Epidemiologia dell'ictus cerebrale

Tracking stroke epidemiology

Essential in order to *manage health services, to optimize economic resources, and to estimate the adequacy of the therapeutic strategies of primary and secondary prevention*

Moreover epidemiological studies allow *to extrapolate the expected number of patients* with cerebrovascular disease that can benefit of stroke units



Ministero della salute

DECRETO 2 aprile 2015, n. 70.

Regolamento recante definizione degli standard qualitativi, strutturali, tecnologici e quantitativi relativi all'assistenza ospedaliera. (15G00084). Pag.

relativi all'assistenza ospedaliera. (15G00084). . Pag. 1

DECRETI, DELIBERE E ORDINANZE MINISTERIALI

Ministero dell'economia e delle finanze

DECRETO 27 maggio 2015.

Individuazione dei criteri per la designazione, da parte dell'Associazione nazionale dei comuni italiani, dei componenti delle sezioni delle commissioni censuarie locali e centrale. (15A04315) Pag. 35

DECRETO 28 aprile 2015.

Annullamento del decreto 11 dicembre 2014, recante il «Rinnovo dell'autorizzazione al Laboratorio chimico merceologico azienda speciale della CCIAA di Napoli, al rilascio dei certificati di analisi nel settore vitivinicolo». (15A04135). Pag. 37



The Impact of Stroke

- 4th/3rd leading cause of death in most Western countries
- A leading cause of long-term disability
- 2nd most expensive illness
- Significant economic burden in terms of health care costs and lost productivity *Most strokes are preventable*

Age-adjusted death rates for CVD and CLRD



Stroke 2014;45:315-353

From the 1970s through 2008 the *age-adjusted stroke incidence rates* in high-income countries declined 42% overall with declines in each subsequent decade of the study

These trends were found across all age groups, with a greater decline in those \geq 75 years of age

A greater than 100% increase in stroke incidence was found in low to middle income countries

Lancet Neurol 2009;8:355-369

Incidence rates of stroke types in high-income countries



The pooled overall age-adjusted incidence rates of ischemic stroke fell by 11%, likely reflecting advances in primary prevention

Lancet Neurol 2009;8:355-369

Early case-fatality rates in high-income countries



During the last decades there was a *nonsignificant reduction* in early stroke case-fatality

The yearly averaged % change in stroke casefatality was -1.1% likely reflecting advances in stroke care and secondary prevention

Lancet Neurol 2009;8:355-369

Decline in stroke mortality rates



The decline in mortality results from *reduced* incidence of stroke and lower casefatality rates and is concurrent with cardiovascular risk factor control interventions

Stroke 2014;45:315-353

US age-standardized death rates attributable to cardiovascular disease 2000 to 2013



Circulation 2016; DOI: 10.1161/CIR.000000000000350

Changes in stroke death rates from 1981 to 2013

Slightly greater decline in age-adjusted stroke death rates in men (-61.4%) than in women (-58.9%) aged ≥18 years

Stroke death rates declined more among people aged ≥65 years (-54.1%) than those aged 45 to 64 years (-53.6%) or those aged 18 to 44 years (-45.9%)

Age-adjusted stroke death rates for adults aged ≥18 years declined by ≈50% or more among all racial groups

Circulation 2016; DOI: 10.1161/CIR.000000000000350

Trends in carotid endarterectomy and stenting procedures



Circulation 2016; DOI: 10.1161/CIR.000000000000350

Epidemiology of Stroke in Europe

Large differences in incidence, prevalence and mortality have been reported between Eastern and Western Europe

This has been attributed to differences in risk factors distribution, with higher levels of hypertension and other risk factors resulting in more severe strokes in Eastern Europe

Eur J Neurol 2015;22:284-291

The EuroHOPE register study



Patients with an index stroke on admission in 2007 were followed for 1 year The number of patients registered per 100,000 European standard population ranged from 77 in Scotland to 407 in Hungary Large differences were observed in medication use

