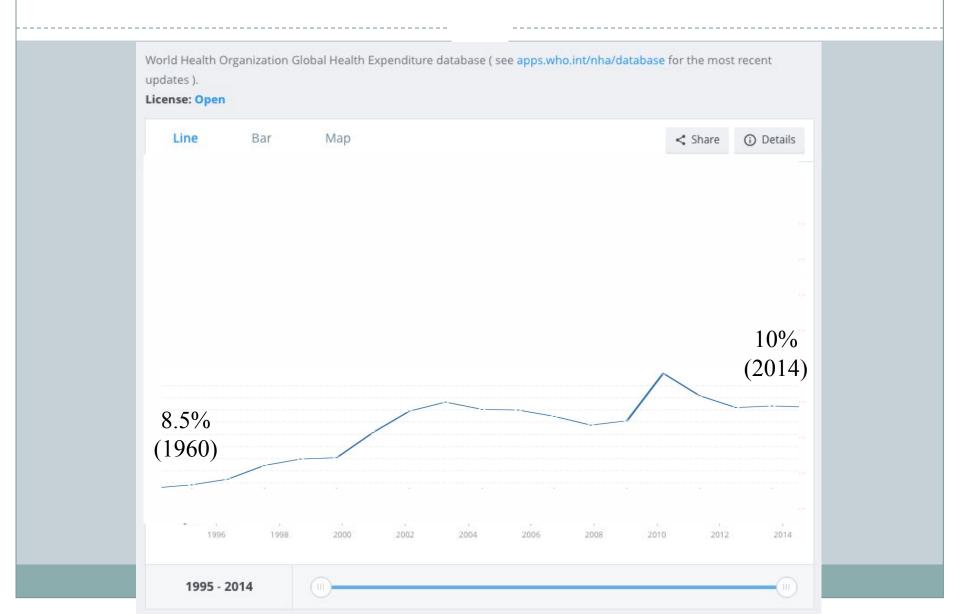




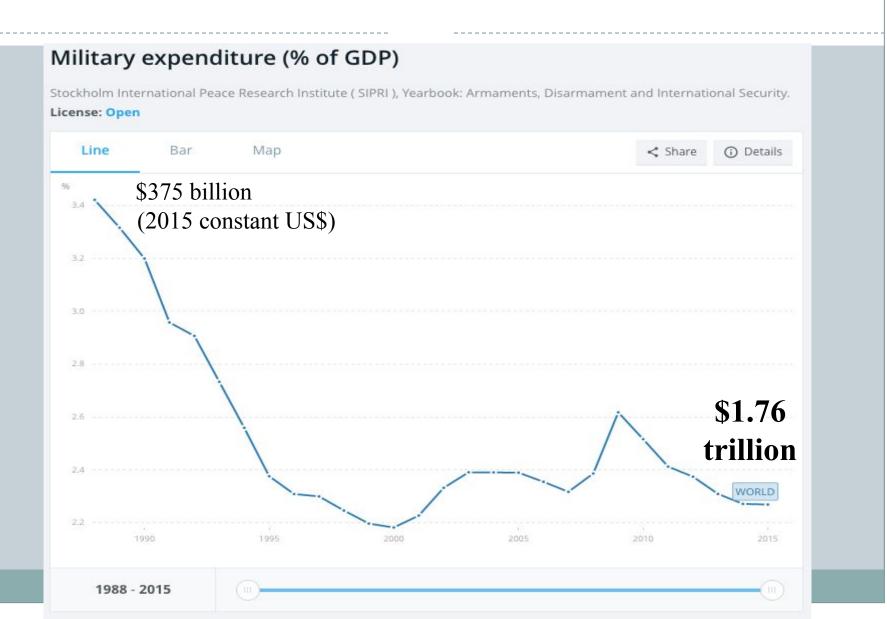
### v. Increase in world health expenditure per capita



### vi. Increase in world health expenditure (% of GNP)



### vii. Increase in world actual military expenditure

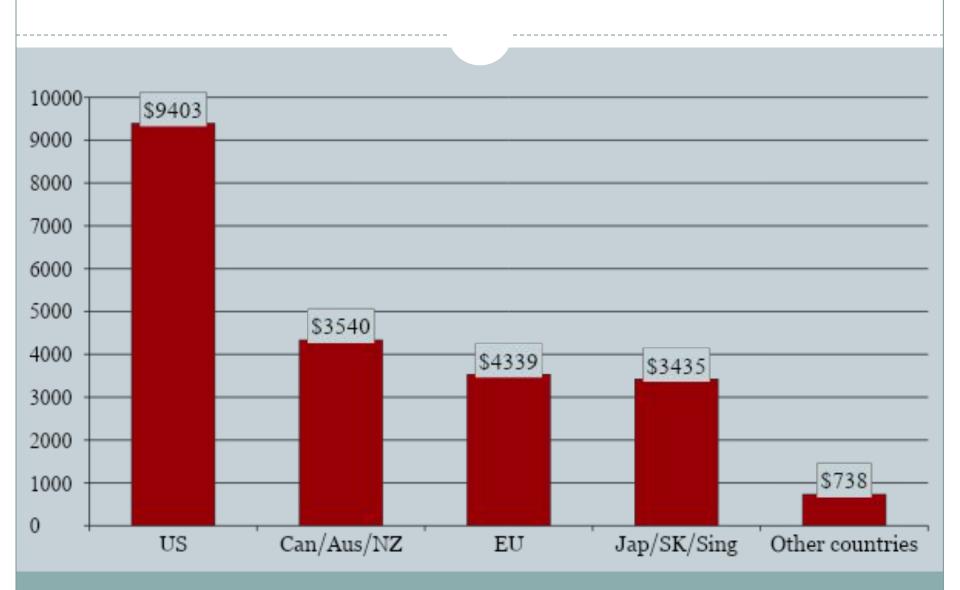


#### II.b. BREAKDOWN OF HEALTH EXPENDITURES IN THE WORLD

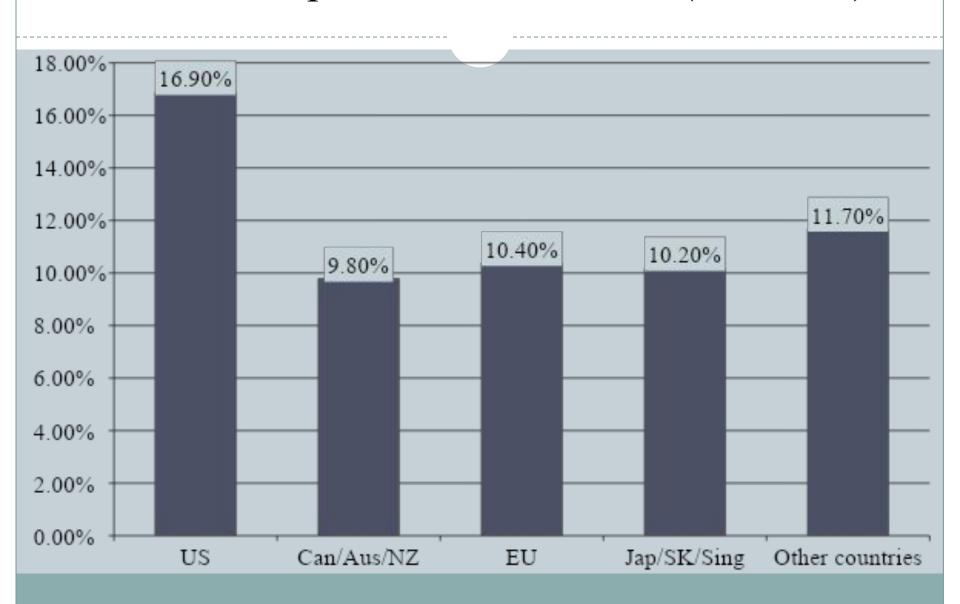




### i. Average Health Expenditure Per Capita

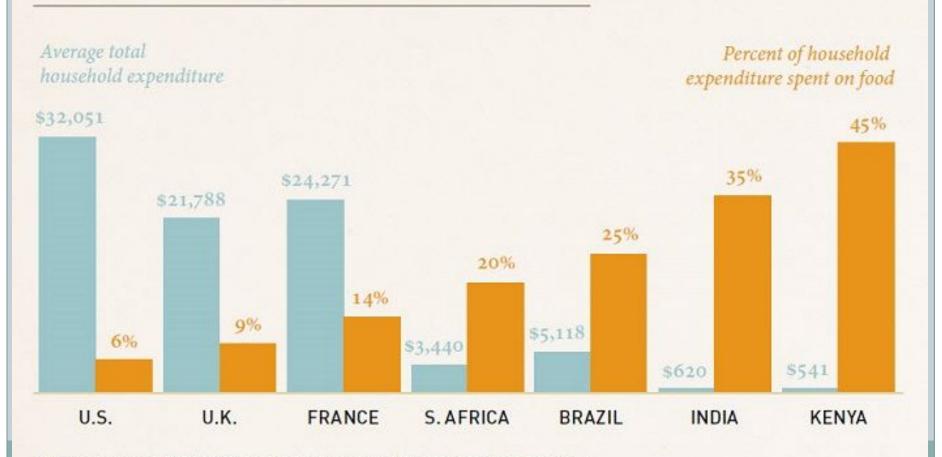


### ii. Health expenditure % of GNP (HE/GNP)



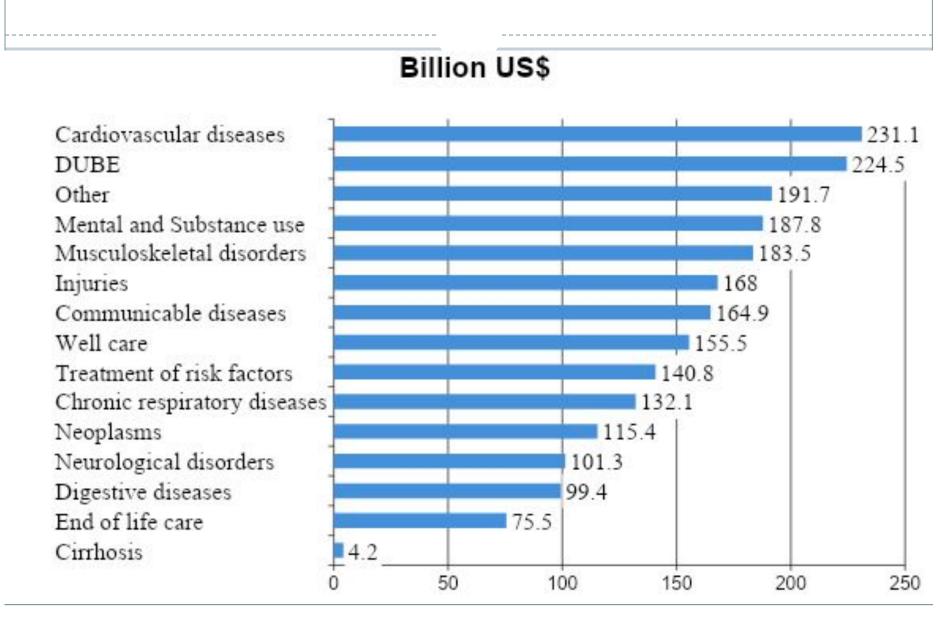
### iii. Expenditure priorities in poor countries

## The Poor Spend a High Percentage of Their Income on Food



Sources: World Bank 2009; U.S. Department of Agriculture 2009; Euromonitor International

### iv. US health expenditures by disease group



### v. UK health expenditures by disease group

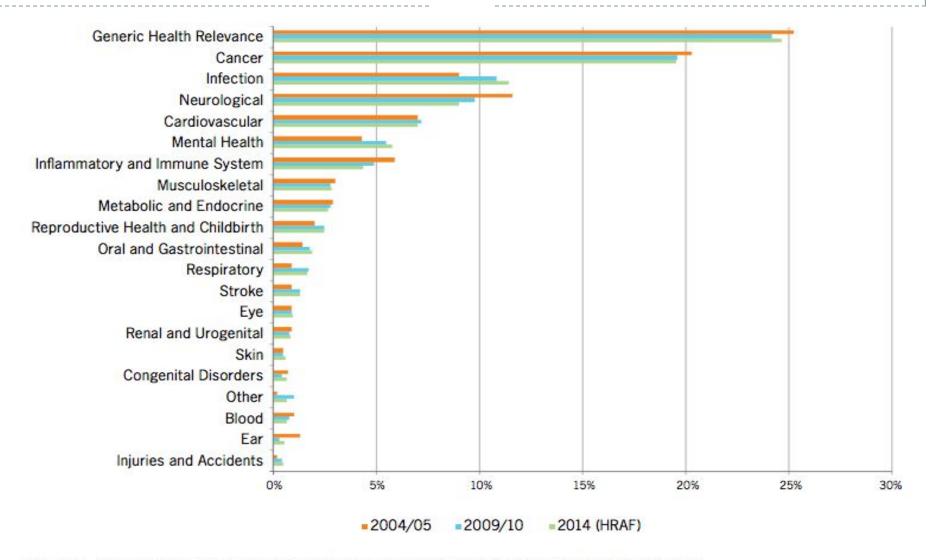
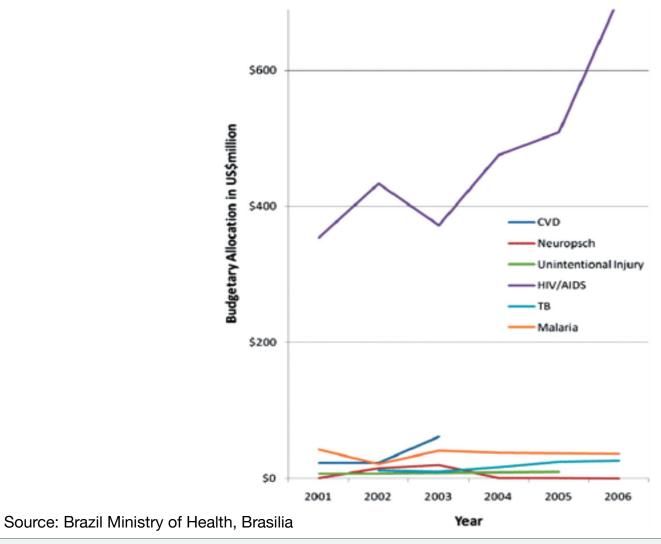
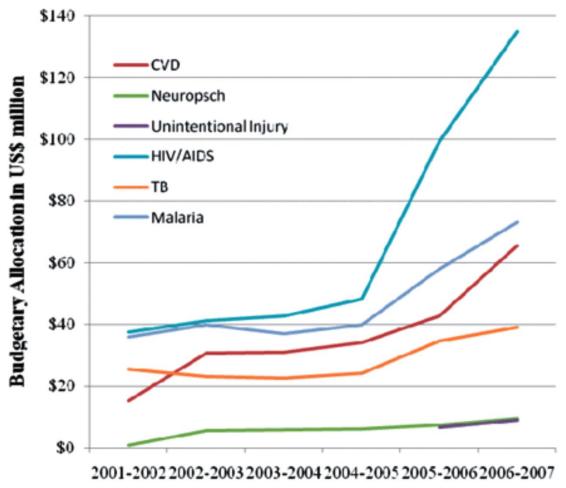


Figure 9 - Proportion of combined spend by health category for 2004/05, 2009/10 and 2014 (HRAF, 12 funders).

