# Secondary Data Analysis

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### **Disclosures**

NIH (R01HL144541, U01NS095869, U01NS106513)

BMS (in-kind study drug for ARCADIA trial)

Roche (ancillary study support for ARCADIA trial)

Medtronic (clinical trial steering committee)

Janssen (clinical trial executive committee)

Javelin Medical (clinical trial steering committee)

Boehringer-Ingelheim (clinical trial adjudication committee)

NovoNordisk (clinical trial adjudication committee)

Deputy Editor, JAMA Neurology



## What is secondary data analysis?

Use of existing data for an analysis that is distinct from the primary reason the data were collected



## Why do secondary data analysis?

Feasibility

Efficiency

**Training** 



## Why do secondary data analysis?

Trainees usually lack time, resources, and research skills 2° analysis lets you do in days what would take years Allows skill acquisition and apprenticeship







#### Methods

Design

We retrospectively studied the rate, timing, and predictors of delayed detection of AF in a cohort of patients with ischemic stroke. To take advantage of the close monitoring and follow-up required in randomized clinical trials, we examined data from patients in the placebo arms of 4 trials: Clomethiazole Acute Stroke Study in Ischemic Stroke (CLASS-I), NXY-059 for Acute Ischemic Stroke (SAINT-I), NXY-059 for the Treatment of Acute Ischemic Stroke (SAINT-II), and Effects of Repinotan in Patients with Acute Ischemic Stroke (mRECT). Data from these trials were obtained from the Virtual International Stroke Trials Archive. Details of CLASS-I, SAINT-I, SAINT-II, and Virtual International Stroke Trials Archive have been published elsewhere. 18-21 The design and results of mRECT were presented in abstract form at the XIV European Stroke Conference.<sup>22</sup> Briefly, all 4 studies were randomized, double-blinded, placebo-controlled trials of neuroprotective agents in acute ischemic stroke. All trials were approved by the institutional review boards at the participating institutions. Our analysis was certified as exempt from review by our institutional review board because the data had been collected for other purposes and lacked patient identifying information.

#### Delayed Detection of Atrial Fibrillation after Ischemic Stroke

Hooman Kamel, MD,\* Kennedy R. Lees, MD,† Patrick D. Lyden, MD,‡§
Philip A. Teal, MD,// Ashfaq Shuaib, MD,¶ Myzoon Ali, MRes,† and
S. Claiborne Johnston, MD, PhD,\* on behalf of the Virtual International Stroke Trials
Archive Investigators





#### Methods

Design

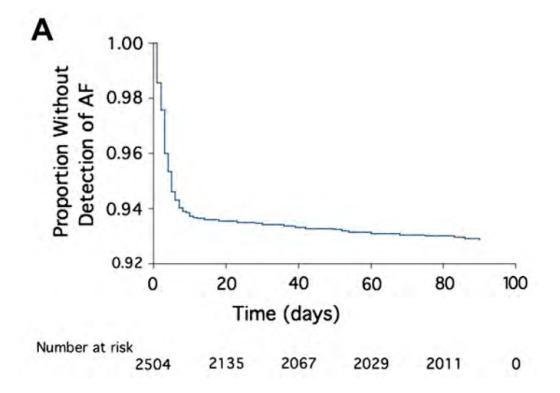
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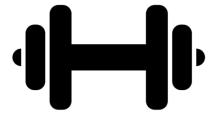
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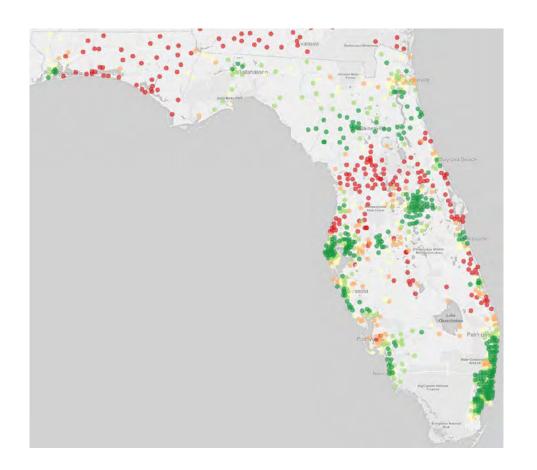
## Why do secondary data analysis?

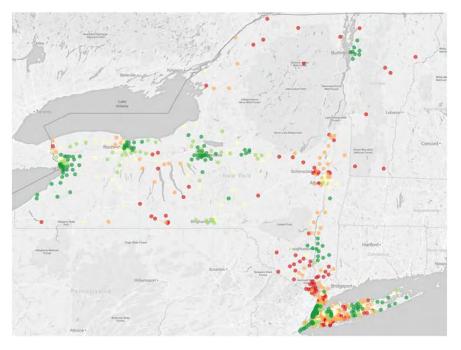
Some projects are not feasible without 2° data analysis











### Stroke

### ORIGINAL CONTRIBUTION

Access to Mechanical Thrombectomy for Ischemic Stroke in the United States

Hooman Karnel<sup>10</sup>, MD, MS; Neal S, Parikh<sup>10</sup>, MD, MS; Abhinaba Chatterjee, BS; Luke K, Kim<sup>10</sup>, MD; Jeffrey L, Saver<sup>10</sup>, MD; Lee H, Schwamm<sup>10</sup>, MD; Kori S, Zachrison<sup>10</sup>, MD, MSc; Raul G, Nogueira<sup>10</sup>, MD; Opeolu Adeoye<sup>10</sup>, MD; Iván Díaz<sup>10</sup>, PhD; Andrew M, Ryan, PhD; Ankur Pandya<sup>10</sup>, PhD; Babak B, Navi<sup>10</sup>, MD, MS



Our study should be considered in light of its limitations. First, we lacked data from the entire United States, particularly from Western states. However, we had comprehensive data from 11 heterogeneous, geographically dispersed states encompassing 80 million residents. Second, we analyzed administrative data rather than review medical records or prospectively ascertain events. However, a comprehensive, population-based analysis across the United States is not possible without administrative data. To mitigate misclassification, we used validated codes to define key variables. Third, we lacked data on the availability of certain important clinical services such as telestroke consultation, and we lacked data on the numbers of interventionalists and the times during the week when interventional treatments were available at each hospital. Fourth, our latest data were from the end of 2018 and thrombectomy access may have improved in the last 2 years. The all-payer claims data we used are released with a lag time of several years, and we used the latest available data, so periodic analyses will be required to assess trends in thrombectomy access.

### Stroke

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**¬** New York-Presbyterian

## How to do secondary data analysis?

Randomized clinical trials

Observational research studies

Registries

Surveys

Administrative data



### Randomized clinical trials

### Advantages

- Relatively large populations with a specific disease
- Rigorous ascertainment of outcomes
- Can explore effects of intervention

### Disadvantages

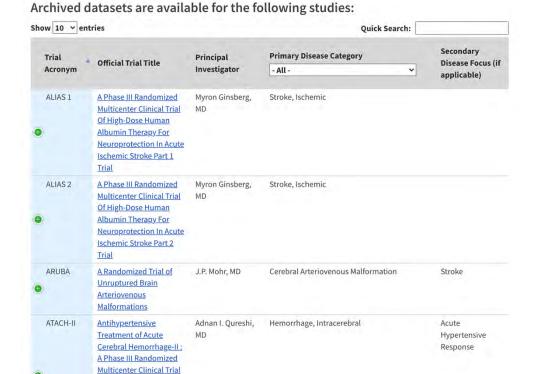
- Not population based
- Need to account for intervention effect
- Usually modest sample sizes



### Randomized clinical trials

### NINDS repository

https://www.ninds.nih.gov/Current-Research/Research-Funded-NINDS/Clinical-Research/Archived-Clinical-Research-Datasets



of Blood Pressure





#### Methods

### **Study Design**

We performed an individual patient-level analysis of patients with ICH enrolled in the Minimally Invasive Surgery Plus Alteplase for Intracerebral Hemorrhage Evacuation phase 3 trial (MISTIE III), <sup>10</sup> Antihypertensive Treatment of Acute Cerebral Hemorrhage (ATACH-2) trial, <sup>11</sup> Intracerebral Hemorrhage Deferoxamine (i-DEF) phase 2 trial, <sup>12</sup> and the multicenter prospective ERICH study. <sup>13</sup> The study and trial protocols were approved by an ethics committee at each enrolling site, and written informed consent was obtained from each participant or their legal surrogate. This study was approved by the institutional review board of Weill Cornell Medicine, New York, New York, and followed the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) reporting guideline (eAppendix in the Supplement).