Eur J Neurol 2015;22:284-291

Proportional frequency of stroke types



Lancet 2005;365:2160-2161

Epidemiology of stroke in Italy

Table 1 Main characteristics of the population-based registers included in the analysis of stroke incidence and case-fatality

		Duration (years)	Population	Total strokes Brain neuro- (n; % men) imaging (%)	Hospita-	Mean age at onset (years \pm SD)			
Study	Study period		at risk (person-years)		Brain neuro- imaging (%)	lization (%)	Men	Women	Total
Acquaviva-Casamas- sima (16)	2001–2002	2	38735	127; 61	94	95	72.5 (NR)	77-5 (NR)	74.5 (NR)
Aeolian Islands (14,20)	1999–2002	3	13 4 3 1	62; 42	82	82	71.5 (<u>+</u> 10.3)	74·6 (±9·6)	72.5 (±9.92)
Aosta (11,12)	1997	1	1 18 72 3	343; 47	97	95	69.3 (NR)	77.2 (NR)	73.3 (NR)
Belluno (15)	1992-1993	1	2 11 389	474; 43	90*	93	NR	NR	NR
L'Aquila (10,17–19, 25–28)	1994–1998	5	2 97 645	4353; 47	88	94	72·6 (±11·9)	76∙7 (±10∙6)	74·8 (±11·4)
Trasimeno (21–24)	1986-1989	3	49218	375; 49	69*	85	NR	NR	NR
Vibo Valentia (16)	1996	1	1 79 186	321; 51	96*	98	71∙9 (<u>+</u> 10•6)	74·2 (<u>+</u> 11·7)	73.0 (<u>+</u> 11.2)

*Only brain computed tomography.NR, not reported.

67-83% IS - 10-20% ICH - 2-4.0% SAH - 1-18% UNDET

Int J Stroke 2011;6:219–227

Epidemiology of stroke in Italy



	IRR	95% CI	Р
Area (reference l	Northern Italy)		
Central	0.96	0.89-1.03	0.23
Southern	0.72	0.64-0.80	<0.001
Age group (refer	ence 0–54 years)		
55-64	10.5	9.3-11.9	<0.001
65–74	32.2	28.8-35.9	<0.001
75–84	63.6	57.0-70.9	<0.001
85+	109.8	97.6-123.4	<0.001
Gender (referend	ce male gender)		
Female	0.78	0.74-0.82	<0.001

The incidence rates increased linearly with age, peaking in subjects aged 85+ years in all the registers

Age group, gender, and geographic area were significant predictors of stroke incidence

Int J Stroke 2011;6:219-227

Comparison of the Age Structure of the Study Population with that of Italy (ISTAT, 2011) and Europe (EUROSTAT, 2011)



Comparison of the Age Structure of the Study Population between 1994-1998 and 2011



Age-structure Changes in Residents ISTAT L'Aquila 1981-1991-2001-2011



High Stroke Incidence in the Prospective Community-Based L'Aquila Registry (1994–1998)

First Year's Results

Antonio Carolei, MD; Carmine Marini, MD; Mario Di Napoli, MD; Giacinto Di Gianfilippo, MD; Paola Santalucia, MD; Massimo Baldassarre, MD; Giorgio De Matteis, MD; Ferdinando di Orio, MD

Stroke 1997;28:2500-2506



Study period



Demographic characteristics

1994-1998

2011-2012

Gender	n	Age (years) mean \pm SD min max
Men	2051	72.7±11.8 5 100
Women	2302	76.7±10.6 16 106
Both	4353	74.8±11.4 5 106

Gender	n	Age (years) mean \pm SD min max
Men	417	72.7±13.2 7 96
Women	441	78.2±12.6 19 103
Both	858	75.5±13.2 7 103

Distribution of Cerebrovascular Events (ICD - IX Revision)