# vi. Brazil's budgetary allocation for health (US\$ million) from 2001 to 2006



# vii. India's budgetary allocation for health (US\$ million) from 2001 to 2007



Year
Source: India Ministry of Finance, available at: http://indiabudget.nic.in

### viii. Summary

- Population
  - Thanks to relatively inexpensive (first) technology transference
- □ □ Gross World Product
- □ □ □ World Health Expenditure
- Most of the world's health expenditure is in wealthy countries
- Most of that expenditure is for chronic diseases in a mature population
- Poorer countries spend more money on food, infectious disease and primary care

### II.c. NEUROSURGERY EXPENDITURES

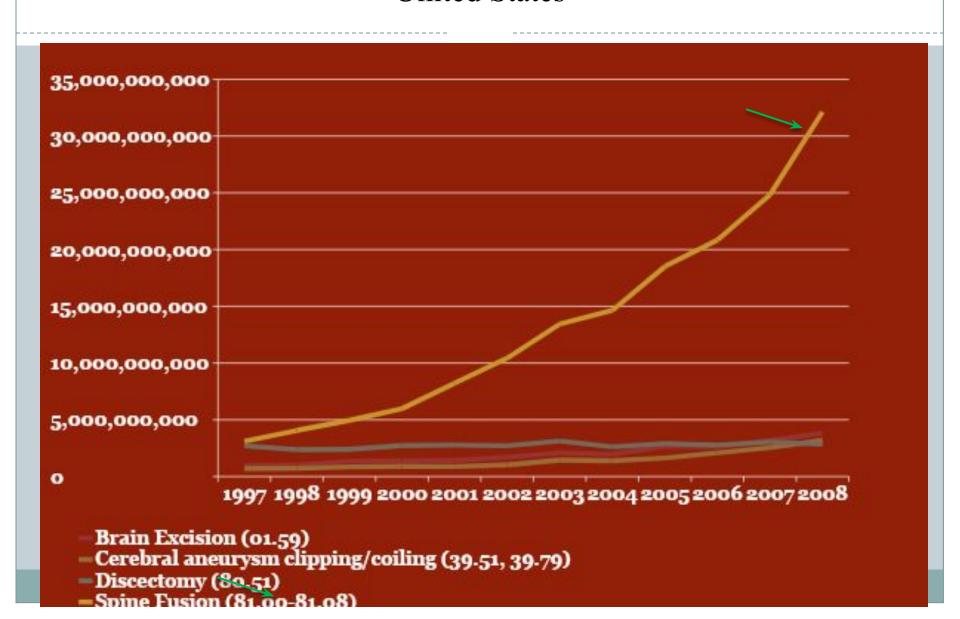


### i. Neurosurgery Expenditures in the United States

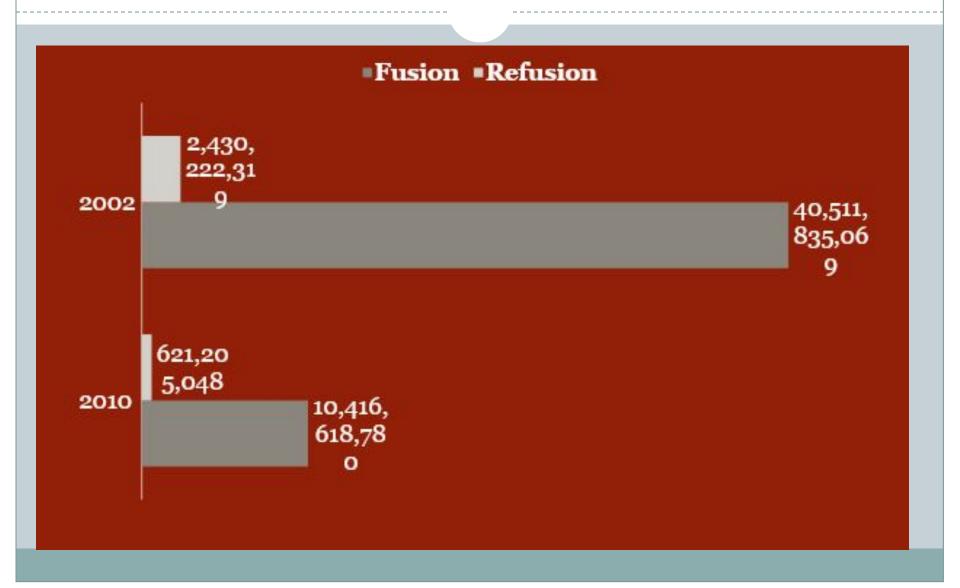
#### Calculated using Center of Medicare & Medicaid data

- Medicare payments to Neurosurgeons as salary in 2014 \$612.417.016
   Medicare makes 20% of total insurance expenditures
- Thus, total Neurosurgery salaries paid by Insurance \$3.062.085.080
   Neurosurgeon's salary makes 20% of all personnel expenditures
- Thus, total personnel expenditure is \$15.310.425.400 Insurance makes 74% of total health expenditures
- Thus, total Neurosurgeon Salary in US \$20.6 billion
- Salaries make 20% of total Health expenditure
- Thus, total Neurosurgery expenditure in US \$103 billion (1% of world health expenditure)

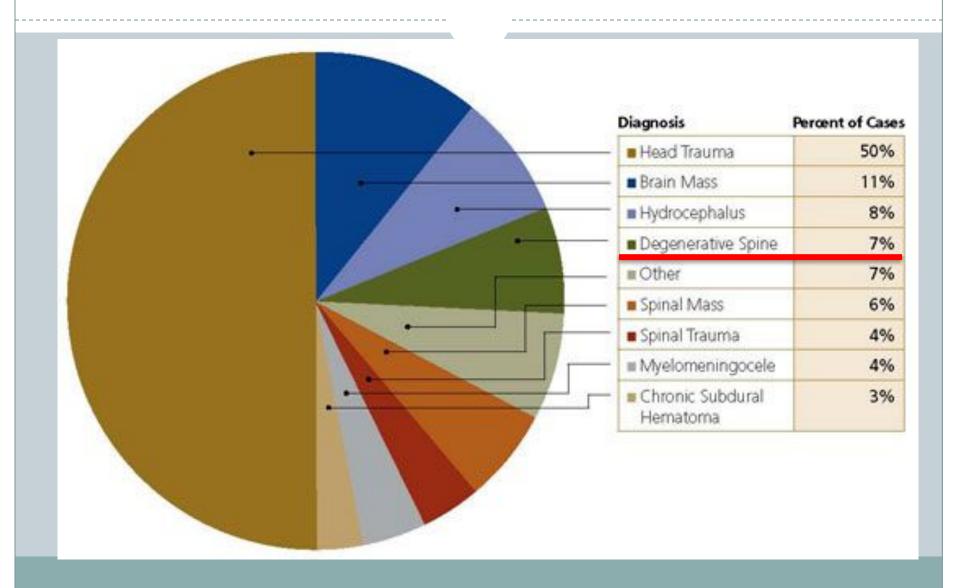
### ii. Aggregate Hospital Charges for Neurosurgical Procedures in the United States



### iii. The "National Bill" of All Spine De Novo Fusions and Refusions

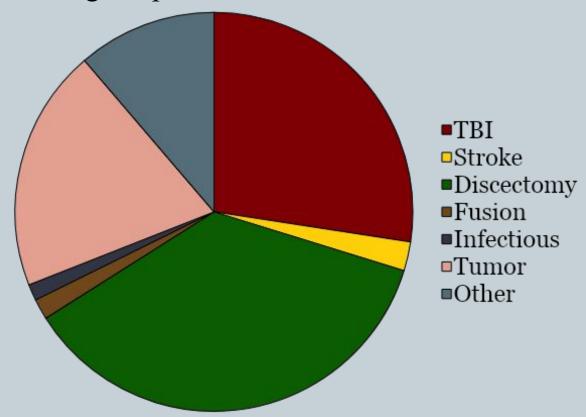


### iv. Neurosurgical Procedures in Ethiopia



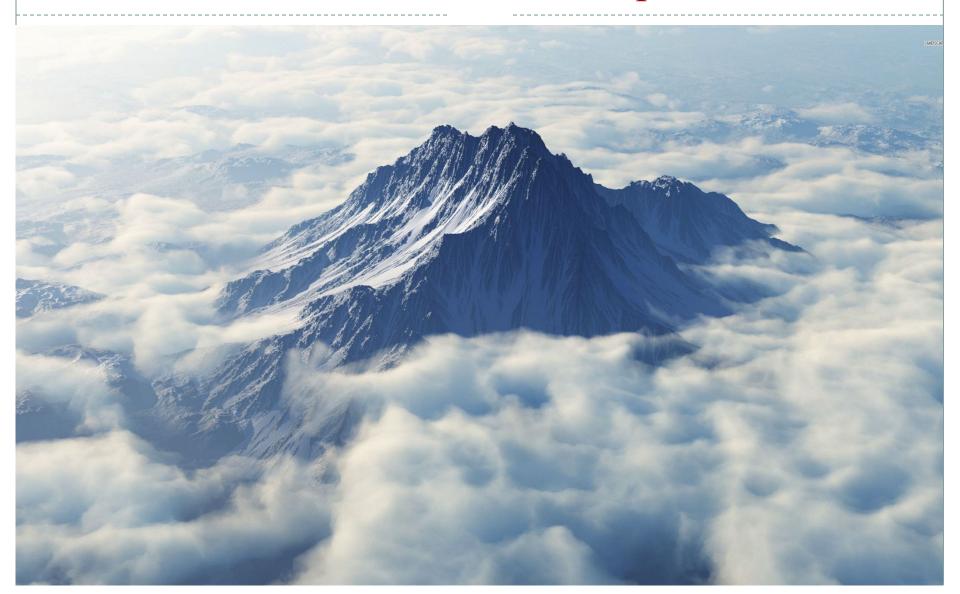
### v. Neurosurgery expenditures in Armenia

- \$2.8 million total neurosurgery expenditure in 2015\*
   \*there are substantial direct unregistered payments to doctors
- 3.101 Neurosurgical procedures in 2015