### JAMA Neurology | Original Investigation

## A Pooled Analysis of Diffusion-Weighted Imaging Lesions in Patients With Acute Intracerebral Hemorrhage

Santosh B. Murthy, MD, MPH; Sung-Min Cho, DO; Ajay Gupta, MD, MS; Ashkan Shoamanesh, MD; Babak B. Navi, MD, MS; Radhika Avadhani, MS; Joshua Gruber, MSPH; Yunke Li, MD; Tatiana Greige, MD; Vasileios-Arsenios Lioutas, MD; Casey Norton, BS; Cenai Zhang, MS; Pitchaiah Mandava, MD, PhD, MSEE; Costantino Iadecola, MD; Guido J. Falcone, MD, ScD, MPH; Kevin N. Sheth, MD; Alessandro Biffi, MD; Jonathan Rosand, MD; Adnan I. Qureshi, MD; Joshua N. Goldstein, MD; Chelsea Kidwell, MD; Issam Awad, MD; Magdy Selim, MD; Daniel F. Hanley, MD; Daniel Woo, MD; Hooman Kamel, MD; Wendy C. Ziai, MD, MPH





Table 3. Multivariable Logistic Regression of Factors Associated With DWI Lesions<sup>a</sup>

Characteristic	OR (95% CI)	P value	
Age, per y	0.98 (0.97-0.99)	.004	
Male sex	1.33 (1.01-1.74)	.04	
Race/ethnicity			
White	1 [Reference]	NA	
Black	1.64 (1.17-2.30)	.004	
Hispanic	0.89 (0.62-1.28)	.54	
Other	0.64 (0.22-1.13)	.42	
Prior anticoagulant therapy	0.63 (0.35-1.13)	.12	
Hematoma volume, baseline (per 10-mL increase)	1.12 (1.02-1.22)	.01	
Presence of IVH	1.07 (0.79-1.43)	.66	
Hematoma location			
Lobar	1 [Reference]	NA	
Deep	0.81 (0.58-1.11)	.19	
Infratentorial	1.09 (0.65-1.82)	.73	
Baseline SBP (per 10-mm Hg increase)	1.13 (1.08-1.18)	<.001	
Delta SBP	1.00 (0.98-1.04)	.49	
Presence of cerebral microbleeds	1.85 (1.39 -2.46)	<.001	
Leukoaraiosis, moderate to severe	1.59 (1.67-2.17)	.003	

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#### Methods

### **Data Source and Study Design**

We performed a retrospective cohort study using data from the Virtual International Stroke Trials Archive ICH (VISTA-ICH). 12 The VISTA database (www.vistacollaboration.org) houses anonymized, individual patient-level data from completed trials. Trials are eligible for inclusion in VISTA-ICH if they meet the following requirements: (1) documented entry criteria into a trial with a minimum of 50 randomized patients with ICH; (2) documented consent or waiver of consent from a local ethics board; (3) baseline assessment within 24 hours of ICH; (4) baseline assessment of neurological deficit at the time of admission; (5) confirmation of ICH by cerebral imaging within 7 days; (6) outcome assessment between 1 and 6 months with a validated stroke scale; and (7) data validation through monitoring. 13 The VISTA cohort used in this study consisted of only patients in the placebo arm of all trials and intervention arms of negative nonsurgical trials. Individual trials that contributed to the dataset used for this analysis were performed with institutional review board or regulatory approval. Individual trials obtained informed consent. Our analysis was approved by the Weill Cornell Medicine institutional review board. The data used in this analysis are restricted per the terms of VISTA-ICH's data use agreement and, therefore, cannot be shared directly with other investigators. However, investigators can obtain access to these data by submitting a formal application to VISTA-ICH.

### Liver Fibrosis Indices and Outcomes After Primary Intracerebral Hemorrhage

Neal S. Parikh, MD, MS; Hooman Kamel, MD; Babak B. Navi, MD, MS;
Costantino Iadecola, MD; Alexander E. Merkler, MD; Arun Jesudian, MD; Jesse Dawson, MD;
Guido J. Falcone, MD, ScD, MPH; Kevin N. Sheth, MD; David J. Roh, MD;
Mitchell S.V. Elkind, MD, MS; Daniel F. Hanley, MD; Wendy C. Ziai, MD, MPH;
Santosh B. Murthy, MD, MPH; on behalf of the VISTA-ICH Collaborators\*





Table 2. Associations Between Liver Fibrosis Indices and Admission Hematoma Volume, Hematoma Expansion, and Outcomes After ICH

	APRI*		FIB-4*		NFS*	
Outcome	OR (95% CI)	P Value	OR (95% CI)	P Value	OR (95% CI)	P Value
Hematoma expansion	1.6 (1.1–2.3)	0.01	1.9 (1.2–3.0)	0.01	1.2 (0.9–1.5)	0.22
All-cause mortality	1.8 (1.1–2.7)	0.01	2.0 (1.1–3.6)	0.02	1.2 (0.9–1.8)	0.22
Death or major disability	1.3 (0.9–1.8)	0.19	1.3 (0.8–2.1)	0.31	0.9 (0.8–1.2)	0.71
	Beta (SE)	P Value	Beta (SE)	P Value	Beta (SE)	P Value
Admission hematoma volume*†	0.20 (0.08)	0.01	0.27 (0.11)	0.01	0.03 (0.05)	0.53

APRI indicates Aspartate Aminotransferase—Platelet Ratio Index; FIB-4, Fibrosis-4; ICH, intracerebral hemorrhage; NFS, Nonalcoholic Fatty Liver Disease Fibrosis Score; and OR, odds ratio.

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<sup>\*</sup>Logarithmic transformation was performed to minimize skewness.

<sup>†</sup>Linear regression was used.

### Observational research studies

### Advantages

- Prospective enrollment, ascertainment, and follow-up
- Wealth of detailed assessments
- Often population based

### Disadvantages

- Small numbers of patients with diseases not of 1° interest
- Incomplete ascertainment of non-study related endpoints
- Trade-off b/w modern diagnostics vs length of follow-up



#### GCNKSS

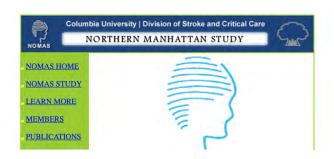
NOME POPULATION STATISTICS FORM ARSTRACT PREPARATION FORM MANUSCRIPT PREPARATION FORM



Greater Cincinnati / Northern Kentucky 5 County Area Population-Based Epidemiology of Stroke Study NIH NINDS (R01NS3078)





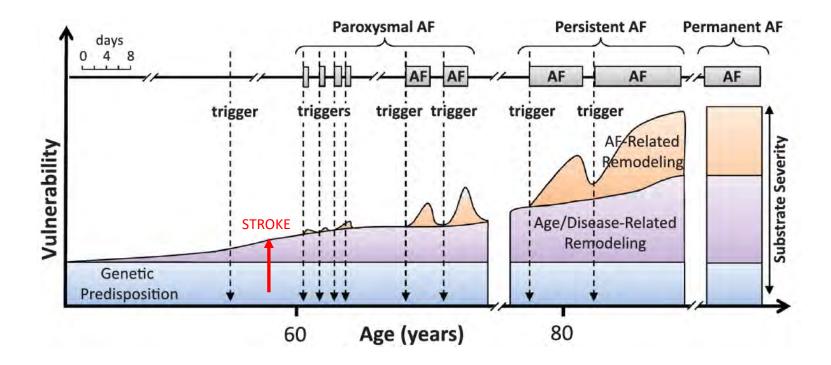








**¬ New York-Presbyterian** 



Cellular and Molecular Electrophysiology of Atrial Fibrillation Initiation, Maintenance, and Progression

Jordi Heijman,\* Niels Voigt,\* Stanley Nattel, Dobromir Dobrev Circ Res. 2014;114:1483-1499





Table 2. Associations Between P-Wave Morphology and Incident Ischemic Stroke

Characteristic	HR Per 1 SD Increase (95% CI)	
P-wave terminal force in lead V <sub>1</sub> , μV*ms	1.21 (1.02-1.44)	
P-wave mean area, μV*ms	1.16 (0.98-1.39)	
P-wave maximum area, μV*ms	1.16 (0.99-1.37)	
P-wave mean duration, ms	1.11 (0.92-1.34)	
P-wave maximum duration, ms	1.12 (0.93-1.35)	

Cl indicates confidence interval; and HR, hazard ratio.

### **Brief Reports**

P-Wave Morphology and the Risk of Incident Ischemic Stroke in the Multi-Ethnic Study of Atherosclerosis

Hooman Kamel, MD; Elsayed Z. Soliman, MD, MS; Susan R. Heckbert, MD, PhD; Richard A. Kronmal, PhD; W.T. Longstreth Jr, MD, MPH; Saman Nazarian, MD, PhD; Peter M. Okin, MD

(Stroke. 2014;45:2786-2788.)