430: subarachnoid hemorrhage; 431-432: intracerebral hemorrhage; 433-434: cerebral ischemia; 436-437: ill-defined cerebrovascular disease

First-ever Stroke

Age distribution







First-ever Stroke Incidence

1994-1998

2011-2012



The decreased stroke incidence over the last two decades suggested that the improvement of prevention strategies might have contributed not only to postpone the age of stroke onset but also to reduce stroke incidence and mortality

Incidence of Cerebrovascular Events (ICD - IX Revision)



30-day Case-fatality rates



1-year Case-fatality rates





	1994-1998	2011-2012
Arterial hypertension	65.3	72.4 懀
Diabetes mellitus	25.5	22.4
Coronary heart disease	24.6	14.3 🦊
Hypercholesterolemia	24.4	18.5 🖊
Smoking habit	23.5	13.3 🦊
Atrial fibrillation	21.7	27.5 懀
Peripheral arterial disease	11.7	5.1 👃
Previous TIA	7.4	5.4

Demographics of Ischemic Stroke

1994-1998

2011-2012

Gender	n	Age (y mean±SD	ears) min	max	Gender	n	Age (ye mean \pm SD	ears) min	
Men	1700	73.4±11.0	5	100	Men	316	72.5±12.9	7	
Women	1894	77.0±10.1	16	100	Women	318	78.6±12.0	19	
Both	3594	75.3±10.7	5	100	Both	634	75.6±12.8	7	

Ischemic Stroke Incidence



Ischemic Stroke Incidence Trend

	Crude rate x1000 (95% CI)		Incidence rate ratio	(95% CI)	P Value
	1994-1998	2011-2012			
All	2.42 (2.39-2.44)	1.06 (0.98-1.15)	0.44	0.40-0.48	<0.001
Men	2.36 (2.32-2.39)	1.09 (0.97-1.21)	0.46	0.41-0.52	<0.001
Women	2.47 (2.44-2.50)	1.04 (0.93-1.16)	0.42	0.37-0.47	<0.001
Age <55 years	0.16 (0.13-0.18)	0.13 (0.09-0.17)	0.82	0.59-1.12	0.216
Age 55-74 years	4.37 (4.15-4.60)	1.39 (1.20-1.60)	0.32	0.27-0.37	<0.001
Age ≥75 years	14.68 (14.04-15.34)	5.51 (4.97-6.08)	0.38	0.34-0.42	<0.001

Risk factors for Ischemic Stroke

	1994-1998	2011-2012
Arterial hypertension	64.2	77.0 懀
Diabetes mellitus	25.7	24.3
Coronary heart disease	26.2	15.9 🦊
Hypercholesterolemia	26.1	20.3
Smoking habit	24.1	13.9 🦊
Atrial fibrillation	24.2	31.5 🕇
Peripheral arterial disease	12.2	6.2
Previous TIA	8.3	5.7

Risk Factors for Ischemic Stroke

1994-1998 vs 2011-2012

Ω

<44







45-54 55-64 65-74 75-84

Age (years)

85+

*P<0,001

Demographics of ICH

1994-1998

2011-2012

Gender	n	Age (y	ears)	
		mean±5D		max
Men	247	70.5±13.1	24	95
Women	302	76.1±11.5	19	106
Both	549	73.6±12.5	19	106

Gender	n	Age (ye	ears)	
		$mean \pm SD$	min	max
Men	77	73.9±13.0	27	92
Women	71	77.8±13.1	25	98
Both	148	75.8±13.2	25	98

ICH Incidence



ICH Incidence Trend

	Crude rate x1000 (95% CI)		Incidence rate ratio	95% CI	P Value
	1994/1998	2011/2012			
All	0.37 (0.36-0.38)	0.25 (0.21-0.29)	0.67	0.56-0.80	<0.001
Men	0.34 (0.31-0.38)	0.27 (0.21-0.33)	0.77	0.60-0.99	0.049
Women	0.39 (0.35-0.44)	0.23 (0.18-0.29)	0.59	0.45-0.76	<0.001
Age <55 years	0.04 (0.03-0.05)	0.03 (0.02-0.05)	0.72	0.36-1.32	0.319
Age 55-74 years	0.69 (0.60-0.79)	0.29 (0.20-0.39)	0.41	0.29-0.57	<0.001
Age ≥75 years	2.04 (1.80-2.29)	1.35 (1.10-1.65)	0.66	0.52-0.83	0.0005