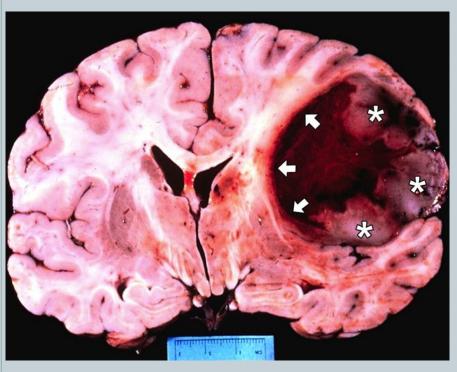


### III. THE VIEW FROM MT. OLYMPUS

-Resource reallocation – a comparative exercise



### Glioblastomas vs ruptured intracranial aneurysms

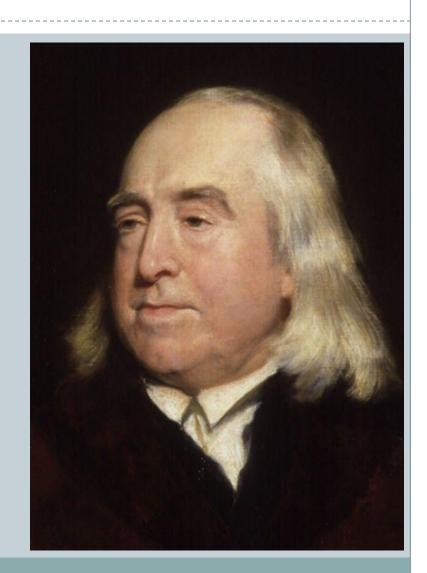




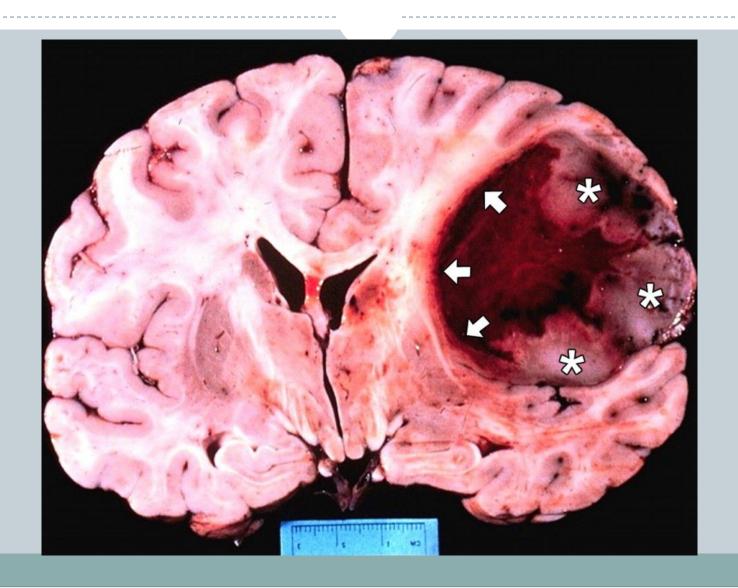
### Jeremy Bentham

Greatest good for the greatest number

1748 - 1832

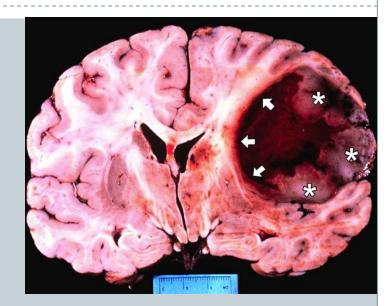


### III.a. Glioblastomas

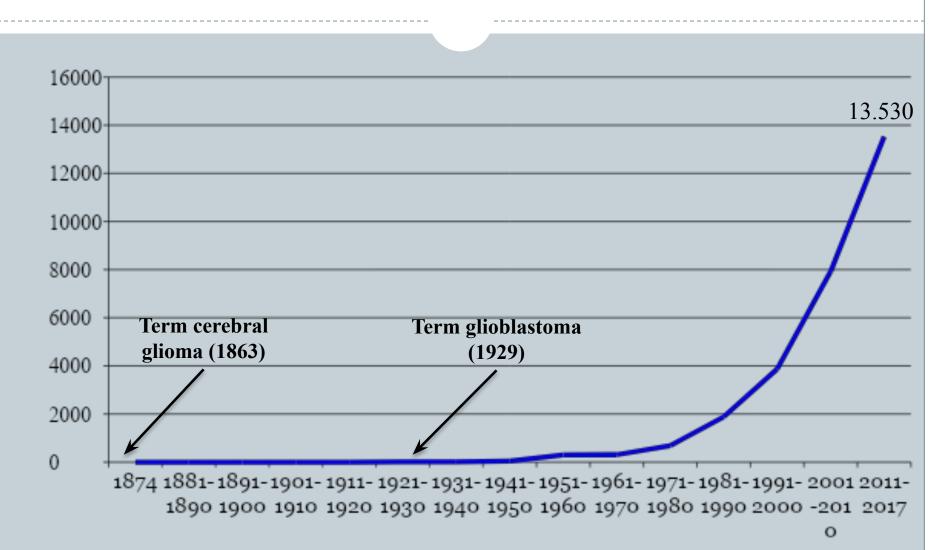


### III.a. Glioblastomas

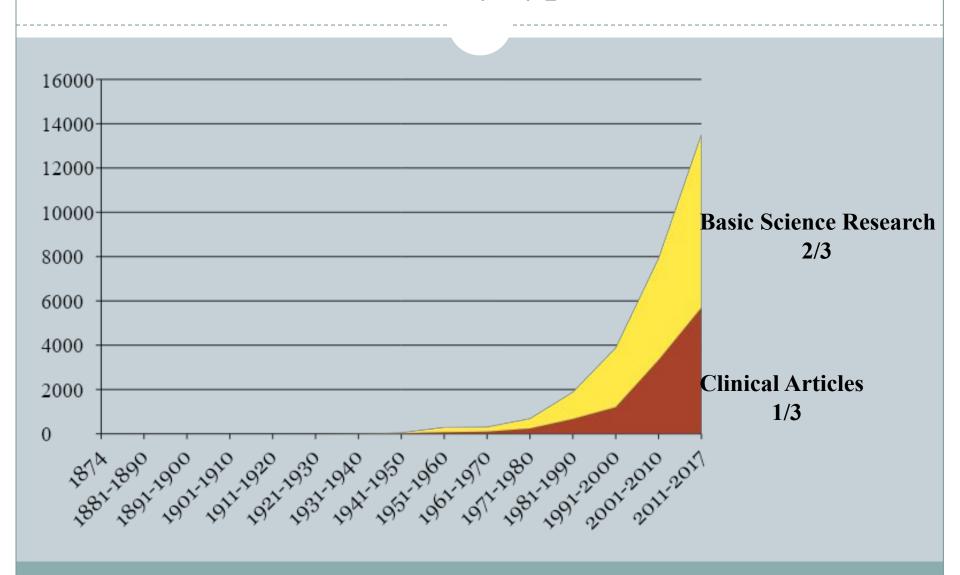
- i. Published articles
- ii. Types of published articles
- iii. Clinical Guidelines/ Standard of care
- iv. Main countries publishing guidelines
- v. Most recent guidelines
- vi. Summary of guidelines
- vii. Guideline limitations
- viii. A little math....
  - 1) Number of Glioblastoma cases in the world
  - 2) How many people with Glioblastoma are NOT receiving the standard of care?



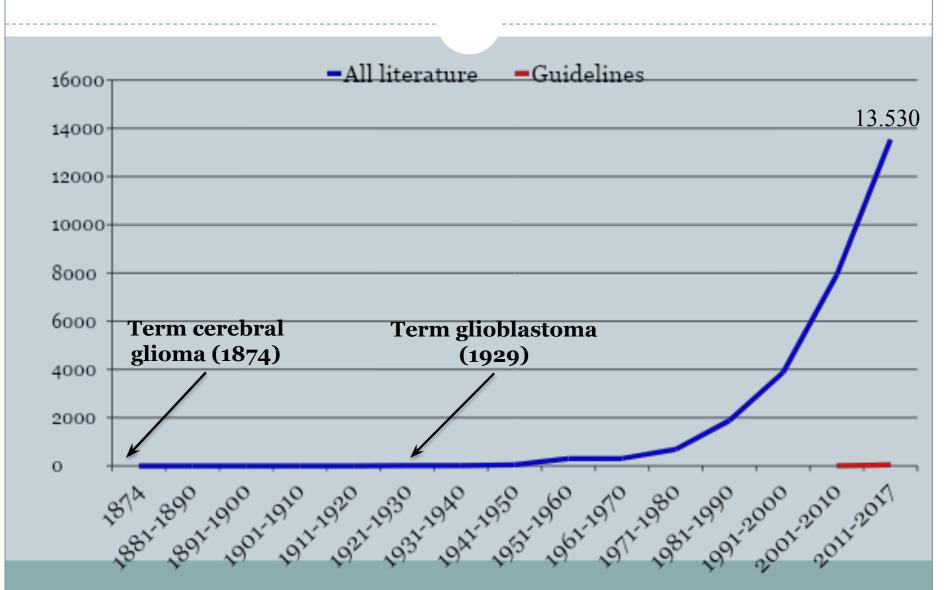
# i. Cited articles related to cerebral gliomas or glioblastomas – 28.577



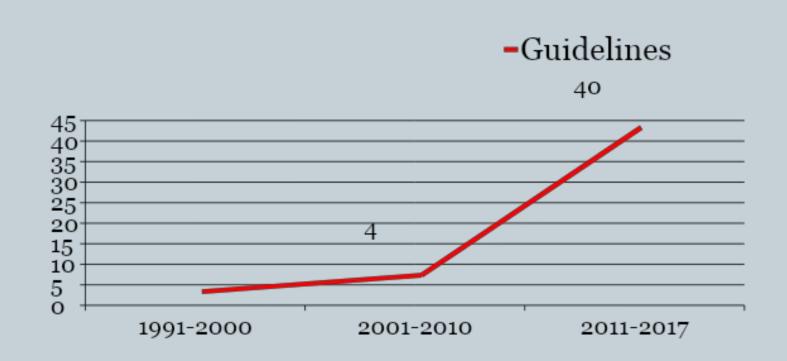
### ii. Cited articles by type of research



# iii. Articles related to cerebral gliomas or glioblastomas – 28.577



# iii. Articles related to clinical <u>guidelines</u> for treatment of gliomas or glioblastomas – 40



# iv. Most published guidelines for GBM are from US or Western Europe

	USA	UK	Switzerland	Germany	Netherlands	Spain	Belgium
2006	1						
2007		1	1				
2008		1					
2009							
2010		1				1	
2011	1						
2012	5	1					1
2013	2						
2014	9		1	1	1		
2015	7	1	1				
2016	3			1	1		
2017	1						



#### National Comprehensive Cancer Network

Your Best Resource in the Fight Against Cancer®

National Comprehensive Cancer Network (NCCN) is an alliance of 27 cancer centers in the United States, most of which are designated by the National Cancer Institute as comprehensive cancer centers.



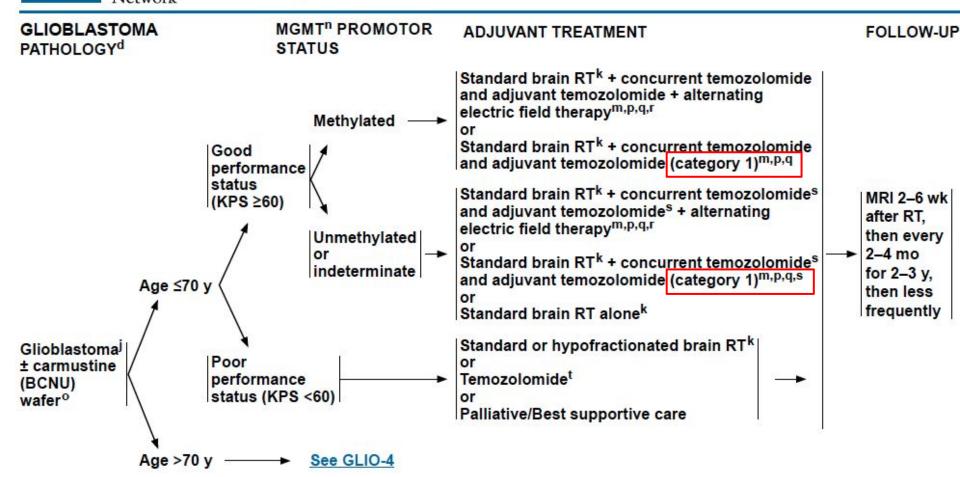
#### v. GBM Guidelines – Adjuvant Treatment (Age <70)

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#### NCCN Guidelines Version 1.2016 Anaplastic Gliomas<sup>a</sup>/Glioblastoma

NCCN C



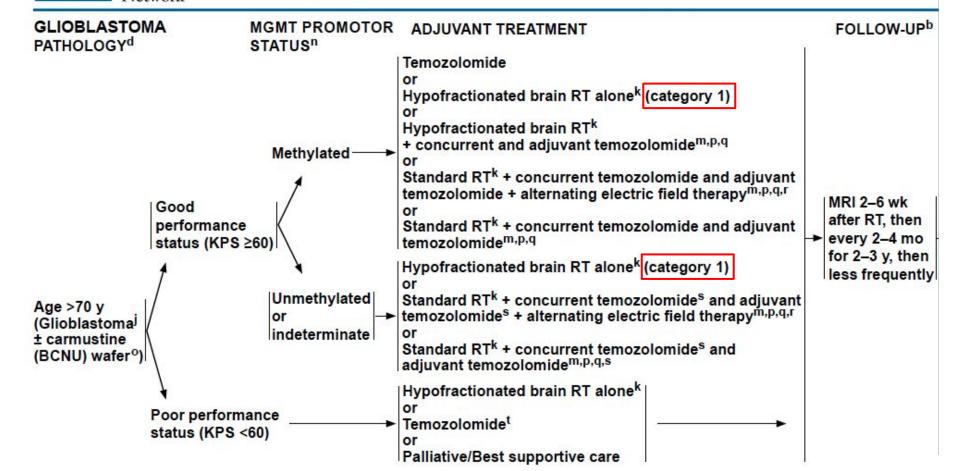
#### v. GBM Guidelines – Adjuvant Treatment (Age >70)

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#### NCCN Guidelines Version 1.2016 Anaplastic Gliomas<sup>a</sup>/Glioblastoma

NCCN G



## vi. Summary of published clinical guidelines

#### Generally addressed issues

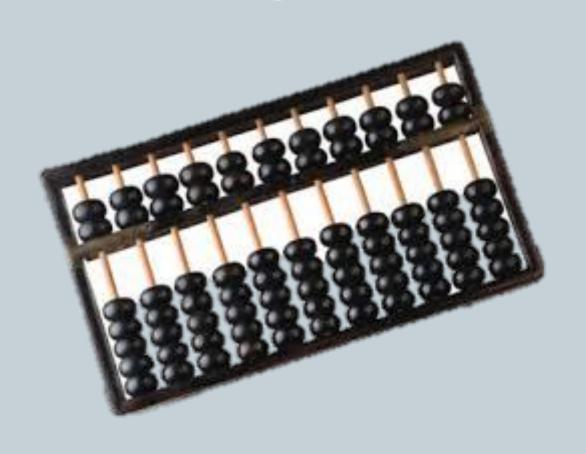
- Who to treat?
- Role of surgery
- Role of chemotherapy
- Role of radiation
- Management of recurrence

#### vii. Guideline limitations

- Applicable only to developed countries (Western world)
- Only 3 out of 40 guidelines support any of the recommendations with partial Grade I evidence
- No Grade I evidence before 2014
- There is no any reference to <u>cost</u>
- Often incompletely address treatment algorythm

### viii. And now a little math...

------ 45



• The question we ask: How many people in the world are actually getting the standard of care?

• If not, why so?

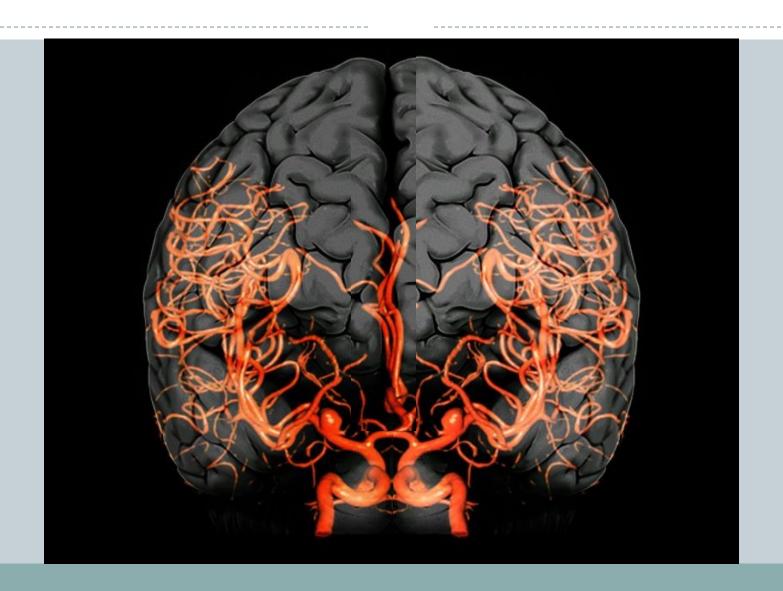
### (1) Number of Glioblastoma cases in the world

- Even in developed places there are limitations to access
- US 13,000 new GBM cases
- US population 320,000,000
- Number of people over 40 is about 130,000,000
- Incidence per older person in the US is 1/10,000
- More or less the world population is about 7.5 billion
- World population over 40 is about 2.8 billion
- Thus if the incidence is 1/10,000 **280,000 people** in the world that will be yearly affected by GBM

# (2) How Many People are NOT Receiving the "Standard of Care"/treatment for Glioblastomas

- How many people are receiving standards of care world wide at the present?
- Estimated yearly incidence of GBM worldwide is 280,000
- $\bullet$  280,000 13,000(US) 41,000 (EU) = 226,000
- About 5% of the remainder of world's population is getting appropriate treatment, which is 11,500
- <u>226,000 11,500 = **214,500**</u>

## III.b. Ruptured intracranial aneurysms



## III.b. Ruptured Intracranial Aneurysms

- i. Published articles
- ii. Clinical Guidelines/ Standard of care
- iii. Main countries publishing guidelines
- iv. Summary of guidelines
- v. Guideline limitations
- vi. A little more math....
  - 1) Number of Ruptured Aneurysms worldwide
  - 2) How Many People are NOT Receiving the Standard of Care?