**¬** New York-Presbyterian

### Association Between Left Atrial Abnormality on ECG and Vascular Brain Injury on MRI in the Cardiovascular Health Study

Hooman Kamel, MD; Traci M. Bartz, MS; W.T. Longstreth Jr, MD, MPH; Peter M. Okin, MD;
 Evan L. Thacker, PhD; Kristen K. Patton, MD; Phyllis K. Stein, PhD;
 Rebecca F. Gottesman, MD, PhD; Susan R. Heckbert, MD, PhD; Richard A. Kronmal, PhD;
 Mitchell S.V. Elkind, MD, MS; Elsayed Z. Soliman, MD, MS

Stroke. 2015;46:711-716

### Electrocardiographic Left Atrial Abnormality and Stroke Subtype in the Atherosclerosis Risk in Communities Study

Hooman Kamel, MD, <sup>1</sup> Wesley T. O'Neal, MD, MPH, <sup>2</sup> Peter M. Okin, MD, <sup>3</sup> Laura R. Loehr, PhD, <sup>4</sup> Alvaro Alonso, MD, PhD, <sup>5</sup> and Elsayed Z. Soliman, MD, MS<sup>6</sup>

ANN NEUROL 2015;78:670-678

## Electrocardiographic Left Atrial Abnormality and Risk of Stroke

Northern Manhattan Study

Hooman Kamel, MD; Madeleine Hunter; Yeseon P. Moon, MS; Shadi Yaghi, MD;Ken Cheung, PhD; Marco R. Di Tullio, MD; Peter M. Okin, MD; Ralph L. Sacco, MD;Elsayed Z. Soliman, MD, MSc, MS; Mitchell S.V. Elkind, MD, MS

Stroke. 2015;46:00-00

## Atrial Cardiopathy and the Risk of Ischemic Stroke in the CHS (Cardiovascular Health Study)

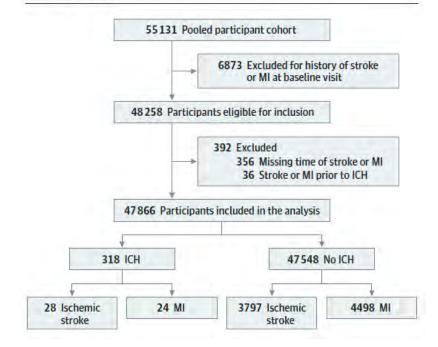
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W.T. Longstreth Jr, MD, MPH

Stroke. 2018;49:00-00





Figure 1. Flowchart Showing Inclusion Criteria for Analysis of Intracerebral Hemorrhage (ICH) and Subsequent Arterial Ischemic Events





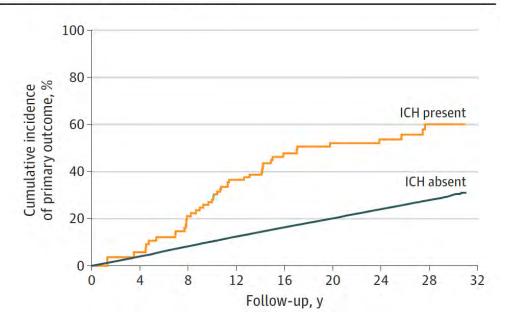
Association Between Intracerebral Hemorrhage and Subsequent Arterial Ischemic Events in Participants From 4 Population-Based Cohort Studies

Santosh B. Murthy, MD, MPH; Cenai Zhang, MS; Ivan Diaz, PhD; Emily B. Levitan, ScD; Silvia Koton, PhD; Traci M. Bartz, MS; Janet T. DeRosa, MS; Kevin Strobino, MS; Lisandro D. Colantonio, MD, PhD; Costantino ladecola, MD; Monika M. Safford, MD; Virginia J. Howard, PhD; W. T. Longstreth Jr, MD; Rebecca F. Gottesman, MD, PhD; Ralph L. Sacco, MD, MS; Mitchell S. V. Elkind, MD, MS; George Howard, DrPH; Hooman Kamel, MD





Figure 2. Kaplan-Meier Analysis of the Risk of an Arterial Ischemic Event After Intracerebral Hemorrhage (ICH)



#### JAMA Neurology | Original Investigation

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## Registries

### Advantages:

- Relatively granular data
- Often prospectively collected
- Standardized definitions

### **Disadvantages:**

- Not population based
- Most diseases lack registries
- Variable availability of follow-up



## Registries

Get With The Guidelines
National Surgical Quality Improvement Program
National Cardiovascular Data Registry
Cornell Acute Stroke Academic Registry (CAESAR)
Athens Stroke Registry



#### Figure 1. Study Population

93314 Medicare patients with acute ischemic stroke who were ≥65 y, discharged alive, and had a medical history of atrial fibrillation/flutter or documented persistent or paroxysmal atrial fibrillation/flutter between October 2011 and December 2014

#### 81652 Excluded

33217 Missing creatinine clearance

15 376 Discharged to hospice, transferred out, left against medical advice, or received comfort measures

12169 Taking an oral anticoagulant prior to stroke

12016 Documented contraindication for anticoagulant therapy

3787 Medical history of renal insufficiency, dialysis, or creatinine clearance <15 mL/min

2556 Not prescribed an oral anticoagulant, had ≥2 oral anticoagulants, or were missing these data at discharge

2531 Missing NIHSS score and were excluded from primary analysis

11662 Patients in primary analysis (1041 hospitals)

7621 Prescribed dabigatran, rivaroxaban, or apixaban at discharge

4041 Prescribed warfarin at discharge



Clinical Effectiveness of Direct Oral Anticoagulants vs Warfarin in Older Patients With Atrial Fibrillation and Ischemic Stroke Findings From the Patient-Centered Research Into Outcomes Stroke Patients Prefer and Effectiveness Research (PROSPER) Study

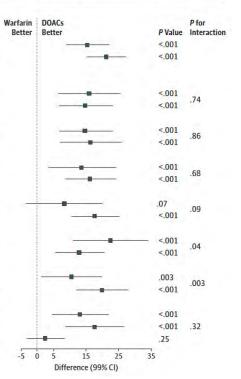
Ying Xian, MD, PhD; Haolin Xu, MS; Emily C. O'Brien, PhD; Shreyarish Shah, MD; Laine Thomas, PhD; Michael J. Pencina, PhD; Gregg C. Fonarow, MD; DalWal M. Olson, RN, PhD; Lee H. Schwamm, MD; Deepalk. Bhatt, MD, MPH; Eric E. Smith, MD, MPH; Deldre Hannah, MSN, RN; Lesley Malsch, BA; Barbara L. Lyte, MS; Eric D. Peterson, MD. MPH; Adrian F. Hennandez, MD, MHS





Figure 2. Association Between Direct Oral Anticoagulants (DOACs) at Discharge and Home Time During the First Year Postdischarge

	DOACs		Warfarin		Adjusted Analysis,	
ource	Patients, No.	Mean (SD)	Patients, No.	Mean (SD)	Difference (99% CI)	
rimary analysis	4041	287.2 (114.7)	7621	263.0 (127.3)	15.6 (9.0-22.1)	
ensitivity analysis	4773	287.1 (114.4)	9420	262.4 (127.4)	21.4 (15.3-27.4)	
ubgroup analyses						
Age, y						
65-80	2100	308.5 (100.6)	3847	284.7 (115.6)	16.2 (6.8-25.5)	
>80	1941	264.3 (124.3)	3774	241.0 (134.6)	15.0 (6.9-23.1)	
Sex						
Female	2277	281.4 (116.9)	4292	257.0 (127.9)	14.9 (6.8-22.9)	
Male	1764	294.8 (111.5)	3329	270.8 (126.1)	16.5 (7.0-26.0)	
Prior coronary arte	ry disease or	myocardial infarct	ion			
Yes	1217	277.7 (118.7)	2412	253.2 (131.9)	13.6 (3.1-24.1)	
No	2824	291.4 (112.8)	5209	267.6 (124.8)	16.4 (8.7-24.0)	
Prior stroke/transic	ent ischemic	attack				
Yes	949	272.9 (123.8)	1749	255.6 (129.3)	8.3 (-3.6-20.1)	
No	3092	291.7 (111.5)	5872	265.3 (126.6)	17.8 (10.4-25.1)	
Diabetes mellitus						
Yes	1012	281.2 (116.4)	1994	248.7 (131.3)	22.6 (11.1-34.1)	
No	3029	289.3 (114.2)	5627	268.1 (125.4)	13.2 (5.7-20.6)	
NIHSS score						
0-4	2295	311.5 (98.6)	3629	300.0 (105.7)	10.6 (1.3-19.8)	
>4	1746	255.4 (126.1)	3992	229.5 (135.6)	20.0 (12.0-27.9)	
Concomitant antip	latelet thera	ру				
Yes	1732	285.2 (116.4)	4629	263.3 (127.5)	13.2 (4.5-21.8)	
No	2309	288.8 (113.5)	2992	262.7 (126.9)	17.7 (8.6-26.7)	
Discharged home	2066	330.5 (81.5)	2957	326.4 (85.9)	2.6 (-3.2-8.3)	



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Clinical Effectiveness of Direct Oral Anticoagulants vs Warfarin in Older Patients With Atrial Fibrillation and Ischemic Stroke Findings From the Patient-Centered Research Into Outcomes Stroke Patients Prefer and Effectiveness Research (PROSPER) Study