Risk factors for ICH

	1994-1998	2011-2012
Arterial hypertension	75.6	68.9
Diabetes mellitus	26.2	21.6
Coronary heart disease	17.9	10.8 🖊
Hypercholesterolemia	17.5	15.5
Smoking habit	20.6	11.5 👃
Atrial fibrillation	10.7	20.3* 🕇
Peripheral arterial disease	9.8	3.4 👃
Previous TIA	3.6	5.4

*including patients with AF and medication-related ICH

Demographics of SAH

1994-1998

2011-2012

Gender	n	Age (y mean±SD	ears) min max	
Men	55	56.2±16.8	15 92	
Women	63	64.7±14.1	28 101	
Both	118	60.7±15.9	15 101	

		Age (ve	ears)	
Gender	n	mean ± SD	min	max
Men	12	61.9±17.6	36	91
Women	27	65.4±13.1	42	89
Both	39	64.4±14.5	36	91





Atrial fibrillation profile

1994-1998

2011-2012

- 4353 First-ever strokes 946 (22%) with AF
- 3594 Ischemic strokes 869 (24%) with AF
- 1316 Ischemic strokes over 80 397 (30%) with AF

- 858 First-ever strokes 235 (27%) with AF
- 634 Ischemic strokes 199 (31%) with AF
- 289 Ischemic strokes over 80 126 (44%) with AF

Incidence of ischemic stroke subtypes in Europe

	Ischemic stroke large vessels	Ischemic stroke small vessels	Ischemic stroke cardioembolic	Ischemic stroke miscellaneous	Unknown ischemic stroke	Total
Total	15.3 (95% CI: 12–19.3)	25.8 (95% CI: 21.5–30.9)	30.2 (95% CI: 25.6–35.7)	2.1 (95% CI: 0.9–3.9)	39.3 (95% CI: 34–45.5)	112.7 (95% CI: 103.6–122.6)
Men	23.6 (95% CI: 17.1–31.6)	35.3 (95% CI: 27.2–45.2)	29.1 (95% CI: 21.7–38.2)	1.4 (95% CI: 0.3–4.1)	40.2 (95% CI: 31.3–50.6)	129.7 (95% CI: 114.3–147)
Women	9.2 (95% CI: 6.1–13.4)	19.8 (95% CI: 15–25.8)	30.8 (95 CI: 24.8–37.8)	2.5 (95% CI: 0.9–5.5)	39.1 (95% CI: 32.5–47)	101.4 (95% CI: 90.5–113.7)

J Neurol Sci 2007;262:85-88

Secular trends in ischemic stroke subtypes



LA: large artery CA: cardioembolic SA: small artery OE: other etiology UE: undetermined

Stroke 2014;45:3208-3213

L'Aquila Stroke Registry 2011-2012



L'Aquila Stroke Registry 2011-2012

CE

Main characteristics of the included

cording to TOAST criteria

	LAA	TOAST CE	220 (34.7)	OC	UND	Р
Main characteristics			/0//121210			
n.% Age ±DS	73 (11.5) 76.33±10.9	220 (34.7) 78.91±12.3		9 (4.6) 97±19.0	220 (34.7) 76.14±11.4	<0.001
Gender Men Women	42 (57.5) 31 (42.2)	91 (41.3) 129 (58.6)	91 (41.3) 129 (58.6)	3 (62.0) L (37.9)	108 (49.0) 112 (50.9)	0.004
Stroke severity and disability NIHSS at onset (median, IQR) mRS at discharge (median, IQR)	6 (3-12) 3 (2-5)	10 (4-17,5) 4 (3-5)	10 (4-17,5)	(2-11) 5 (1-5)	6 (3-12) 3 (2-4)	<0.001 <0.001
Case fatality rates 30-day mortality 1-year mortality	13.7 20.5	26.8 38.2	4 (3-5)	17.2 24.1	16.3 24.1	<0.001 <0.001