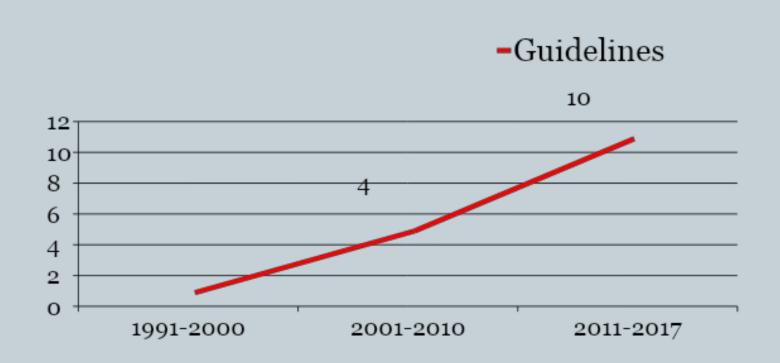
# i. Cited articles related to ruptured intracranial aneurysms - 26.742



ii. Cited articles related to ruptured intracranial aneurysms (26.742) and Clinical Guidelines (14)



# ii. Articles related to <u>clinical guidelines</u> for treatment of ruptured intracranial aneurysms - 12



# iii. Most published guidelines for ruptured aneurysms are from US

	USA	UK	Australia	South Korea	Spain	Japan
2002						1
2006	1				1	
2010					1	
2011					1	
2012	1					
2014				1		
2015	2		1			
2016	2	1				1

### iv. Summary of published clinical guidelines

#### Generally addressed issues

- Risk factors for and prevention of aSAH
- Natural history and outcome of aSAH
- Clinical manifestations and diagnosis of aSAH
- Medical measures to prevent rebleeding after aSAH
- Surgical and endovascular methods of treatment of ruptured cerebral aneurysms
- Hospital characteristics and systems of care
- Anesthetic management during surgical and endovascular treatment
- Management of cerebral vasospasm/ hydrocephalus/ seizures/ medical complications after aSAH

#### v. Guideline limitations

- 2 out of 52 recommendations have Level A evidence
- There is no any reference to <u>cost</u>
- Based on a single large clinical trial
- Not all treatment algorithms are included

#### vi.1. Number of Ruptured Aneurysms worldwide

- Incidence 9.1/100.000
- Calculated number of new cases/year worldwide **673.849**
- How many people are receiving standards of care world wide at the present?
- $673.849 29.500(US) 47,000(EU) \approx 600,000$
- About 5% of the remainder of world's population is getting appropriate treatment, which is 30.000
- <u>600,000 -30,000 = **570.000**</u>

# vi.2. How Many People are NOT Receiving the Standard of Care for Glioblastoma and Ruptured Aneurysms worldwide

● Glioblastomas – 214.500

■ Ruptured intracranial aneurysms – 570.000

#### IV

• Why not <u>worldwide access</u> to <u>standard of care</u>/ clinical guidelines/ treatment of <u>Glioblastomas</u> and <u>Ruptured Aneurysms</u>?

Supply?



Cost?

## **SUPPLY**



## IV.a. Worldwide general supply difficulties...

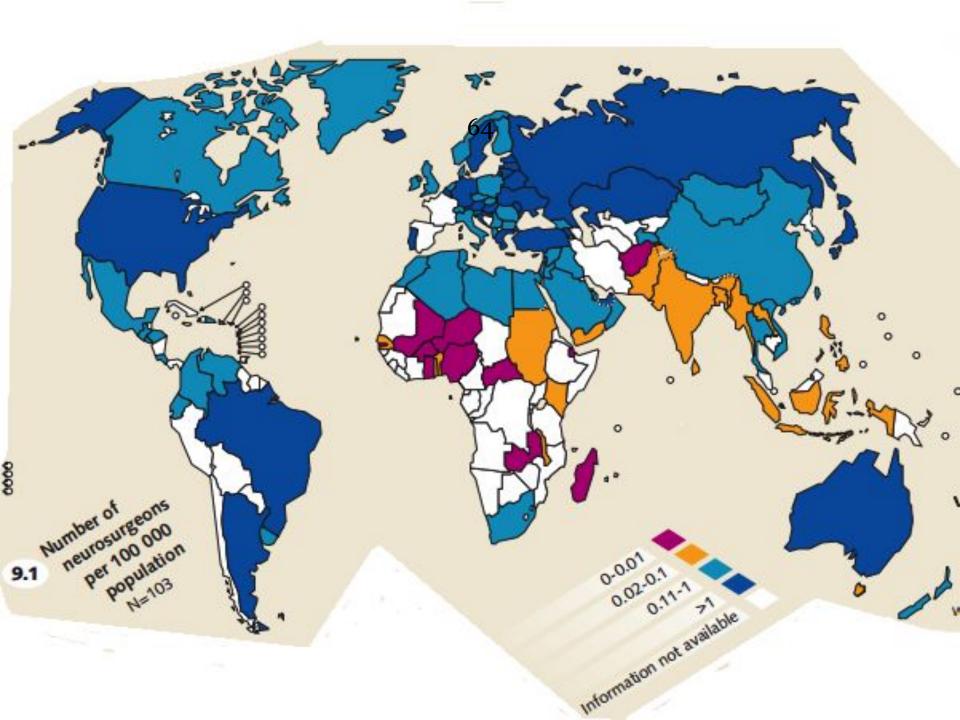
- i. Mal-Supply of Physician Work-force
- ii. Neurosurgeons Around the World
- iii. Thailand a personal story
- iv. Absence of radiographic diagnostic facilities
  - v. Tertiary Care: China

# i. Mal-Supply of Physician Work-force



## ii. Supply: Neurosurgeons Around the World

- A total of 33 193 neurosurgeons are reported to be available in 103 countries in 2004.
- The median number of neurosurgeons per 100 000 population is:
- 0.01 in Africa
- 0.76 in the Americas
- 0.03 in South-East Asia
- 0.37 in the Eastern Mediterranean
- 1.02 in Europe



## iii. Thailand - a personal story





## iv. Absence of radiographic diagnostic facilities

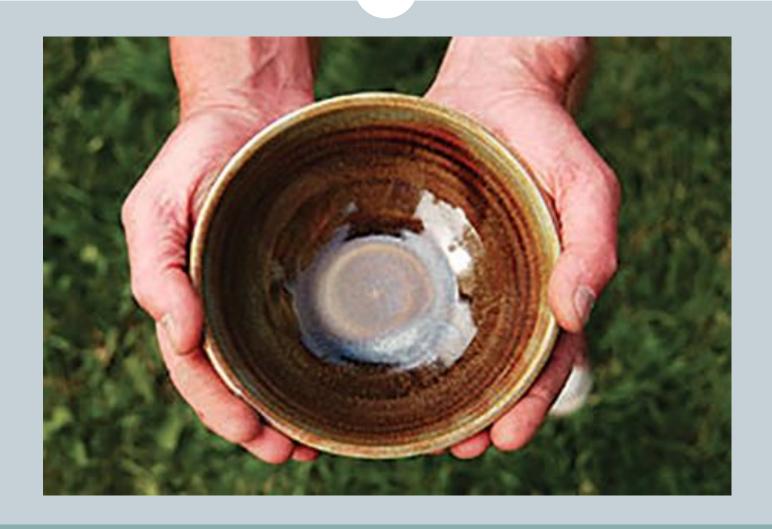


Table 123 (page 1 of 2). Number of magnetic resonance imaging (MRI) units and computed tomography (CT) scanners: Selected countries, selected years 1990–2009

Updated data when available, Excel, PDF, and more data years: http://www.cdc.gov/nchs/hus/contents2011.htm#123.

[Data are based on reporting by Organisation for Economic Co-operation and Development (OECD) countries]

Country	1990	1995	2000	2007	2008	2009	1990	1995	2000	2007	2008	2009
	Number of MRI units per million population				Number of CT scanners per million population							
Australia 1	0.6	2.9	†3.5	5.1	5.6	5.9	13.8	††20.5	††26.1		•••	††38.7
Austria			11.0	17.7	18.0	18.4			26.1	30.0	29.6	29.3
Canada <sup>2</sup>	0.7	1.4	2.5	6.7		8.0	7.2	8.0		12.7		13.9
Czech Republic <sup>3</sup>		1.0	1.7	4.4	5.0	5.7		6.7	9.6	12.9	13.3	14.1
Denmark			5.4			15.4			11.4	18.5	21.5	23.7
Estonia				5.2	8.2	7.5				11.2	14.9	14.9
Finland	1.8	4.3	9.9	15.3	16.2	16.9	9.8	11.8	13.5	16.5		20.4
France			1.7	5.5	6.1	6.5			7.0	10.4	10.9	11.1
Greece				17.9	19.6	21.7				29.0	30.6	33.8
Hungary 4	0.1	1.0	1.8	2.8	2.8	2.8	1.9	4.6	5.7	7.3	7.1	7.2
celand	3.9	7.5	10.7	19.3	18.8	21.9	11.8	18.7	21.3	32.1	31.3	34.5
and a seed				8.5	9.0	†11.9	4.3		-1.0	14.3	14.5	15.3
		0.9	1.4	2.0	2.1	1.9	4.5	1.6	5.7	8.5	8.8	9.4
Srael		307773	7.8	18.5	20.1	21.6			21.1	30.1	30.9	31.7
Italy <sup>6</sup>	0.4	• • • •				4111	55.0	***				The state of
Japan	6.1	0.5	0.0	40.4	43.1	44.0	55.2	00.0	25.0	07.4	97.3	00.0
Luxembourg	2.6	2.5	2.3	10.4	12.4	14.2	5.2	26.9	25.2	27.1	26.9	26.3
Mexico				1.5	1.7	1.9				4.0	4.2	4.3
Netherlands 8	0.9	3.9	•••	7.6	10.4	11.0	7.3	• • • •	• • • •	7.8	10.3	11.3
New Zealand				8.8	9.6	9.7	3.5		8.8	12.3	12.4	14.6
Poland				2.7	2.9	3.7			4.4	9.7	10.9	12.4
Portugal <sup>9</sup>				8.9						26.0		
Republic of Korea		3.9	5.4	16.0	17.6	19.0		15.5	28.4	37.1	36.8	37.1
Slovak Republic 10			1.1	5.7	6.1	6.1				13.7	13.7	13.3
Slovenia				3.5	4.5	4.5				10.9	12.4	11.9
Switzerland										††31.4	††32.0	††32.8
Turkey				5.4	†7.2	8.9	1.6			7.7	†10.6	11.6
United Kingdom <sup>11</sup>			<b>††5.6</b>		115.6				<b>††5.3</b>		117.4	
United States <sup>12</sup>		12.3	110.0	25.9	110.0				110.0	34.3	110.00	

## v. Tertiary Care: China

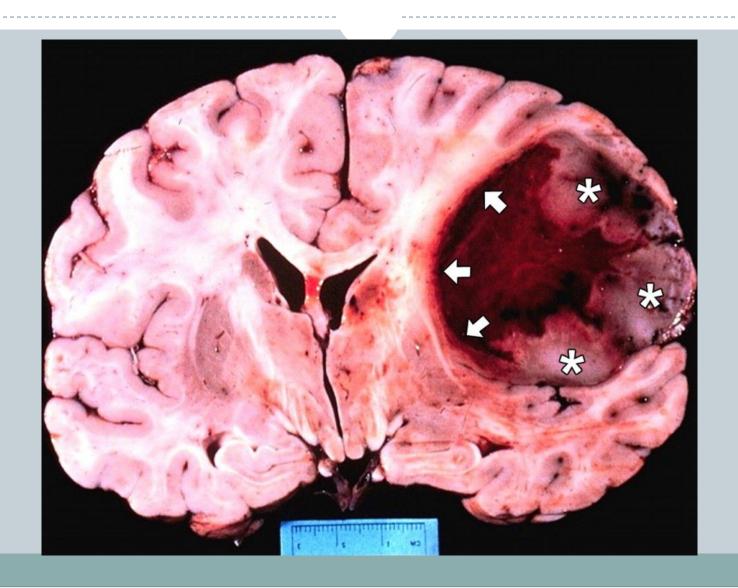


# v. China: Tertiary Hospital Care Imbalanced

	Name of Hospital	Total Score	Research Score	Total Score
1	Beijing Union hospital	80.00	10.95	90.95
2	Sichuan Huaxi hospital	69.42	20.00	89.42
3	PLA General hospital	58.16	14.52	72.68
4	Ruijin Shanghai Jiaotong Univ.	42.92	11.90	54.83
5	Xijing 4 <sup>th</sup> Military Medical Univ.	33.99	12.99	46.98
6	Zhongshan, Fudan Univ.	32.83	13.66	46.48
7	Huashan, Fudan Univ.	34.70	10.23	44.93
8	First hospital, Peking Univ.	32.43	9.10	41.53
9	People's Hospital, Peking Univ.	29.25	8.88	38.13
10	Tongji hospital,	26.43	11.13	37.56
11	First Hospital, Zhongshan Univ.	25.15	12.26	37.41
12	Peking Univ. 3 <sup>rd</sup> Hospital,	18.42	9.29	27.72
13	Union hospital, Tongji Med. College	15.61	10.85	26.45
14	First hospital, Zhejiang Univ.	13.13	13.09	26.22
15	Changhai, 2 <sup>nd</sup> Military Med. Univ.	14.35	11.25	25.60
16	First hospital, China Med. Univ.	15.05	10.17	25.21
17	Fuwai Heart-Vascular Disease Hospital	12.75	12.43	25.16
18	Xiangya 2 <sup>nd</sup> hospital. Zhongnan Univ.	13.53	9.06	22.59
19	Renji hospital, Jiaotong Univ.	12.82	9.64	22.46
20	Southern Hospital, Southern Medical Unv.	12.98	9.32	22.29

Table 6. Top 20 hospitals in China (2011)

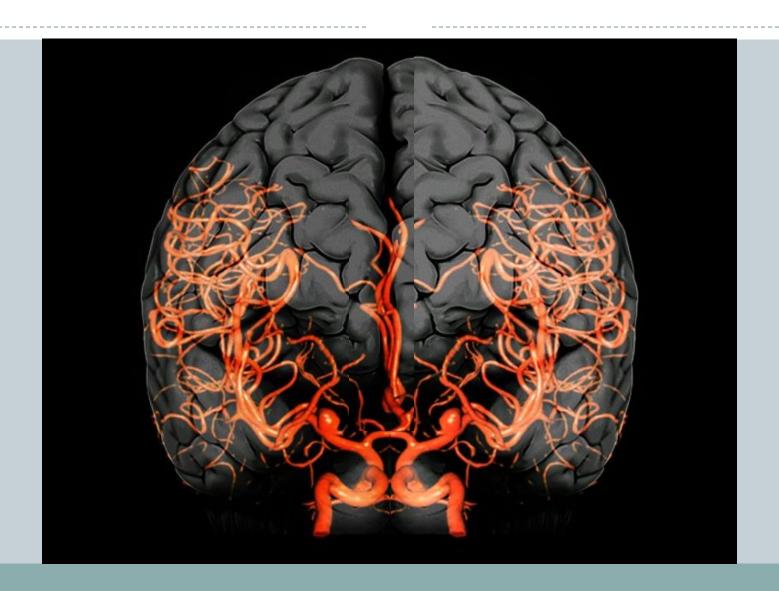
# IV.b. Supply difficulties for Glioblastomas



## i. Neuropathologists around the world

- American Association of Neuropathologists has 800 members from around the world
- Number of neuropathologists in UK: 98