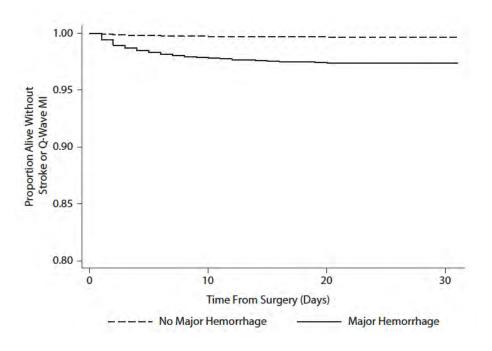
Ying Xian, MD, PhD, Haolin Xu, MS, Emily C. O'Brien, PhD; Shreyansh Shah, MD; Laline Thomas, PhD, Michael J. Pencina, PhD; Gregg C. Fonarow, MD; DalWal M. Olson, RN, PhD; Lee H. Schwamm, MD; Deepak L. Bhatt, MD, MPH; Eric E. Smith, MD, MPH; Deldre Hannah, MSN, RN; Lesley Maisch, BA; Barbara L. Lyte, MS; Eric D. Peterson, MD, MPH; Adrian F. Hennandez, MD, MHS





Table 2. Results of Proportional Hazards Analysis and Sensitivity Analyses of the Association Between Major Perioperative Hemorrhage and Subsequent Stroke or Q-Wave Myocardial Infarction

	HR for Subsequent Stroke or Q-Wave MI (95% CI)
Major perioperative hemorrhage	
Unadjusted analysis*	4.1 (3.4-4.9)
Primary adjusted analysis†	2.6 (2.2-3.1)
Sensitivity analysis of possible misclassification of major hemorrhage‡	4.2 (3.0–5.7)
Sensitivity analysis of missing preoperative creatinine values§	2.6 (2.2–3.1)
Sensitivity analysis of missing preoperative hematocrit values	2.6 (2.2–3.2)
Sensitivity analysis of surgical classifications¶	2.4 (2.0-2.8)







Association Between Major Perioperative Hemorrhage and Stroke or Q-Wave Myocardial Infarction

Hooman Kamel, S. Claiborne Johnston, John C. Kirkham, Christopher G. Turner, Jorge R. Kizer, Richard B. Devereux and Costantino Iadecola





## CAESAR piggybacks onto GWTG—Stroke

### Data quality in the American Heart Association Get With The Guidelines-Stroke (GWTG-Stroke): Results from a National Data Validation Audit

Ying Xian, MD, PhD, <sup>a</sup> Gregg C. Fonarow, MD, <sup>b</sup> Mathew J. Reeves, PhD, <sup>c</sup> Laura E. Webb, CCRP, <sup>a</sup> Jason Blevins, MPH, <sup>a</sup> Vladimir S. Demyanenko, MS, <sup>a</sup> Xin Zhao, MS, <sup>a</sup> DaïWai M. Olson, PhD, RN, <sup>a</sup> Adrian F. Hernandez, MD, MHS, <sup>a</sup> Eric D. Peterson, MD, MPH, <sup>a</sup> Lee H. Schwamm, MD, <sup>d</sup> and Eric E. Smith, MD, MPH <sup>c</sup> Durham, NC Los Angeles, C4; East Lansing, MI; Boston, MA; and Alberta, Canada

#### JAMA | Original Investigation

Association of Preceding Antithrombotic Treatment With Acute Ischemic Stroke Severity and In-Hospital Outcomes Among Patients With Atrial Fibrillation

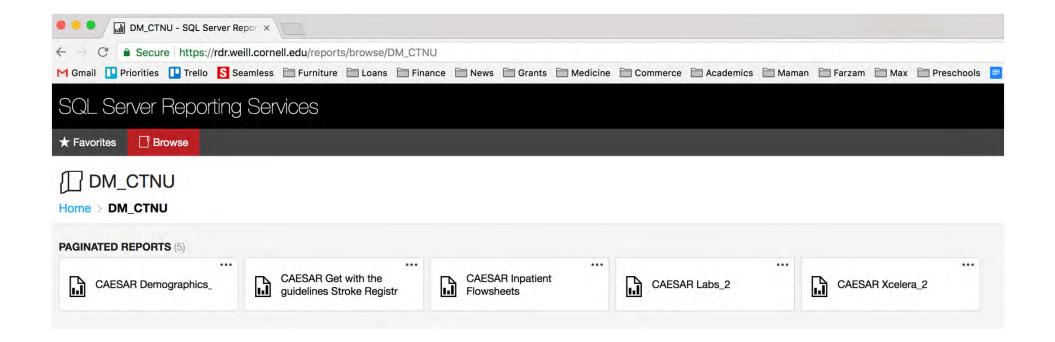
Ying Xian, MD, PhD; Emily C. O'Brien, PhD; Li Liang, PhD; Haolin Xu, MS; Lee H. Schwamm, MD; Gregg C. Fonarow, MD; Deepak L. Bhatt, MD, MPH; Eric E. Smith, MD, MPH; DaiWai M. Olson, PhD, RN; Lesley Maisch, BA; Deidre Hannah, MSN, RN; Brianna Lindholm, BA; Barbara L. Lytle, MS; Michael J. Pencina, PhD; Adrian F. Hernandez, MD, MHS; Eric D. Peterson, MD, MPH

Hospital type	Level	Composite accuracy	P	
Overall	-	96.1	_	
No. of ischemic	0-100	96.2	.87	
stroke discharges	101-300	96.0		
Transferred and	301+	96.0		
No. of beds	0-100	96.2	.19	
	101-200	96.1		
	201-300	96.3		
	301-500	96.1		
	501+	95.3		
Region	West	96.1	<.001	
	South	95.5		
	Midwest	96.6		
	Northeast	96.5		
Teaching status	Academic	96.4	.01	
	Nonacademic	95.8		
Primary stroke center	Yes	96.2	.67	
	No	96.1		
Paul Coverdell hospital	Yes	96.5	.12	
	No	96.1		

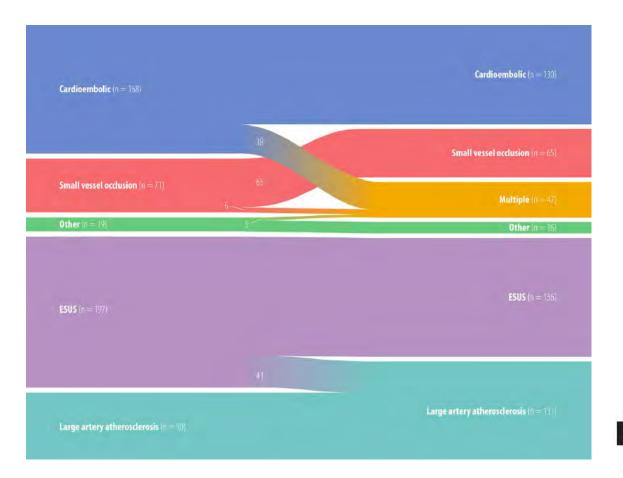
Table II Overall data accuracy among CWTG Stroke hospitals



## Research Data Repository









### **Original Contribution**

Reclassification of Ischemic Stroke Etiological Subtypes on the Basis of High-Risk Nonstenosing Carotid Plaque

Hooman Kamel, MD; Babak B. Navi, MD, MS; Alexander E. Merkler, MD; Hediyeh Baradaran, MD; Iván Díaz, PhD; Neal S. Parikh, MD, MS; Scott E. Kasner, MD; David J. Gladstone, MD, PhD; Costantino Iadecola, MD; Ajay Gupta, MD, MS



### Plans to link CAESAR to NY SPARCS







## Surveys

### Advantages:

- Nationally representative
- Outpatient data available in some surveys
- Longer time span of available data

### Disadvantages:

- No longitudinal follow-up
- Cannot account for multiple visits by same patient
- Often mostly limited to administrative data



#### RESEARCH LETTER

### National Trends in Ambulance Use by Patients With Stroke, 1997-2008

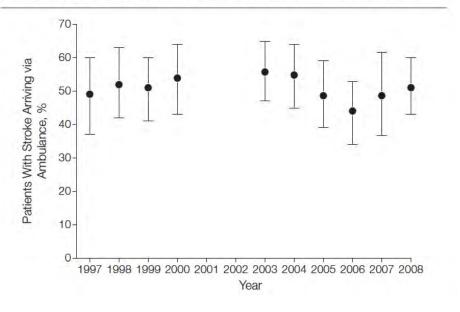
To the Editor: Thrombolytic therapy improves outcomes after ischemic stroke, but most patients are ineligible because they do not present in time. <sup>1</sup> This has prompted efforts to educate people to call 911 for signs of stroke because ambulance transportation results in faster arrival at the emergency department (ED). <sup>2</sup> Regional studies have suggested suboptimal ambulance use among patients with stroke, <sup>3</sup> but none has examined a nationally representative population or temporal trends since the approval of thrombolysis.