L'Aquila Stroke Registry 2011-2012

Risk factors distribution according to TOAST criteria

	Total	(634)	LAA	(73)	CE (2	220)	SAC) (92)	00	(29)	UND	(220)	Р
Risk factor	N	%	Ν	%	Ν	%	Ν	%	Ν	%	Ν	%	
Hypertension	488	77.0	56	76.7	180	81.8	79	85.9	13	44.8	160	72.7	<0.001
Diabetes	154	24.3	23	31 . 5	55	25.0	27	29.3	8	27.6	41	18.6	0.154
Dyslipidemia	130	20.5	16	21.9	41	18.6	29	31.5	4	13.8	40	18.2	0.052
Atrial fibrillation	200	31.5	2	2.7	173	78.6	11	12.0	1	3.4	13	5.9	<0.001
Coronary heart disease	101	15.9	10	13.7	47	21.4	18	19.6	4	13.8	22	10.0	0.029
Peripheral artery disease	39	6.2	6	8.2	15	6.8	6	6.5	1	3.4	11	5.0	0.863
Cigarette smoking	88	13.9	16	21.9	19	8.6	16	17.4	9	31.0	28	12.7	0.074
Alcohol abuse	80	12.6	14	19.2	25	11.4	11	12.0	2	6.9	28	12.7	0.471

Crude incidence of ischemic stroke subtypes by age groups 2011-2012



Definitions of TIA

Time-based a sudden, focal neurological deficit of presumed vascular origin lasting less than 24 hours

Tissue-based a transient episode of neurological dysfunction caused by focal brain, spinal cord, or retinal ischemia, without acute infarction

Acute neurovascular syndrome (ANS) relatively brief symptom duration (symptoms may persist several hours but less than a day) who do not receive a detailed diagnostic evaluation

Cerebral infarction with transient symptoms events that last <24 hours but are associated with cerebral infarction

Stroke 2009;40:2276–2293

TIA Incidence - Tissue-based Crude rate 25/100,000



TIA Incidence - All definitions Crude rate 35/100,000



TIA Incidence in the District of L'Aquila

TIA definition	Total number of included TIA	Crude Incidence (x100,000)	95% CI	Standardized* incidence	95% CI
Tissue-based	151	25.3	21.5-29.6	20.0	16.3-24.1
All definitions	210	35.2	30.7-40.2	28.6	24.1-33.5

CI confidence interval *to the 2011 European population

THANKS!

L'Aquila Stroke registry Distribution of lacunar and nonlacunar strokes



Nonlacunar stroke Lacunar stroke

Demographics of lacunar stroke

1994-1998

2011-2012

Gender	n	Age (y mean±SD	ears) min	max
Men	252	70.6±10.2	35	97
Women	239	74.9±09.4	36	91
Both	491	72.7±10.1	35	97

Gender	n	Age (ye mean±SD	ears) min	max
Men	57	67.0±10.6	45	90
Women	35	74.5±12.1	46	89
Both	92	69.8±11.7	45	90

Risk Factors for Lacunar and Nonlacunar Stroke Multivariate analysis



Lacunar Stroke Incidence



Lacunar Stroke Incidence

Men

Women

IRR=0.56 (95% CI 0.42-0.74); p<0.001



IRR=0.37 (95% CI 0.25-0.52); p<0.001



1994-1998 • 2011-2012

Lacunar vs nonlacunar stroke 1-year survival



Lacunar

Nonla<u>cunar</u>

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CI confidence interval *to the 2011 European population