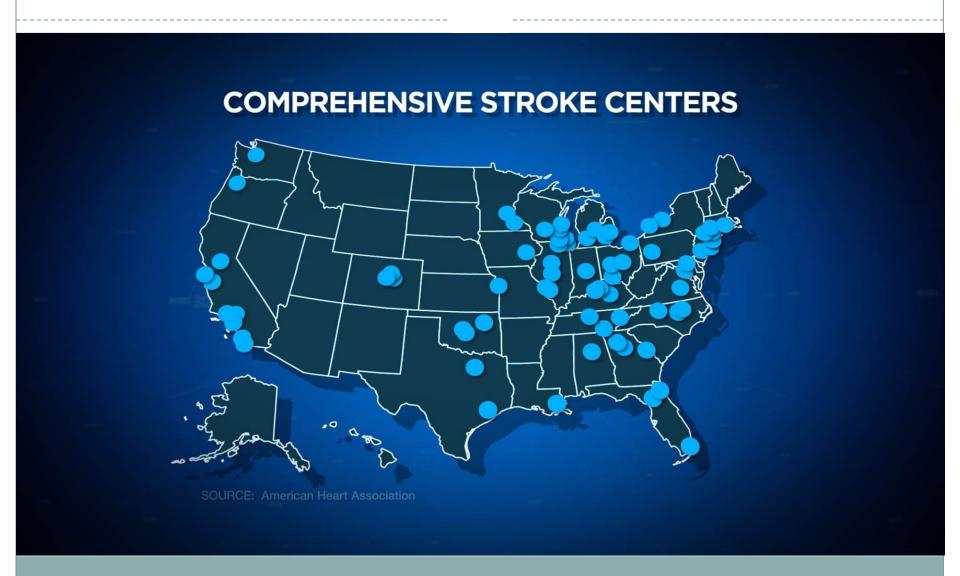
## IV.c. Supply difficulties for ruptured aneurysms



### i. Interventional Neuroradiologists

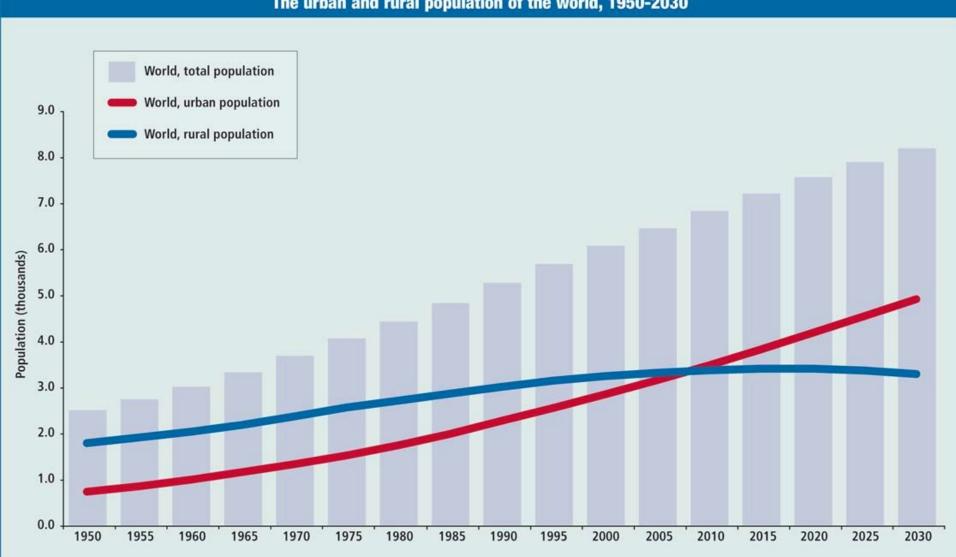
 The number of physicians in the United States with an active interest and special competency in this field is estimated to be around 600

## ii. 103 Comprehensive Stroke Centers in US

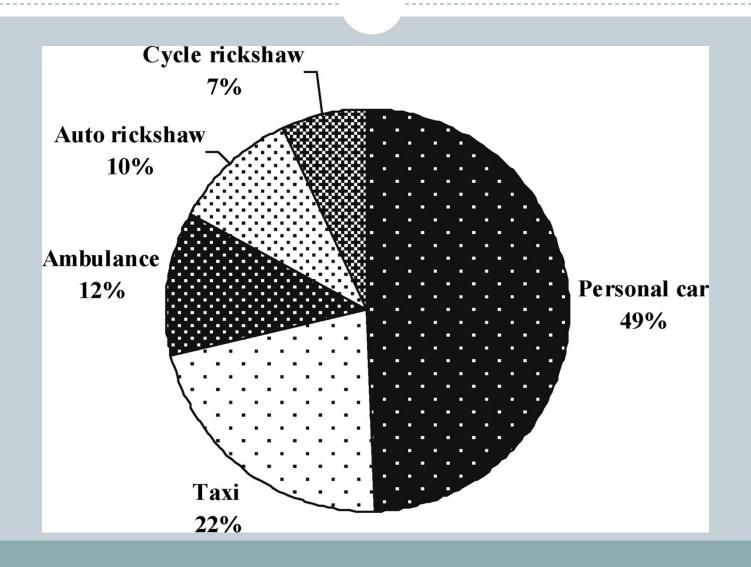


### iii. World Urbanization (United Nations)

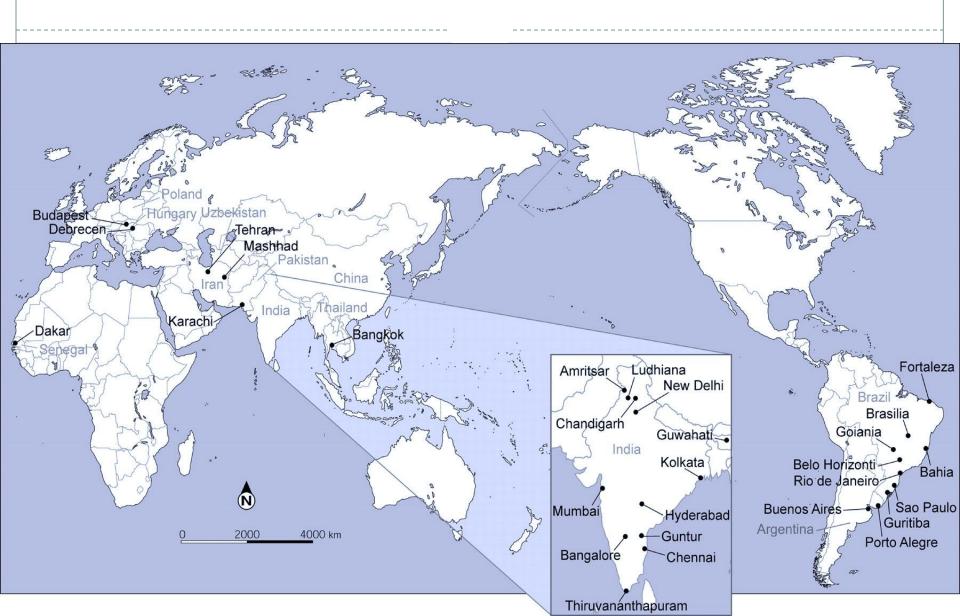
The urban and rural population of the world, 1950-2030



### iv. Modes of transport of stroke patients in India



## v. Stroke centers in developing world



## COST



#### IV.d. Cost issues for Glioblastomas\*

- i. US Cost data
- ii. EU Cost data
- iii. Amount of GNP spent on healthcare
- iv. Developing world cost data
  - v. How much would it cost to treat Glioblastomas in this population?
- vi. % of people who can independently pay for reduced costs of GBM



#### i. US Cost data

• Currently in the U.S., the cost of temozolomide (Temodar, Temodal) treatment is over \$10,000 per month. If you add in Gliadel Wafers, it adds another \$10,000 to your initial resection surgery. And treatment with Avastin + CPT-11 can be over \$50,000 per year.

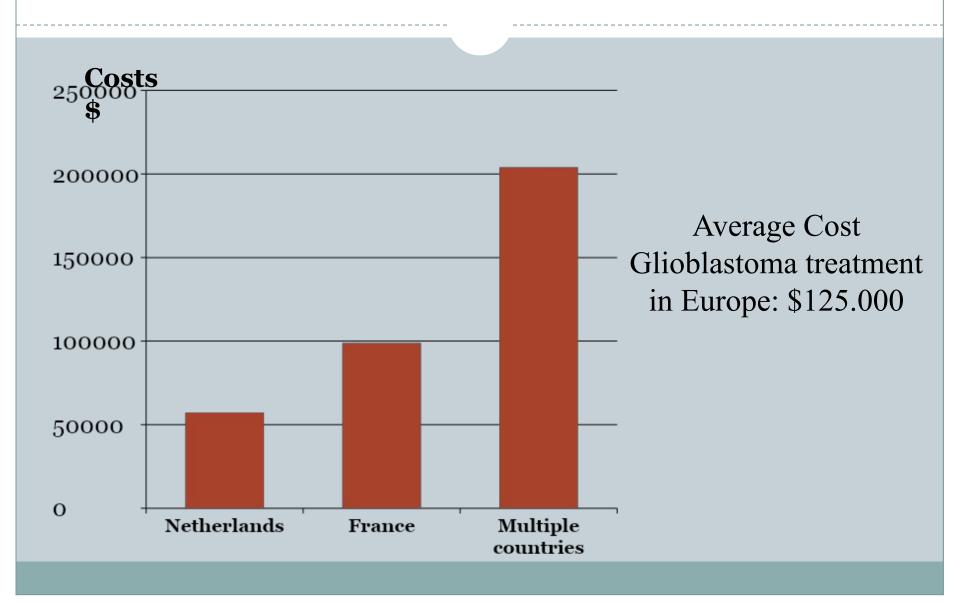
#### Cost data in the USA:

- $\Box$  Dx
  - MRI/CT-Scan
  - **\$1200**
  - Biopsy
- Neuropathology
- Craniotomy
  - □ \$50,000-\$70,000
- Radiation
  - □ \$7,000**-**\$20,000

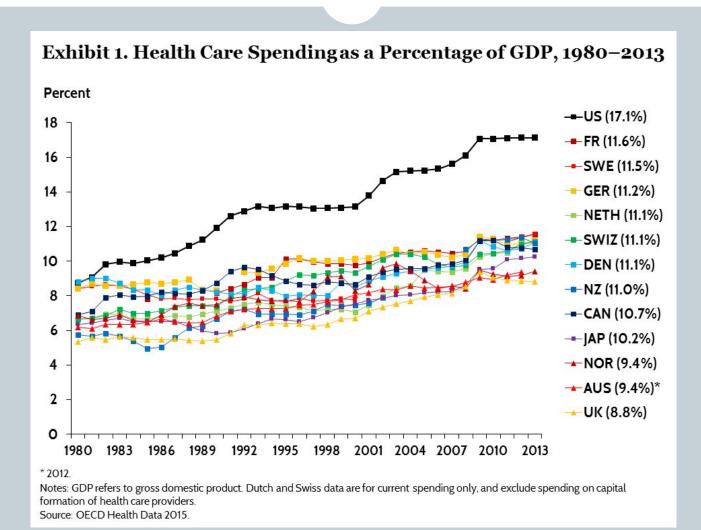
#### ii. US Cost data, cont.

- Chemotherapy with the drug temozolomide is the current standard of treatment for GBM
  - □ \$51/5 capsules (6 times treatment)
- Hospice
  - □ \$150/day
- Meds
  - Seizure meds
  - Steroids
- Follow up scans
- Hospice Care
- Pain Management

#### ii. Average Glioblastoma Treatment Costs in Europe



#### ii. Amount of GNP Spent on Healthcare



#### ii. EU Cost data

- Thus we estimate the cost of GBM treament in the United
   States would be \$250,000
- Estmated cost of GBM treatment in EU is \$125.000

### iii. Developing world cost data

#### **USA**

- 1. Neurosurgery consultation \$378
- 2. Labs \$400
- 3. Imaging (MRI)  $-6 \times $600$
- 4. Hospital costs \$8.813
- 5. Surgery costs \$25.204
- 6. BCNU costs \$10.000
- 7. Genetic study \$550
- 8. Immunohistology \$416
- 9. Radiation costs \$11.520
- 10. TMZ (initial) \$12.136
- 11. TMZ (maintenance) \$2.041/month
- 12. Electric field therapy \$21.000/month
- 13. Other medications \$300/month
- 14. Hospice \$4.500/month

**Total:** \$73.017 + \$23.341/month + Hospice

#### **INDIA**

- 1. Neurosurgery Consultation \$11
- 2. Labs \$119
- 3. Imaging (MRI)  $-6 \times $400$
- 4. Hospital costs \$467
- 5. Surgery costs \$3.500
- 6. BCNU costs \$4.285
- 7. Genetic study \$243
- 8. Immunohistology \$124
- 9. Radiation costs \$6.898
- 10. TMZ (initial) \$1.410
- 11. TMZ (maintenance) \$365/month
- 12. Electric field therapy NA
- 13. Other medications \$180/month
- 14. Hospice NA

**Total:** \$19.457 + \$545/month + Hospice

# iv. How much would it cost to treat Glioblastomas in this population?

## For typical patient with an average length of survival net calculated cost will be:

• US  $-220,000 \times $250,000 = $55 \text{ billion}$ 

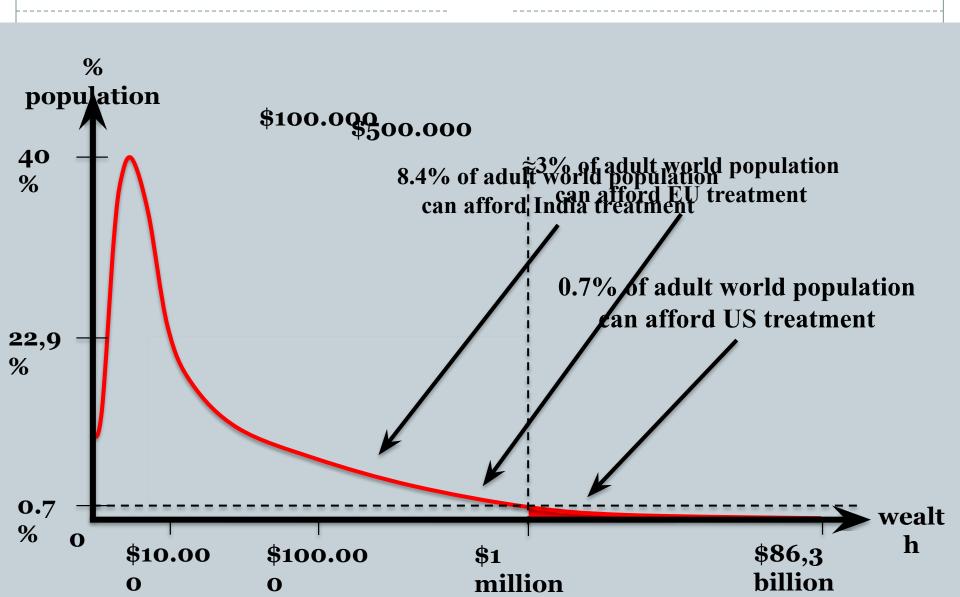
#### VS

•  $EU - 220,000 \times $125,000 = $27.5 \text{ billion}$ 

#### VS

• India -  $220,000 \times $27.414 = $6 \text{ billion}$ 

## v. % of people who can independently pay for reduced costs of GBM



### IV.e. Cost issues for ruptured aneurysms\*

- i. Costs of aneurysm treatment
- ii. How much would it cost to treat ruptured aneurysms in this population?
- iii. % of people who can independently pay for reduced costs of aneurysm treatment