Methods. We analyzed data collected by the National Hospital Ambulatory Medical Care Survey (NHAMCS) between 1997 and 2008. An nationally representative random sample of 340 to 408 EDs was surveyed annually, reflecting a participation rate of 87% to 98% and constituting approximately 10% of US EDs. Staff used structured forms to collect data about a systematic random sample of patients over a random 4-week period. Analysis of this publicly available deidentified data set was exempt from evaluation by our institutional review boards.

We included patients with a primary diagnosis of ischemic stroke, defined by International Classification of Diseases, Ninth Revision codes that have been validated for identifying patients with acute stroke and used in other studies.5 Additionally, we included patients with subarachnoid hemorrhage, intracerebral hemorrhage, and transient ischemic attack because these can present similarly to ischemic stroke. Our outcome was arrival at the ED via ambulance. We used survey visit weights provided by the NHAMCS to estimate the national proportion of patients diagnosed with stroke in the ED each year who arrived by ambulance. We examined trends within subgroups defined by characteristics associated with ambulance use: age, sex, race, payment source, geographic region, and stroke subtype.6 We performed sensitivity analyses limited to ischemic stroke and excluding patients not admitted to the hospital or with additional ED diagnoses besides stroke.

A survey-weighted  $\chi^2$  test for trend was used to examine the statistical significance of changes in ambulance use over time. We used multiple logistic regression to analyze yearly trends in ambulance use for stroke while controlling for covariates. The threshold of statistical significance was a 2-sided  $\alpha$  level of .05. Statistical analysis was performed with Stata SE version 11 (StataCorp).

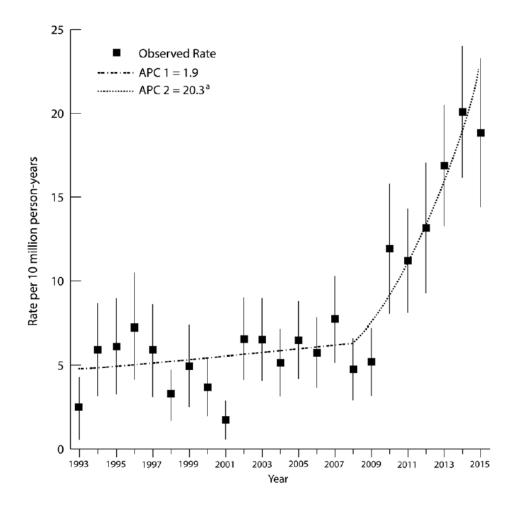
**Figure.** Proportion of Patients With Stroke Presenting to the Emergency Department via Ambulance From 1997 Through 2008

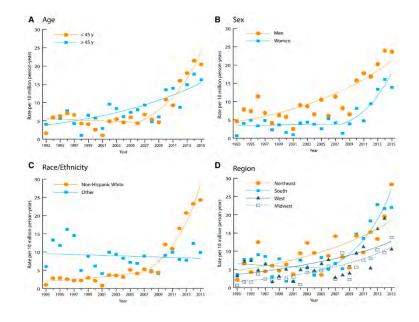


Hooman Kamel, MD Babak B. Navi, MD Jahan Fahimi, MD, MPH

JAMA, March 14, 2012-Vol 307, No. 10







#### National Trends in Hospitalizations for Stroke Associated With Infective Endocarditis and Opioid Use Between 1993 and 2015

Setareh Salehi Omran, MD; Abhinaba Chatterjee, BS; Monica L. Chen, BA; Michael P. Lerario, MD; Alexander E. Merkler, MD; Hooman Kamel, MD

Stroke March 2019



#### Administrative data

#### Advantages:

- Extensive longitudinal follow-up
- Adequate power even for rare diseases
- Can be population based

#### Disadvantages:

- Data limited to ICD codes
- No medication data
- No lab or imaging data



#### What are administrative data?

In using the term administrative data, we refer broadly to information generated during routine encounters between patients and the healthcare system. Such data may arise from hospitalizations, emergency department visits, clinic appointments, encounters for diagnostic testing, or pharmacy dispensing. Although the data may be collected primarily for administrative, regulatory, or billing purposes, the information can also be used for research. Administrative data can be obtained from many sources, including state governments that collect data from hospitals and emergency departments; payers such as Medicare, Medicaid, and commercial insurers; quality improvement registries; and federal surveys such as the Nationwide Inpatient Sample.

#### InterSECT

International Section for Early Career and Training

Jump Starting Your Clinical Research Career Using Administrative Data Sets for Stroke Research

Alexander E. Merkler, MD; Neal S. Parikh, MD; Hooman Kamel, MD

Stroke October 2018





### Very useful for health services research

The NEW ENGLAND JOURNAL of MEDICINE

#### SPECIAL ARTICLE

#### Readmissions, Observation, and the Hospital Readmissions Reduction Program

Rachael B. Zuckerman, M.P.H., Steven H. Sheingold, Ph.D., E. John Orav, Ph.D., Joel Ruhter, M.P.P., M.H.S.A., and Arnold M. Epstein, M.D.

JAMA | Original Investigation

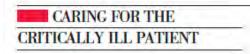
#### Association of Changing Hospital Readmission Rates With Mortality Rates After Hospital Discharge

Kumar Dharmarajan, MD, MBA; Yongfei Wang, MS; Zhenqiu Lin, PhD; Sharon-Lise T. Normand, PhD; Joseph S. Ross, MD, MHS; Leora I. Horwitz, MD, MHS; Nihar R. Desai, MD, MPH; Lisa G. Suter, MD; Elizabeth E. Drye, MD, SM; Susannah M. Bernheim, MD, MHS; Harlan M. Krumholz, MD, SM





#### Also useful for clinical research



# Incident Stroke and Mortality Associated With New-Onset Atrial Fibrillation in Patients Hospitalized With Severe Sepsis

Allan J. Walkey, MD, MSc
Renda Soylemez Wiener, MD, MPH
Joanna M. Ghobrial, MD
Lesley H. Curtis, PhD
Emelia J. Benjamin, MD, ScM

JAMA, November 23/30, 2011-Vol 306, No. 20



### But high potential for MIS-use

Research Practice No. by Domain	Required Research Practices for Conducting Studies Using the NIS
Data interpretation	
1	Identifying observations as hospitalization events rather than unique patients <sup>4,12</sup>
Research design	
2	Not performing state-level analyses <sup>11</sup>
3	Limiting hospital-level analyses to data from years 1988-2011 <sup>10,14</sup>
4	Not performing physician-level analyses 13,15
5	Avoiding use of nonspecific secondary diagnosis code to infer in-hospital events 16-20
Data analysis	
6	Using survey-specific analysis methods that account for clustering, stratification, and weighting <sup>6</sup>
7	Accounting for data changes in trend analyses spanning major transition periods in the data set (1997-1998 and 2011-2012) <sup>14,21</sup>

Table 2. Total Number of Instances of Nonadherence to Required Research Practices per Study for Publicatio
in 2015-2016 Using the National Inpatient Sample (NIS)

No. of nstances of Nonadherence, No. (%) of Studies				Estimates of Nonadherence for the Universe of NIS Studies (N = 1082) <sup>a</sup>		
to Required Practices	Overall (N = 120)	Journal Impact Factor <10 (n = 96) <sup>b</sup>	Journal Impact Factor ≥10 (n = 24) <sup>b</sup>	No. (95% CI)	% (95% CI)	
0	18 (15.0)	10 (10.4)	8 (33.3)	114 (50-177)	10.5 (4.7-16.4)	
1	28 (23.3)	21 (21.9)	7 (29.2)	229 (143-315)	21.2 (13.2-29.1)	
2	36 (30.0)	32 (33.3)	4 (16.7)	342 (244-440)	31.6 (22.6-40.7)	
3	30 (25.0)	25 (26.0)	5 (20.8)	269 (178-360)	24.9 (16.5-33.3)	
≥4	8 (6.7)	8 (8.3)	0	85 (28-142)	7.8 (2.5-13.1)	

Research

JAMA | Original Investigation

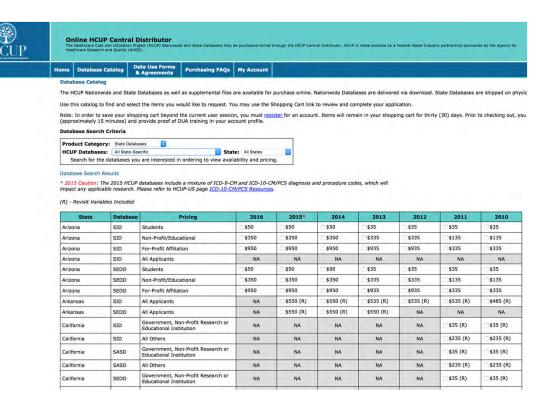
Adherence to Methodological Standards in Research Using the National Inpatient Sample

Rohan Khera, MD; Suveen Angraal, MBBS; Tyler Couch, BS; John W. Welsh, BS; Brahmajee K. Nallamothu, MD, MPH; Saket Girotra, MD, SM; Paul S. Chan, MD, MSc; Harlan M. Krumholz, MD, SM