\*Supply costs are not included in this section



# i. Costs of treatment of aneurysms are relatively well investigated



Richard Dodel et al., Stroke. 2010;41:2918-2923

## i. Costs of aneurysm treatment

- US cost \$42.000
- Averag EU cost \$36.000
- India cost \$6.000<sup>1</sup>

# ii. How much would it cost to treat ruptured aneurysms in this population?

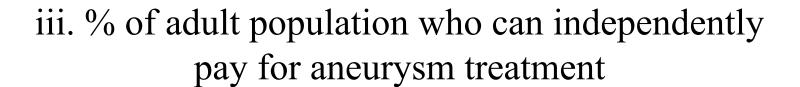
• US  $-570,000 \times $42,000 = $24 \text{ billion}$ 

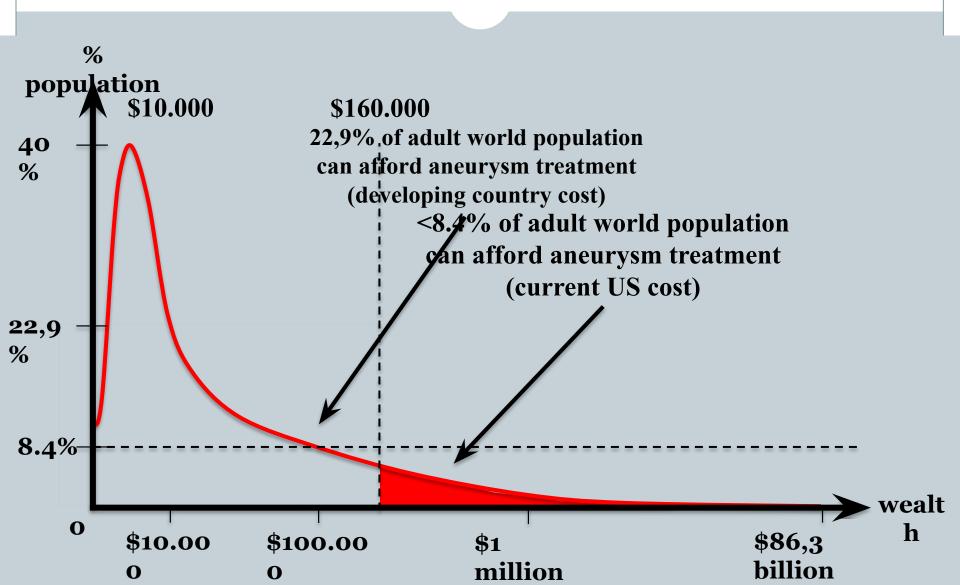
VS

•  $EU - 570,000 \times $36,000 = $20.5 \text{ billion}$ 

VS

India -  $570,000 \times $6,000 = $3.4 \text{ billion}$ 





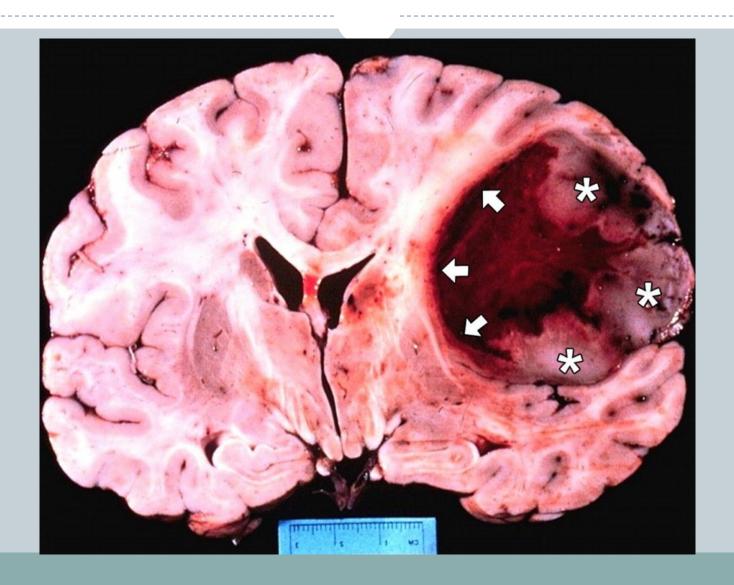
#### IV.f.

• What is the comparative benefit of world standards for each disease process?



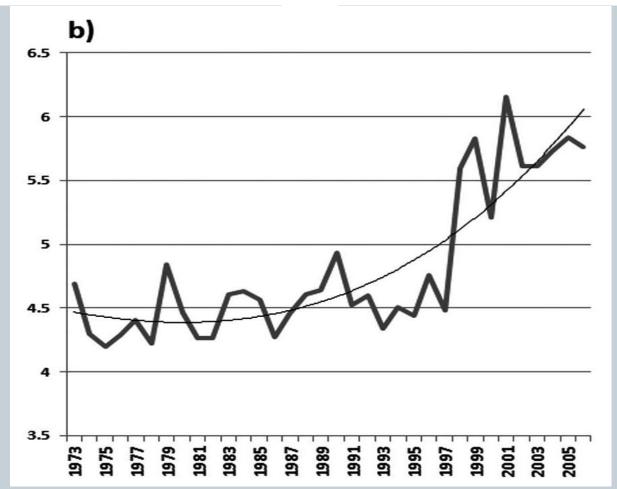


## i. Benefit of clinical guidelines - Glioblastomas



#### Benefit of clinical guidelines: Survivability in GBM

----- 95



Kaplan-Meier plot, 21,783 GBM patient yearly median survival development over time

#### Outcome (GBM)

Median survival without treatment is 3 months<sup>1</sup>

Median survival with maximal treatment is 14.6 months<sup>2</sup>

Treatment adds 11,6 months or about 1 year of life

1. Schapira, Anthony H.V. (2007). <u>Neurology and clinical neuroscience</u>.

Philadelphia: Mosby Elsevier. p. 1336. ISBN 9780323070539.

2. World Cancer Report 2014. World Health Organization.

2014. pp. Chapter 5.16. <u>ISBN</u> 9283204298.

### Outcome (GBM)

Median survival without treatment is 3 months

Median survival with maximal treatment is 14.6 months

Treatment adds 11,6 months or about 1 year of life

- ✓ Number of patients/year 214.500
- ✓ 214.500 years of life gained thanks to treatment
- Amount of money needed to treat these people in

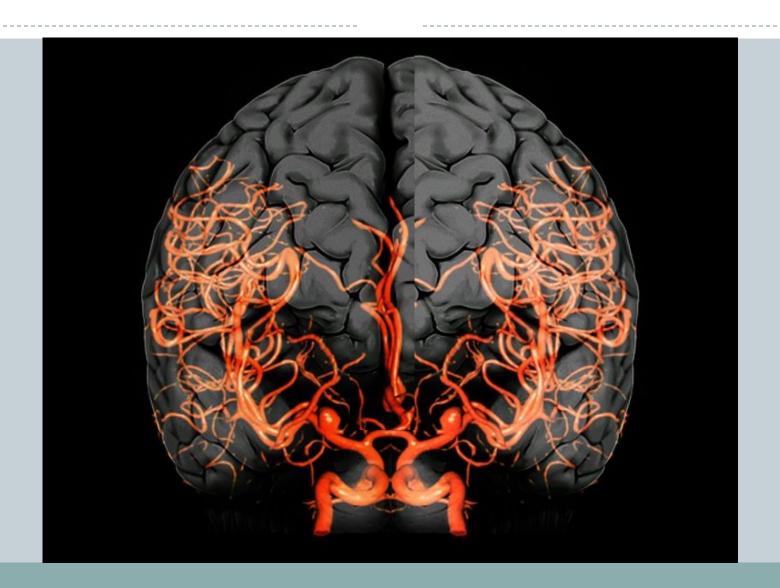
US - \$55 billion vs in India - \$6 billion

or

\$256.500 per year of life gained (US cost)

\$28.000 per year of life gained (India cost)

#### ii. Benefit of clinical guidelines – ruptured aneurysms



# Benefit of Guidelines Natural history of ruptured aneurysms w/wo treatment

Natural History of Ruptured, but Untreated Intracranial Aneurysms (Stroke, March 1, 2017)

Stroke. 2017 Mar 1. pii: STROKEAHA.116.015933. doi: 10.1161/STROKEAHA.116.015933. [Epub ahead of print]

#### Natural History of Ruptured but Untreated Intracranial Aneurysms.

Korja M1, Kivisaari R2, Rezai Jahromi B2, Lehto H2.

#### Author information

<sup>1</sup>From the Department of Neurosurgery, University of Helsinki and Helsinki University Hospital, Finland. miikka.korja@hus.fi.

#### BACK! cohort

#### The 1-year mortality rate was 75% for good-grade patients

relatively large hospital cohort.

**METHODS:** Patients admitted to the study hospital between 1968 and 2007 with saccular but untreated ruptured intracranial aneurysms were identified from the hospital aneurysm registry of 6850 patients. The study cohort included only patients who were followed up until death and for whom the date of symptom onset and the date of hospital admission were available.

RESULTS: For 510 patients identified, the median survival time from symptom onset to death was 20 days. The 1-year mortality rate was 65%, but varied substantially by admission delays and clinical status on admission, being lowest (13%) for patients admitted later than a month after symptom onset and highest (89%) for poor-grade patients. The 1-year mortality rate was 75% for good-grade patients admitted within a week.

**CONCLUSIONS:** Mortality rates for patients with untreated ruptured intracranial aneurysms are even worse than presented in the historical study. When discussing with subarachnoid hemorrhage patients and their relatives about treatment options, the presented natural history figures are of use.

© 2017 American Heart Association, Inc.

KEYWORDS: aneurysm, ruptured; cohort studies; intracranial aneurysm; natural history; subarachnoid hemorrhage

<sup>&</sup>lt;sup>2</sup>From the Department of Neurosurgery, University of Helsinki and Helsinki University Hospital, Finland.

## Benefit of Guidelines: Natural history of ruptured aneurysms w/wo treatment

Natural History of Ruptured and Treated Intracranial Aneurysms (ISAT)

#### **AHA/ASA Guideline**

#### **Guidelines for the Management of Aneurysmal Subarachnoid Hemorrhage**

A Guideline for Healthcare Professionals From the American Heart Association/American Stroke Association

Natural History and Outcome of aSAH

Although the case fatality of aSAH remains high world wide,5 mortality present in  $\approx 20\%$  of aSAH patients and is associated with in industrialized One study in the

per year from 1977 1994. Onlers have shown that case fatality rates decreased from 57% in the mid-1970s to 42% in the mid-1980s, 11 whereas rates from the mid-1980s to 2002 are reported to be anywhere from 26% to

36%.6,12,13,18,20,61,62 Mortality rates vary widely across published epidemiological studies, ranging from 8% to 67%.59 Regional variations become apparent when numbers from different studies are compared. The median mortality rate in epidemiological studies from the United States has been 32% versus 43% to 44% in Europe and

27% in Japan.<sup>59</sup> These numbers are based on studies that

mortality rate in epidemiological studies from the United States has been 32% versus 43% to 44% in Europe and

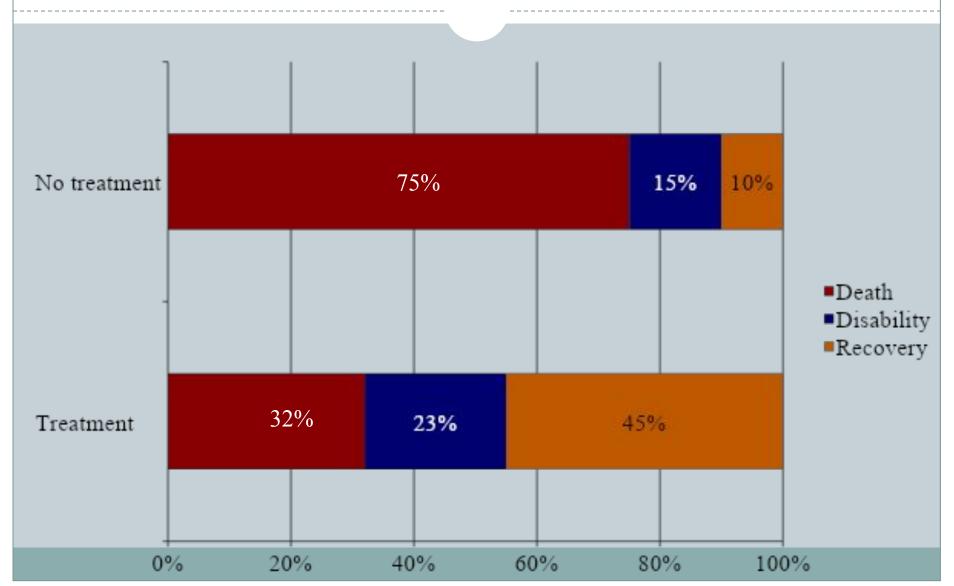
ified Rankin Scale 3) and 6.5% being functionally dependent (modified Rankin Scale score of 4-5) 1 year after aSAH. Furthermore, scales that are relatively insensitive to cognitive

poorer functional recovery and lower quality of life. 65 Cogdemonstrated that intellectual impairment is very prevalent

after aSAH. Although cognitive function tends to improve over the first year,64 global cognitive impairment is still present in ≈20% of aSAH patients and is associated with poorer functional recovery and lower quality of life.65 Cognitive deficits and functional decline are often compounded by mood disorders (anxiety, depression), fatigue, and sleep disturbances.66 Therefore, scales assessing well-being and quality of life can be particularly useful in the integral assessment of patients with aSAH, even among those who regain functional independence. 67,68 Behavioral and psychocial difficulties, as well as poor physical and mental durance, are some of the most commonly encountered

ctors accounting for the inability of otherwise independent tients to return to their previous occupations. 66,68

# Benefit of Guidelines: Natural history of ruptured aneurysms w/wo treatment



### Outcome (Ruptured aneurysms)

#### Primary impact

- 1. Decrease in mortality -43% (289.755/year)
- 2. Increase in disability -8% (53.907/year)
- 3. Increase in recovery -35% (235.847/year)
- ✓ Mean age at presentation 55.74
- ✓ Proper treatment can add about 20 years of productive life
- $\checkmark$  289.775 X 20 = 5.8 mln years of life gained thanks to treatment

or

\$4.862 per year of life gained (US cost)