### Statewide all-payer claims data

FL, NY, GA, MA, ...
Deidentified
Longitudinal tracking
25% US population
No outpatient data
No medication data







### Different possible designs

Descriptive – simple, robust

Associations – more difficult, more interesting



Table 2. Risk of Adverse Outcomes During Hospitalization for Patent Foramen Ovale Closure

Characteristic*	Rate of Adverse Outcomes (95% CI)		
Total adverse outcomes	7.0% (5.9%–8.2%)		
Individual adverse outcomes†			
Atrial fibrillation/flutter	3.7% (2.9%-4.6%)		
Vascular complication	3.0% (2.3%-3.9%)		
Hematoma/hemorrhage only	2.7% (2.0%–3.5%)		
Cardiac tamponade/perforation	0.5% (0.2%-0.9%)		
Death	0.3% (0.1%-0.6%)		
Pneumothorax/hemothorax	0.1% (0%-0.3%)		

#### Safety Outcomes After Percutaneous Transcatheter Closure of Patent Foramen Ovale

Alexander E. Merkler, MD\*; Gino Gialdini, MD\*; Shadi Yaghi, MD; Peter M. Okin, MD; Costantino Iadecola, MD; Babak B. Navi, MD, MS; Hooman Kamel, MD

Table 3. Risk of Adverse Outcomes After PFO Closure, Stratified by Subgroup

Characteristic*	No Adverse Outcome	Adverse Outcome	P Value
Age			<0.001
≤60 y	1176 (95.1)	61 (4.9)	
>60 y	579 (89.1)	71 (10.9)	
Sex			0.15
Female	844 (92.1)	72 (7.9)	
Male	911 (93.8)	60 (6.2)	
Race†			0.49
White	1212 (93.2)	88 (6.8)	
Other	505 (92.3)	42 (7.7)	
Indication for PFO closure			0.002
Ischemic stroke	463 (90.1)	51 (9.9)	
Transient ischemic attack	1292 (94.1)	81 (5.9)	
Medical comorbidities‡			<0.001
0	580 (94.9)	31 (5.1)	-
1	536 (94.2)	33 (5.8)	
2	327 (94.2)	20 (5.8)	
≥3	312 (86.7)	48 (13.3)	



 The role of empirical oral anticoagulation in patients with cardiomyopathy, reduced EF, and a history of stroke/TIA should also be investigated further in future studies because these patients were largely underrepresented in RCTs of prophylactic oral anticoagulation in patients with cardiomyopathies and reduced EF.

#### 5.4.5. Patent Foramen Ovale

Referenced	tecommendations for PFO deferenced studies that support recommendations are summarized in nline Deta Supplaments 38 and 39.						
COR	LOE	Recommendations					
1	C-EO	In patients with a nonlacunar ischemic stroke of undetermined cause and a PFO, recommendations for PFO closure versus medical management should be made jointly by the patient, a cardiologist, and a neurologist, taking into account the probability of a causal role for the PFO.					

the number needed to treat with device closure to prevent 1 recurrent stroke was 131 during 1 person-year of follow-up or 13 during 10 person-years of follow-up, which may be clinically important in this generally young population. Analysis of administrative claims data showed a 4.9% rate of serious periprocedural complication, including AF, in patients ≤60 years of age.<sup>562</sup> RCT data of PFO closure in patients >60 years of age are extremely limited,<sup>553</sup> and the rate of serious periprocedural complications in this older age group is significantly higher (10.9%).<sup>562</sup>

#### Recommendation-Specific Supportive Text

Recommendations for secondary stroke prevention in a patient with a PFO should be based on joint input from a neurologist with expertise in vascular neurology and a cardiologist with expertise in PFO closure (Figure 5).<sup>563,564</sup> Although 1 small trial with 120 patients did include some patients

562. Merkler AE, Gialdini G, Yaghi S, Okin PM, ladecola C, Navi BB, Kamel H. Safety outcomes after percutaneous transcatheter closure of patent foramen ovale. *Stroke*. 2017;48:3073–3077. doi: 10.1161/STROKEAHA.117.018501

#### Stroke

#### AHA/ASA GUIDELINE

# 2021 Guideline for the Prevention of Stroke in Patients With Stroke and Transient Ischemic Attack

A Guideline From the American Heart Association/American Stroke Association

Reviewed for evidence-based integrity and endorsed by the American Association of Neurological Surgeons and Congress of Neurological Surgeons.

Endorsed by the Society of Vascular and Interventional Neurology

The American Academy of Neurology affirms the value of this statement as an educational tool for neurologists.

Dawn O, Kleindorfer, MD, FAHA, Chair; Amytis Towfighi, MD, FAHA, Vice Chair; Seemant Chaturvedi, MD, FAHA; Kevin M. Cockroft, MD, MSc, FAHA; Jose Gutierrez, MD, MPH; Debbie Lombardi-Hill, BS, FAHA; Hooman Karnel, MD; Walter N, Keman, MD; Sleven J, Kittner, MD, MPH, FAHA; Enrique C, Leira, MD, MS, FAHA; Olive Lennon, PhD; James F, Meschia, MD, FAHA; Thanh N, Nguyen, MD, FAHA; Peter M, Pollak, MD; Pasquale Santangeli, MD, PhD; Anjai Z, Sharrief, MD, MPH, FAHA; Sidney C, Smith Jr, MD, FAHA; Tanya N, Turan, MD, MS, FAHA; Enda S, Williams, MD, FAHA





#### **Original Investigation**

# Perioperative Atrial Fibrillation and the Long-term Risk of Ischemic Stroke

Gino Gialdini, MD; Katherine Nearing, MD; Prashant D. Bhave, MD; Ubaldo Bonuccelli, MD; Costantino Iadecola, MD; Jeff S. Healey, MD; Hooman Kamel, MD

JAMA August 13, 2014 Volume 312, Number 6

**RESULTS** Of 1729 360 eligible patients, 24 711 (1.43%; 95% CI, 1.41%-1.45%) had new-onset perioperative atrial fibrillation during the index hospitalization and 13 952 (0.81%; 95% CI, 0.79%-0.82%) experienced a stroke after discharge. In a Cox proportional hazards analysis accounting for potential confounders, perioperative atrial fibrillation was associated with subsequent stroke both after noncardiac and cardiac surgery.

	Cumulative Rate of Hospitalization			
Type of Surgery	Perioperative Atrial Fibrillation	No Perioperative Atrial Fibrillation	Hazard Ratio (95% CI)	
Noncardiac	1.47 (1.24-1.75)	0.36 (0.35-0.37)	2.0 (1.7-2.3)	
Cardiac	0.99 (0.81-1.20)	0.83 (0.76-0.91)	1.3 (1.1-1.6)	

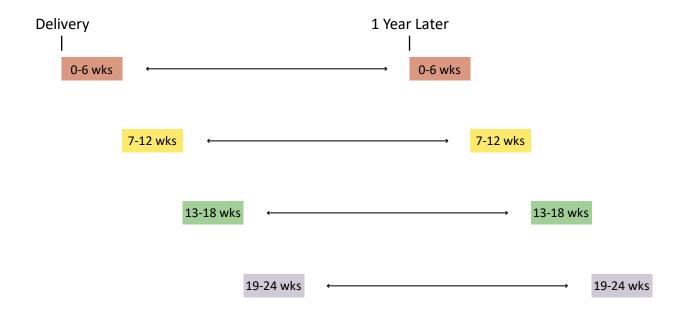
The association with stroke was significantly stronger for perioperative atrial fibrillation after noncardiac vs cardiac surgery (P < .001 for interaction).

**CONCLUSIONS AND RELEVANCE** Among patients hospitalized for surgery, perioperative atrial fibrillation was associated with an increased long-term risk of ischemic stroke, especially following noncardiac surgery.





### Duration of postpartum thrombosis

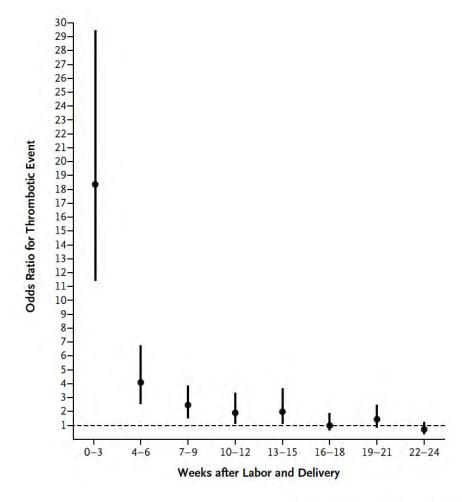




#### ORIGINAL ARTICLE

#### Risk of a Thrombotic Event after the 6-Week Postpartum Period

Hooman Kamel, M.D., Babak B. Navi, M.D., Nandita Sriram, B.S., Dominic A. Hovsepian, B.S., Richard B. Devereux, M.D., and Mitchell S.V. Elkind, M.D.





### Medicare data includes outpatient visits

### Byproduct of claims submitted by providers Key clinical information:

- Basic demographic information
- ICD diagnosis codes (in- and out-patient)
- ICD procedure codes (from hospital)
- CPT procedure codes (in- and out-patient)
- Date of death
- Date of insurance start/stop



#### Medicare data

#### Advantages:

- Includes ambulatory diagnoses
- CPT procedure codes more granular
- More accurate censoring
- Potentially linkable to medication data

#### Disadvantages:

- 65 years of age and older only
- Population-based?
- Expensive/cumbersome



#### Medicare Limited Datasets are attractive

#### **Traditional Medicare datasets:**

- Identifiable
- Need IRB approval
- More cumbersome DUA

#### Limited datasets:

- Essentially deidentified
- Much less expensive
- Straightforward DUA



#### JAMA Neurology | Original Investigation

# Association Between Cirrhosis and Stroke in a Nationally Representative Cohort

Neal S. Parikh, MD; Babak B. Navi, MD, MS; Yecheskel Schneider, MD; Arun Jesudian, MD; Hooman Kamel, MD JAMA Neurology Published online June 5, 2017

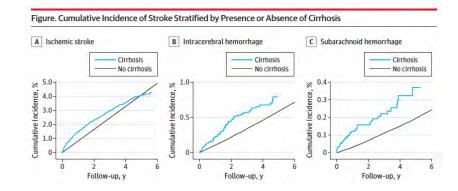


Table 3. Association Between Cirrhosis and Stroke in Medicare Beneficiaries by Type of Cirrhosis and Type of Stroke

	HR (95% CI)					
No. of Patients	All Stroke	Ischemic Stroke	ICH	SAH		
15 586	1.4 (1.3-1.5)	1.3 (1.2-1.5)	1.9 (1.5-2.4)	2.4 (1.7-3.5)		
3255	1.5 (1.2-1.8)	1.4 (1.2-1.7)	2.3 (1.5-3.7)	1.6 (0.7-4.0)		
11 164	1.5 (1.3-1.6)	1.4 (1.2-1.5)	2.1 (1.6-2.7)	2.8 (1.9-4.1)		
6043	1.7 (1.5-2.0)	1.6 (1.4-1.9)	2.5 (1.8-3.5)	2.8 (1.7-5.0)		
5542	1.5 (1.3-1.7)	1.3 (1.1-1.5)	2.5 (1.8-3.5)	3.1 (1.9-5.2)		
13 384	1.4 (1.3-1.6)	1.3 (1.2-1.5)	2.1 (1.6-2.6)	2.4 (1.6-3.5)		
	Patients 15 586 3255 11 164 6043 5542	No. of Patients All Stroke  15 586 1.4 (1.3-1.5)  3255 1.5 (1.2-1.8)  11 164 1.5 (1.3-1.6)  6043 1.7 (1.5-2.0)  5542 1.5 (1.3-1.7)	No. of Patients         All Stroke         Ischemic Stroke           15 586         1.4 (1.3-1.5)         1.3 (1.2-1.5)           3255         1.5 (1.2-1.8)         1.4 (1.2-1.7)           11 164         1.5 (1.3-1.6)         1.4 (1.2-1.5)           6043         1.7 (1.5-2.0)         1.6 (1.4-1.9)           5542         1.5 (1.3-1.7)         1.3 (1.1-1.5)	No. of Patients         All Stroke         Ischemic Stroke         ICH           15 586         1.4 (1.3-1.5)         1.3 (1.2-1.5)         1.9 (1.5-2.4)           3255         1.5 (1.2-1.8)         1.4 (1.2-1.7)         2.3 (1.5-3.7)           11 164         1.5 (1.3-1.6)         1.4 (1.2-1.5)         2.1 (1.6-2.7)           6043         1.7 (1.5-2.0)         1.6 (1.4-1.9)         2.5 (1.8-3.5)           5542         1.5 (1.3-1.7)         1.3 (1.1-1.5)         2.5 (1.8-3.5)		



### Merging in external data can add value

Medicare datasets include county codes and physician NPI numbers

#### Possible to merge in:

- County demographics and socioeconomic indices
- Physician board-certification, etc



#### RESEARCH LETTER

#### Medical Specialties of Clinicians Providing Mechanical Thrombectomy to Patients With Acute Ischemic Stroke in the United States

JAMA Neurology Published online January 25, 2018

Hooman Kamel, MD Caroline D. Chung, BA Gbambele J. Kone, BS Ajay Gupta, MD Nicholas A. Morris, MD Matthew E. Fink, MD Babak B. Navi, MD, MS

Table. Clinician Specialty Among Cases of Mechanical Thrombectomy for Ischemic Stroke in a 5% Sample of Medicare Beneficiaries

	No. (%)		
Clinician Specialty	CMS <sup>a</sup>	NPIb	Google <sup>c</sup>
Radiology	341 (61.4)	328 (59.2)	316 (56.9)
Neurosurgery	110 (19.8)	99 (17.9)	95 (17.1)
Neurology	91 (16.4)	108 (19.5)	131 (23.6)
Other	13 (2.4)	20 (3.4)	13 (2.4)

Abbreviations: CMS, Centers for Medicare and Medicaid Services; NPI, National Provider Identifier.



<sup>&</sup>lt;sup>a</sup> Provider specialty as self-designated at the time of enrollment in Medicare.

<sup>&</sup>lt;sup>b</sup> Provider specialty as self-designated on the NPI application.

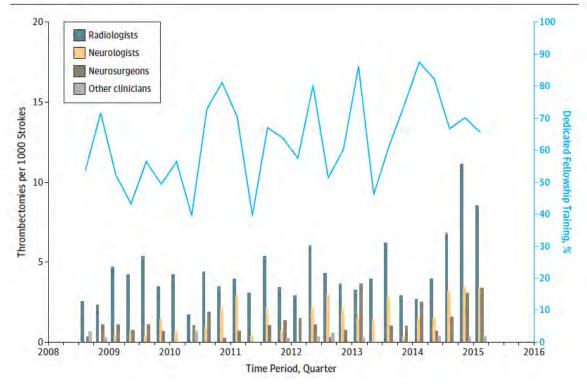
<sup>&</sup>lt;sup>c</sup> Provider specialty as ascertained from review of publicly available online records.

#### RESEARCH LETTER

#### Medical Specialties of Clinicians Providing Mechanical Thrombectomy to Patients With Acute Ischemic Stroke in the United States

JAMA Neurology Published online January 25, 2018

Hooman Kamel, MD Caroline D. Chung, BA Gbambele J. Kone, BS Ajay Gupta, MD Nicholas A. Morris, MD Matthew E. Fink, MD Babak B. Navi, MD, MS Figure. Temporal Changes in Stroke-Related Mechanical Thrombectomy Rates and Backgrounds of Performing Physicians





#### Commercial claims data

Essentially same as Medicare but <65 years of age

Can include medications, lab results

Optum, Thomson Reuters, Premier, etc.

Extremely expensive

Population-based?



#### **Original Investigation**

## Population-Level Evidence for an Autoimmune Etiology of Epilepsy

Mei-Sing Ong, PhD; Isaac S. Kohane, MD, PhD; Tianxi Cai, PhD; Mark P. Gorman, MD; Kenneth D. Mandl, MD, MPH

**IMPORTANCE** Epilepsy is a debilitating condition, often with neither a known etiology nor an effective treatment. Autoimmune mechanisms have been increasingly identified.

**OBJECTIVE** To conduct a population-level study investigating the relationship between epilepsy and several common autoimmune diseases.

**DESIGN, SETTING, AND PARTICIPANTS** A retrospective population-based study using claims from a nationwide employer-provided health insurance plan in the United States, Participants were beneficiaries enrolled between 1999 and 2006 (N = 2 518 034).

MAIN OUTCOMES AND MEASURES We examined the relationship between epilepsy and 12 autoimmune diseases: type 1 diabetes mellitus, psoriasis, rheumatoid arthritis, Graves disease, Hashimoto thyroiditis, Crohn disease, ulcerative colitis, systemic lupus erythematosus, antiphospholipid syndrome, Sjögren syndrome, myasthenia gravis, and celiac disease.

**RESULTS** The risk of epilepsy was significantly heightened among patients with autoimmune diseases (odds ratio, 3.8; 95% CI, 3.6-4.0; P < .001) and was especially pronounced in children (5.2; 4.1-6.5; P < .001). Elevated risk was consistently observed across all 12 autoimmune diseases,

**CONCLUSIONS AND RELEVANCE** Epilepsy and autoimmune disease frequently co-occur; patients with either condition should undergo surveillance for the other. The potential role of autoimmunity must be given due consideration in epilepsy so that we are not overlooking a treatable cause.