\$690 per year of life gained (India cost)

## Conclusion N1

 The cost and supply shortfalls prevent universal applications of clinical guidelines across the world



## ✓ Conclusion N2

 Thus, treating ruptured aneurysms is about <u>50 times</u> more cost-effective than treating Glioblastomas



VS

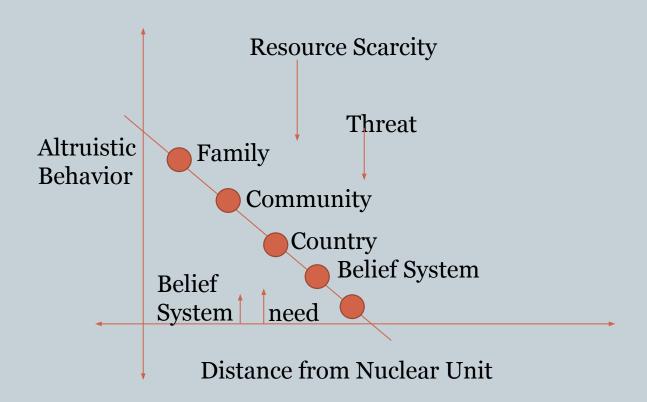


#### V. THE VIEW FROM MT. OLYMPUS

-How can worldwide healthcare resource dysfunction be improved? (diminish cost/ increase supply/ better resource reallocation/ combined efforts to improve clinical outcomes)



### V.a. Altruistic coefficient graph



## V.b. A COLLABORATIVE ARRANGEMENT BETWEEN THE DEVELOPED AND DEVELOPING WORLD



## V.c. The developed world's efforts



### V.c. The developed world's efforts

- i. Better resource allocation
- ii. Income transference
  - 1) Direct Cash transference
  - 2) Charity
  - 3) Universal Insurance
  - 4) Government mandated
- iii. Transference of Technology
- iv. Profit motive
  - v. Used Equipment

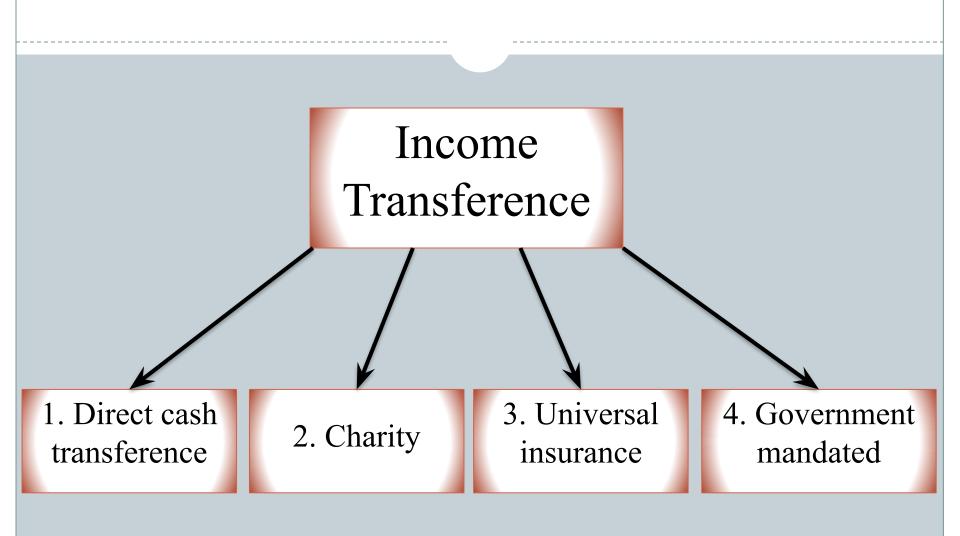
#### V.c. The developed world's efforts (continued)

- vi. Members of the International Patent System
- vii. MD Anderson Cancer Center
- viii. Establish Institutions in Developing Countries to Deal with Tertiary Neurosurgical Problems
  - ix. Advanced Medical Technology in Full Time Use
  - x. Technology around the world
  - xi. Telemedicine

# i. Better resource allocation with development of appropriate methodological system



#### ii. Income Transference



# 1) Direct Cash transference– Glioblastomas, ruptured aneurysms

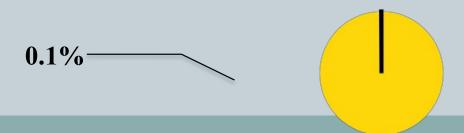
- US \$55 billion (0.6% of WHE)
- EU \$27.5 billion (0.3% of WHE)
- India \$6 billion (0.06% of WHE)

Glioblastomas

- US \$24 billion (0.25% of WHE)
- EU \$20.5 billion (0.2% of WHE)
- India \$3.4 billion (0.035% of WHE)

Ruptured aneurysms

- Combined amount: \$9.4 \$79 billion
- A relatively trivial amount considering the world's GNP of \$77.9 trillion and the amount the world spends on healthcare approximately \$9.5 trillion



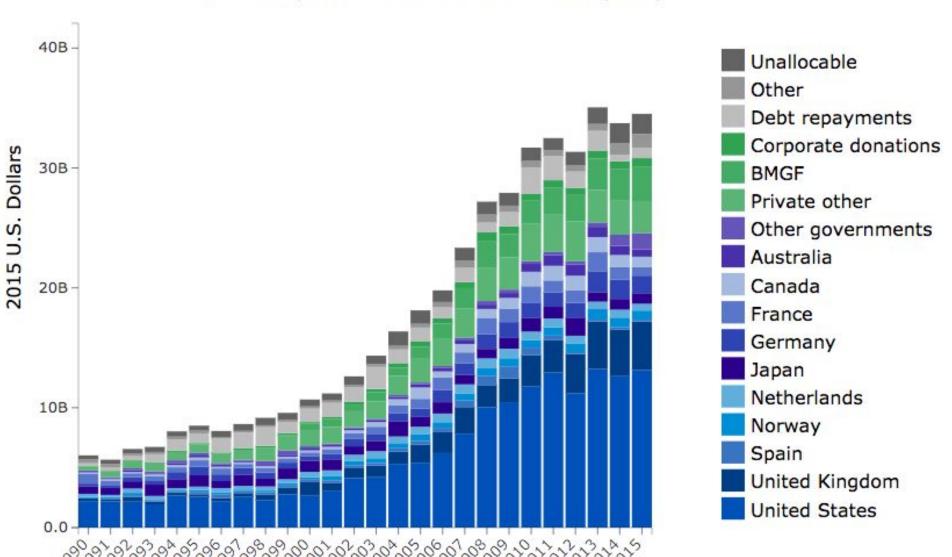
#### 2) Charity in the world

- Total amount of charity in the world
  - \$998.3 billion
  - o Gross amount of charity money on world healthcare \$40 billion

- US charity
  - \$29.8 billion out of \$373.2 billion is allocated to world health

## 2) Charity

Sources of Development Assistance for Health (DAH)

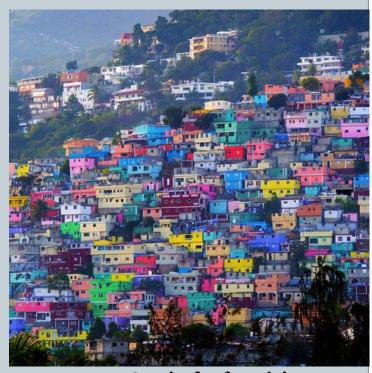


#### 3) Universal insurance

- Health expenditure per capita in US is \$9.000/year
- For a family of four will be \$36.000/year
- Thus, developing countries cannot afford it.

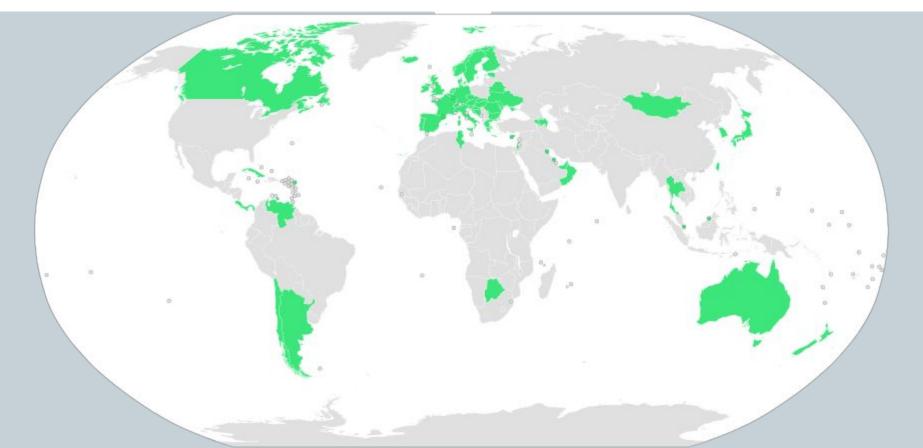
#### Haiti

- Population: 10.711.000<sup>(World bank)</sup>
- GDP: 8.765 billion<sup>(World bank)</sup>
- GNI per capita: \$810<sup>(World bank)</sup>
- Population older than 40: 2.5 million (23%) (Wikipedia)
- Calculated number of GBM per year: 250
- Treatment cost: \$62.5 million
- Even if we treat GBM 10 times cheaper,
   it will still be impossible in Haiti.



Capital of Haiti Port-au-Prince

### 4) Government mandated: Universal healthcare, but not across borders



58 countries with universal health care in 2009.

58 countries with legislation mandating universal health care, along with > 90% health insurance coverage, and > 90% skilled birth attendance.

## iii. Transference of Technology



## iii. Transference of Technology, cont.

- Mercer university in Vietnam:
   Manufacturing prosthetic limbs for landmines victims in Vietnam
- \$200 in Vietnam vs \$10,000 in US





## iv. Profit motive





# v. Used Equipment



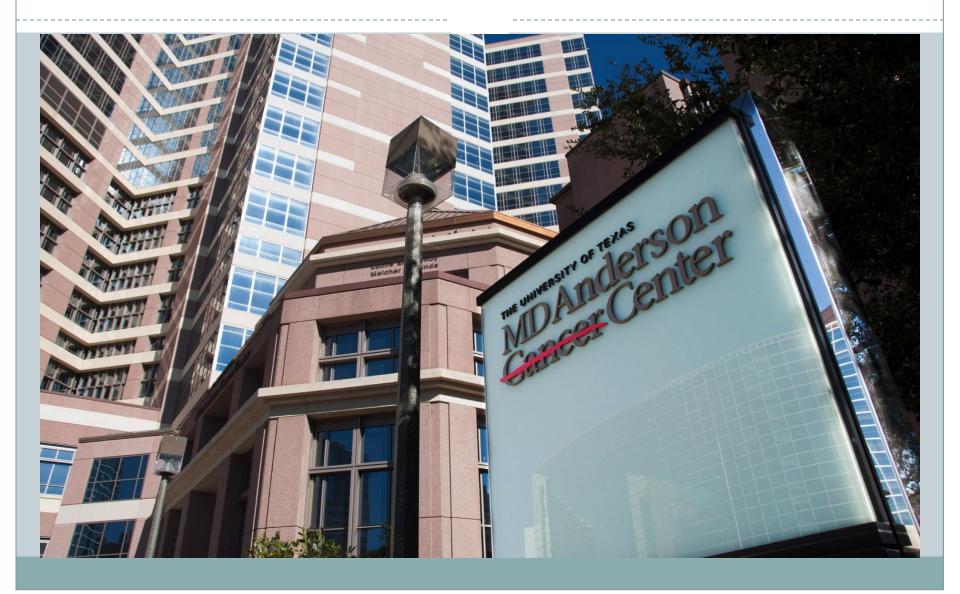
#### v. Used Medical Equipment

- Mexico imports medical equipment, instruments, disposable and dental products totaled US \$10.58 billion in 2015.
- This represented about 80 % of the medical equipment and instrument market and approximately 40 % of the disposable products and dental materials.
- Information on import regulations for pre-owned medical devices was available for 106 markets.
- Of 106 markets, 85 markets appear to permit the unrestricted importation of used or refurbished medical equipment on the same terms as new.
- In 2005 India lifted its restrictions, and Niger and Slovak Republic submitted reports in 2005 and 2007 respectively for the first time and indicated that there are no restrictions. Sixteen markets impose restrictions. Five generally prohibit the importation of pre-owned devices.

## vi. Members of the International Patent System



#### vii. MD Anderson Cancer Center



# viii. Establish Institutions in Developing Countries to Deal with Tertiary Neurosurgical Problems



# ix. Advanced Medical Technology in Full Time Use Expensive Technology around the Clock



#### x. Technology around the world

Top 20 countries for scientific output are first world countries with the USA publishes twice as much as the 2 countries after it in the list (Japan, Germany)

# The top 20 countries for scientific output

Top 20 countries by the number of scientific publications (1999-2009)

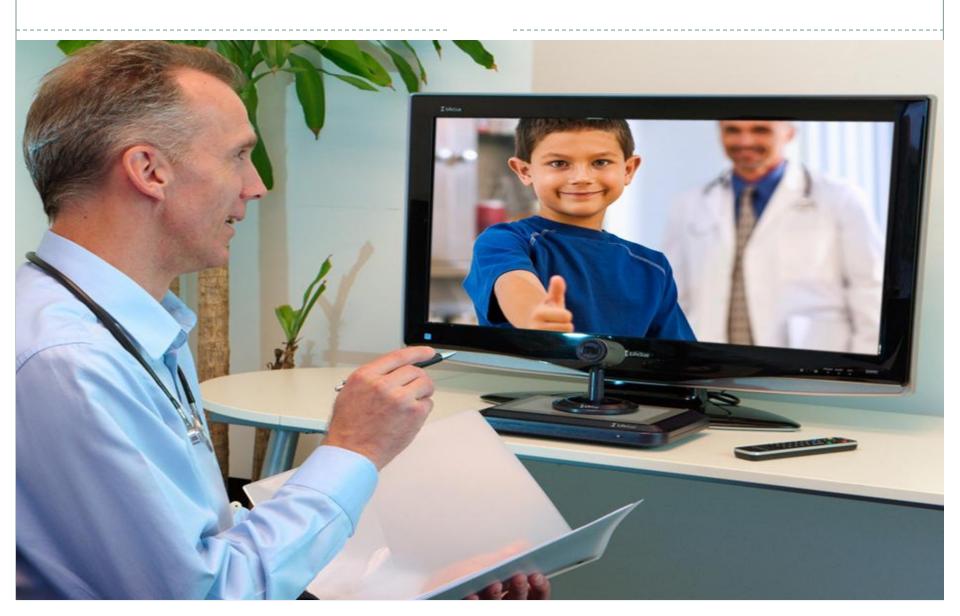


<sup>\*</sup> Numbers indicate the number of papers in millions (M)

 However, in regards to healthcare inquiries there needs to be higher grade clinical research studies with better demarcation of clinical expenditures.