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Figure 1. Risk of Epilepsy in Children (<18 Years) and Nonelderly Adults (≤65 Years) With Autoimmune Disease Compared With Individuals Without Autoimmune Disease

Autoimmune Disease	No.	Epilepsy, %	OR (95% CI)					
Antiphospholipid syndrome	5423	3.2	9.0 (7.7-10.5)	<u></u>			$\vdash$	
SLE	9696	2.5	7.4 (6.5-8.4)				$\vdash \blacksquare \vdash$	
Type 1 diabetes mellitus	43704	1.8	5.2 (4.9-5.6)				H	
Myasthenia gravis	1070	1.7	4.9 (3.1-7.8)			<b>—</b>		
Celiac disease	1885	1.5	4.5 (3.1-6.5)			_	$\dashv$	
Sjögren syndrome	3614	1.5	4.3 (3.2-5.6)			_	1	
Rheumatoid arthritis	22890	1.2	3.5 (3.1-4.0)			H		
Crohn disease	8774	1.1	3.1 (2.5-3.8)		$\vdash$			
Ulcerative colitis	10690	0.9	2.5 (2.1-3.1)		<b>⊢</b> ■	$\dashv$		
Hashimoto thyroiditis	9830	0.8	2.4 (2.0-3.0)		<b>—</b>	$\dashv$		
Graves disease	9758	0.8	2.2 (1.8-2.8)		<b>—</b>	+		
Psoriasis	23 542	0.7	1.9 (1.6-2.3)		$\vdash$			
Any of the above	135394	1.3	3.8 (3.6-4.0)					
				1.0	2.0	4.0	8.0	16.0
					0	R, Log Sca	ale	



### Admin data are a great resource for trainees

Publicly available

Not disease specific

Can be quickly certified as IRB exempt



Figure. Cumulative Incidence of Stroke Stratified by Presence or Absence of Cirrhosis

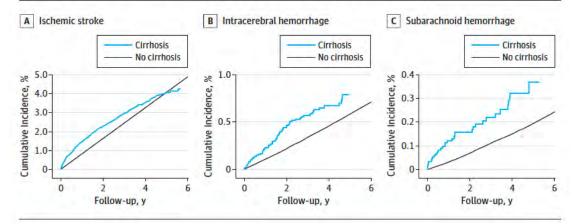


Table 3. Association Between Cirrhosis and Stroke in Medicare Beneficiaries by Type of Cirrhosis and Type of Stroke

		HR (95% CI)					
Type of Cirrhosis	No. of Patients	All Stroke	Ischemic Stroke	ICH	SAH		
Cirrhosis	15 586	1.4 (1.3-1.5)	1.3 (1.2-1.5)	1.9 (1.5-2.4)	2.4 (1.7-3.5)		
Alcohol-related cirrhosis	3255	1.5 (1.2-1.8)	1.4 (1.2-1.7)	2.3 (1.5-3.7)	1.6 (0.7-4.0)		
Non-alcohol-related cirrhosis	11 164	1.5 (1.3-1.6)	1.4 (1.2-1.5)	2.1 (1.6-2.7)	2.8 (1.9-4.1)		
Decompensated cirrhosis	6043	1.7 (1.5-2.0)	1.6 (1.4-1.9)	2.5 (1.8-3.5)	2.8 (1.7-5.0)		
Cirrhosis diagnosed by GI	5542	1.5 (1.3-1.7)	1.3 (1.1-1.5)	2.5 (1.8-3.5)	3.1 (1.9-5.2)		
Cirrhosis with imaging or biopsy <sup>a</sup>	13 384	1.4 (1.3-1.6)	1.3 (1.2-1.5)	2.1 (1.6-2.6)	2.4 (1.6-3.5)		

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### Association Between Cirrhosis and Stroke in a Nationally Representative Cohort

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#### ARTICLE

#### Prevalence of Cervical Artery Dissection Among Hospitalized Patients With Stroke by Age in a Nationally Representative Sample From the United States

Yahya B. Atalay, MD, Pirouz Piran, MD, Abhinaba Chatterjee, BS, Santosh Murthy, MD, MPH, Babak B. Navi, MD, MS, Ava L. Liberman, MD, Joseph Dardick, BS, Cenai Zhang, MS, Hooman Kamel, MD, MS, and Alexander E. Merkler, MD, MS

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#### Circulation

#### RESEARCH LETTER

### Association Between Cervical Artery Dissection and Aortic Dissection

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# Association between Pregnancy and Cervical Artery Dissection

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ANN NEUROL 2020;88:596-602



#### JAMA Neurology | Original Investigation

#### Distributional Validity and Prognostic Power of the National Institutes of Health Stroke Scale in US Administrative Claims Data

Hamidreza Saber, MD, MPH; Jeffrey L. Saver, MD

#### RESEARCH LETTER

#### Prehospital Treatment of Status Epilepticus in the United States

JAMA November 16, 2021 Volume 326, Number 19

Elan L. Guterman, MD, MAS James F. Burke, MD, MS Karl A. Sporer, MD

JAMA Neurology | Original Investigation

# Risk of Pregnancy-Associated Stroke Across Age Groups in New York State

Eliza C. Miller, MD; Hajere J. Gatollari, MPH; Gloria Too, MD; Amelia K. Boehme, PhD, MSPH; Lisa Leffert, MD; Mitchell S. V. Elkind, MD, MS; Joshua Z. Willey, MD, MS

Circulation: Cardiovascular Quality and Outcomes

#### **ORIGINAL ARTICLE**

Utilization and Availability of Advanced Imaging in Patients With Acute Ischemic Stroke

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Songmi Lee, MS
Rania Abdelkhaleq<sup>®</sup>, BS
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Sunil A. Sheth<sup>®</sup>, MD



### Methodological considerations

Understand source data

Beware of misclassification

Beware of nonrepresentative samples

Data management and ethics



#### Must understand nature of source data

#### Misunderstandings of source data

- Assuming longitudinal design when data are cross-sectional
- Effects of undocumented mortality
- Physician claims versus hospital diagnoses
- Secular trends in coding



#### Must understand nature of source data

#### Know your dataset inside and out

- Read documentation
- Attend training
- Read papers
- Speak with experts



#### Beware of misclassification

Power ≠ misclassification

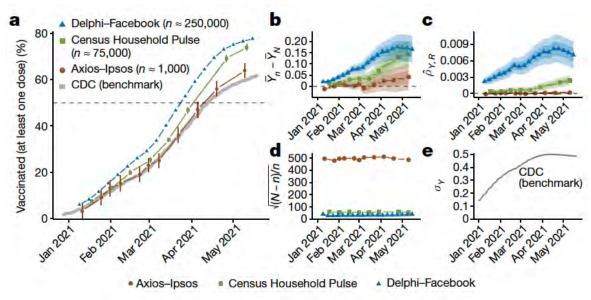
Misclassification dilutes ability to detect differences

Cannot be fully overcome by increasing sample size

Thought experiment: what if you gave half your treatment group placebo instead?



### Beware of nonrepresentative samples



# Unrepresentative big surveys significantly overestimated US vaccine uptake

Nature | www.nature.com
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### Key message

Include patients who reflect target population, in sufficient numbers

Measure the right things, and do it accurately

### Key message

Include patients who reflect target population, in sufficient numbers

Measure the right things, and do it accurately

NO AMOUNT OF STATISTICAL WIZARDRY CAN HELP YOU IF YOU FAIL IN THESE TWO STEPS



### Data management

#### Security

- Always use an encrypted, ITS-tagged computer
- Strict password hygiene
- Be aware of phishing
- Do not use unapproved cloud services for research data



### Data management

#### Reproducibility

- Start with source data file
- Use a script to manipulate/analyze data
- Save resulting file as a different file
- Errors can be corrected without affecting the source data

### Data management

#### Audit trail

- At all times, act as if FDA, DHHS, IRB, and Office of Research Compliance are about to come and audit your study
- Clearly organized and named folders and file names
- Detailed comments in your analysis script
- Readme files, notes, whatever you need to be able to remember and explain source of data and its analysis from start to finish



### Ethics and regulatory approvals

Ensure all research activities have IRB approval Request IRB exemption for analysis of deidentified data Request expedited review and waiver of informed consent for:

- Minimal risk
- Research not practicable without exemption

Anything else will require full-board review and informed consent



#### Make a win-win deal with your chair/chief/mentor:

- They provide modest start-up funding (\$1,000-\$5000)
- You provide the time/effort to acquire, learn, clean dataset
- Provides crucial career development for you and becomes lasting departmental resource



Master the literature on a topic
Be alert to new ideas and techniques
Observe closely
Allow time for reflection and creativity
Find a good mentor
Be tenacious



Remember: Initial projects are primarily to build skills Rigorously assess feasibility before committing time Know when to call off a project



Think creatively about diagnoses, not clinical data
Think through biases and minimize them
Use validated codes or validate codes yourself
Lots of sensitivity analyses
Be careful about making strong claims
Remember Bayes, guard against p-hacking



Form collaborations with researchers involved in prospective research studies in your area of interest Reach out to investigators at your institution, network at meetings

Be enthusiastically persistent!



#### Maintain a balanced research diet

- Chart reviews (limit your portions!)
- Administrative data
- Secondary analyses of prospective data
- Ancillary studies
- Prospective observational studies
- Clinical trials



#### We love to collaborate!

Please feel free to reach out anytime

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