## xi. Telemedicine



## V.d. What Can Developing Countries do?



#### V.d. What Can Developing Countries do?

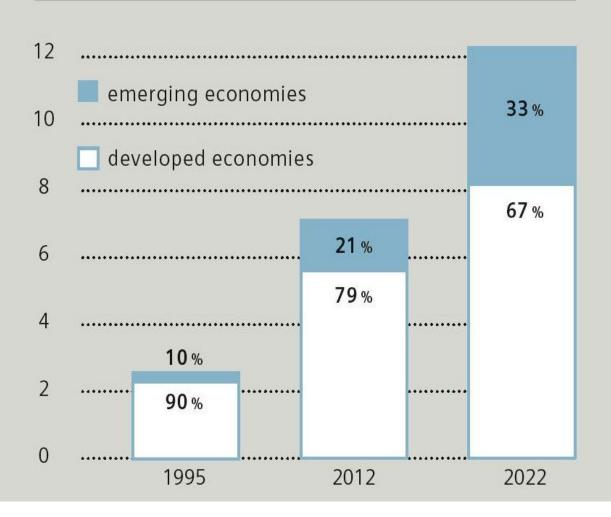
- i. Become more prosperous
- ii. Develop cheaper technology exportable to the 1<sup>st</sup> world
- iii. Become innovators
- iv. Discover outrider operative variables
  - 1) Discover unusual disease presentations
  - 2) Human genome variability
  - 3) Discover possible environmental causable agents
  - T cell & variability of the immune system
  - 5) Differing successful treatment modalities

#### V.d. What Can Developing Countries do? (continued)

- v. Participate in Clinical Trials
- vi. Western trained physicians from developing countries serving as a conduit for tertiary technology
- vii. Better statistics

#### i. Become more prosperous

#### Global health expenditure (in trillion US\$) 5)

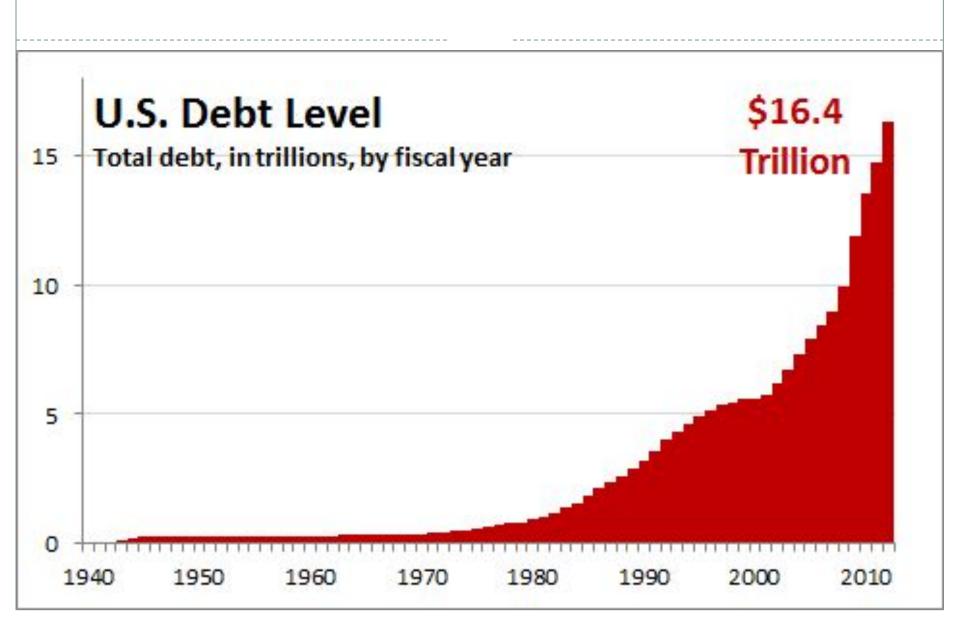


#### ii. Burr hole with ultrasound

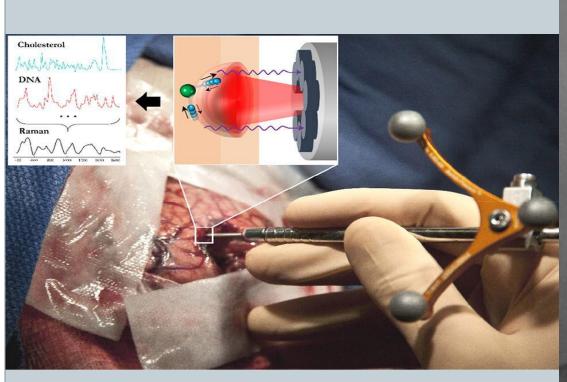
- ☐ They can experiment with less expensive options and see if they work, which could not be done in the US
- ☐ Ultrasound (\$200) vs CT-Scan (\$1200)



#### **US** National Debt



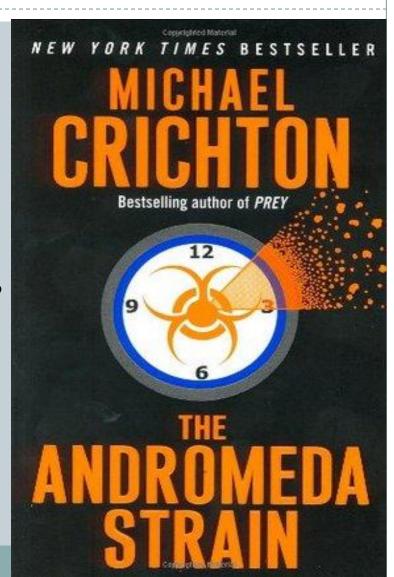
### iii. Become Innovators-Raman technique



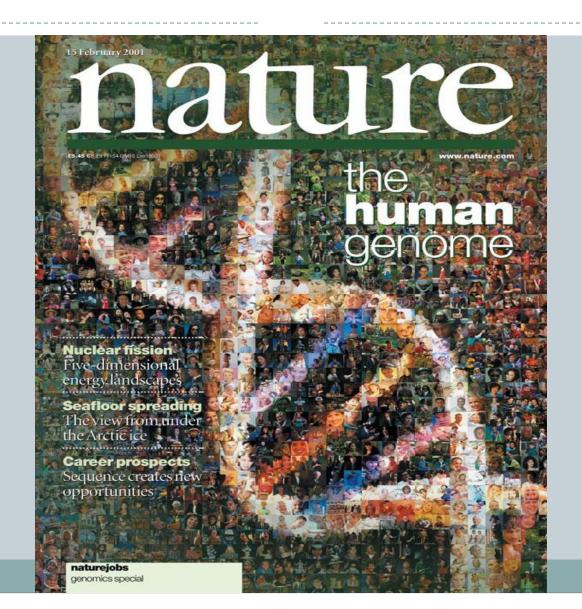


### iv.1. Discover Unusual disease Presentations And Treatment Patterns (find outrider operative variable)

- Maybe certain DNA protects against GBM/Glioma
- Maybe everybody has infections, can an infection lead to a GBM cure?



### iv.2. The human genome variability



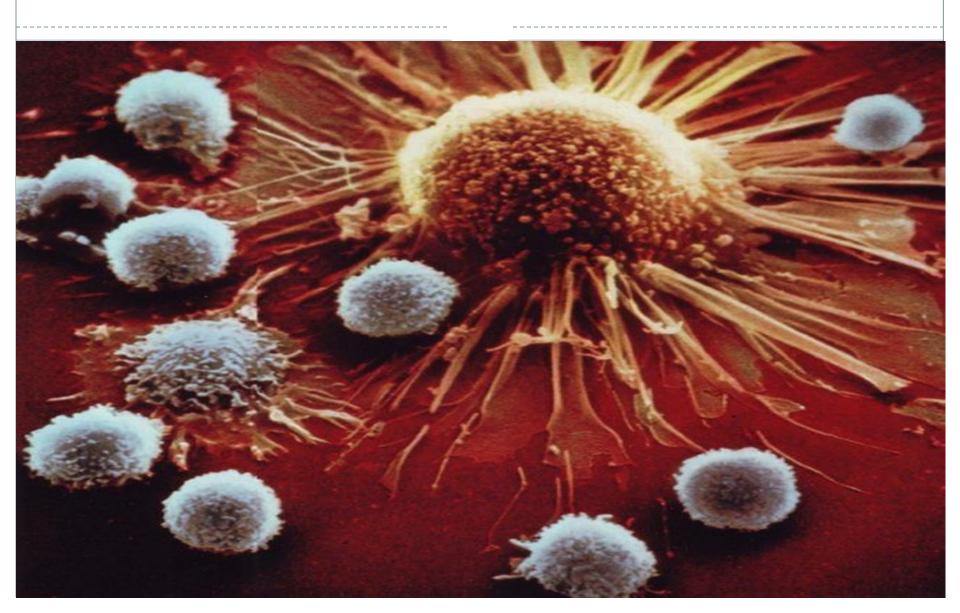
## iv.2. African tribe (Massai warriors)



### iv.3. Pollution and cancer



iv.4. T-cell & variability of the immune system

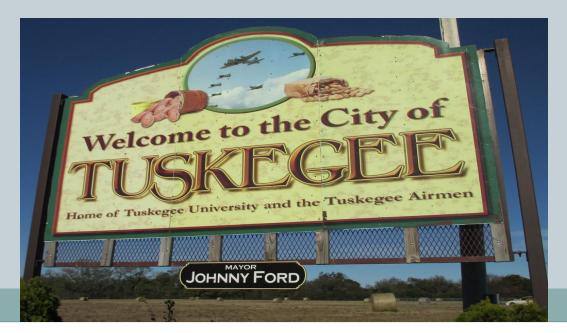


#### iv.5. 5 years survival rates in comparison to the USA

Glioblastoma	US	1995- 2010	4.7	4.4- 5.0
	Korea∞	1994- 2004	8.9	
	<b>⇒</b> US¹⁵⁴	1997- 2000	0.1	
ICD-O-3 morphology codes 9440 <b>–</b> 9442	UK & Ireland: (EUROCARE)	1995- 2002	2.2	1.6- 2.9
	Northern Europe:(EUROCARE)	1995- 2002	1.9	1.2- 2.9
	Central Europe: (EUROCARE)	1995- 2002	4.4	3.2- 5.9
	Eastern Europe: (EUROCARE)	1995- 2002	2.2	1.0- 4.4
	Southern Europe:(EUROCARE)	1995- 2002	2.8	1.8- 4.3

## v. Participate in Clinical Trails - Avoid the mistakes made in The Tuskegee Studies of untreated syphilis

■ The study was conducted between 1932 and 1972. The study initially involved 600 african american men — 399 with syphilis, 201 who did not have the disease. The study was conducted without the benefit of patients' informed consent.



### The Constant Gardner

 Unethical clinical trials in Africa for financial gain



# vi. Such physicians could pioneer technology and improve care arrangements for their home countries

 14% of the approximately 20,000 new residents and interns in the US are not US citizens



 Those physicians in western countries that come from developing countries could have the responsibility to their countries of origin to facilitate treatment of GBM

# vii. Better Statistics/Incidence is poorly established world wide...

#### Canada

o "Currently, the Canadian medical system does not track statistics on primary brain tumors. Complete and accurate data is needed to facilitate the research that will lead to a better understanding of this disease and improved diagnosis and treatment."

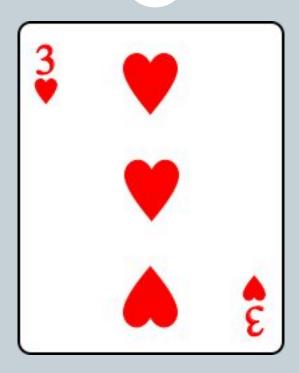
#### Korea

• Despite its importance, there are no accurate statistics on primary CNS tumors in Korea.

#### Sudan

- The reason there are few statistics on glioma incidence by country is because it is so expensive to diagnose
- We believe world statistics are greatly understated do to lack of available diagnostic medicine
- Furthermore, the world's population is younger than the typical age of diagnosis for Glioma/GBM, causing under-reported numbers.

### V.e. A third solution

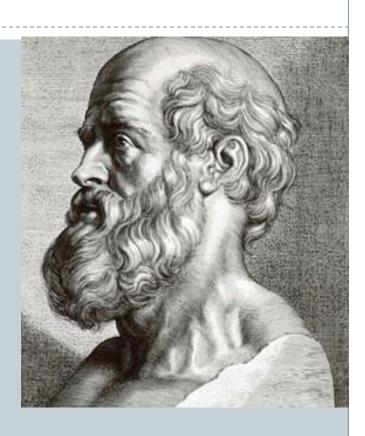


Physician Responsibility

## ii. Various Physician oaths

- The Oath of Hippocrates of Kos, 5th century BC
- Declaration of Geneva of the World Medical Association
- An Oath that "Bears the Name of Hippocrates"
- American Medical Association Code of Ethics (post-1980)





## iii. Responsibilities in the Hippocratic Oath

- I swear to fulfill, to the best of my ability and judgment, this covenant:...
- I will respect the hard-won scientific gains of those physicians in whose steps I walk, and gladly share such knowledge as is mine with those who are to follow.
- I will apply, for the benefit of the sick, all measures which are required, avoiding those twin traps of overtreatment and therapeutic nihilism.
- I will remember that there is art to medicine as well as science, and that warmth, sympathy, and understanding may outweigh the surgeon's knife or the chemist's drug.
- I will not be ashamed to say "I know not," nor will I fail to call in my colleagues when the skills of another are needed for a patient's recovery.
- I will respect the privacy of my patients, for their problems are not disclosed to me that the world may know. Most especially must I tread with care in matters of life and death. Above all, I must not play at God.
- I will remember that I do not treat a fever chart, a cancerous growth, but a sick human being, whose illness may affect the person's family and economic stability. My responsibility includes these related problems, if I am to care adequately for the sick.
- I will prevent disease whenever I can but I will always look for a path to a cure for all diseases.
- I will remember that I remain a member of society, with special obligations to all my fellow human beings, those sound of mind and body as well as the infirm.
- If I do not violate this oath, may I enjoy life and art, respected while I live and remembered with affection thereafter. May I always act so as to preserve the finest traditions of my calling and may I long experience the joy of healing those who seek my help.

## iv. Suggested additions to the oath

1. The responsibility of health care expenses sharing

2. Responsibility of spread of new medical technology



- "We are **Doctors Without Borders**/Médecins Sans Frontières (**MSF**). We help people worldwide where the need is greatest, delivering emergency **medical** aid to people affected by conflict, epidemics, disasters, or exclusion from health care"
- Founded: December 20, 1971, Paris, France
- Headquarters: Geneva, Switzerland
- Number of employees: 36,482







## THANK YOU

