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Trends in emergency department visits and hospitalizations for anaphylaxis among US older adults: 2006-2014

Anna Chen Arroyo, MD, MPH; Lacey B. Robinson, MD; Rebecca E. Cash PhD, MPH; Mohammad Kamal Faridi, MPH; Kohei Hasegawa MD, Carlos A. Camargo Jr., MD, DrPH

Address manuscript correspondence to: Anna Chen Arroyo, MD, MPH, 300 Pasteur Drive MC 5236, Stanford, CA 94305, Phone: 650-723-3200 Fax: 650-503-0169 Email: annaarroyo@stanford.edu

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Acute Allergic Reactions	ICD-9-0	9-CM Code				
Drug Allergy	693.0	Dermatitis due to drug				
	995.2	Unspecified adverse effect due to unspecified drug				
	995.1	Angioneurotic edema				
	995.4	Shock due to Anesthesia				
Venom Allergy	989.5	Toxic effect of venom				
Food Allergy	693.1	Dermatitis due to food				
	995.60	Anaphylactic shock due to unspecified food				
	995.61	Anaphylactic shock due to peanuts				
	995.62	Anaphylactic shock due to crustaceans				
	995.63	Anaphylactic shock due to fruits & vegetables				
	995.64	Anaphylactic shock due to tree nuts & seeds				
	995.65	Anaphylactic shock due to fish				
	995.66	Anaphylactic shock due to food additives				
	995.67	Anaphylactic shock due to milk products				
	995.68	Anaphylactic shock due to eggs				
	995.69	Anaphylactic shock due to other specified food				
	995.70	Adverse food reaction, not classified elsewhere				
Other Allergy	995.0	Other anaphylactic shock				
	995.3	Allergy, unspecified				
	708.0	Allergic urticaria				
	999.4	Anaphylactic reaction to serum				
	999.41	Anaphylactic reaction due to administration of blood and blood products				
	999.42	Anaphylactic reaction due to vaccination				
	999.49	Anaphylactic reaction due to other serum				

Table E1. ICD-9-CM diagnostic codes used for acute allergic reactions

		c coues and algorithm used for anaphynanis				
Anaphylaxis Method I	995.60	Anaphylactic shock due to unspecified food				
	995.61	Anaphylactic shock due to peanuts				
	995.62	Anaphylactic shock due to crustaceans				
	995.63	Anaphylactic shock due to fruits & vegetables				
	995.64	Anaphylactic shock due to tree nuts & seeds				
	995.65	Anaphylactic shock due to fish				
	995.66	Anaphylactic shock due to food additives				
	995.67	Anaphylactic shock due to milk products				
	995.68	Anaphylactic shock due to eggs				
	995.69	Anaphylactic shock due to other specified food				
	995.0	Other anaphylactic shock				
Anaphylaxis Method II ¹⁴						
Symptoms						
Skin-mucosal tissue involvement	Conjunctival edema (372.73), Edema of eyelid (374.82), Edema of larynx (478.6), Edema of pharynx or nasopharynx (478.25), Flushing (782.62), Urticaria (708), Allergic urticaria (708.0), Idiopathic urticaria (708.1), Urticaria due to cold and heat (708.2), Dermatographic urticaria (708.3), Vibratory urticaria (708.4), Cholinergic urticaria (708.5), Other specified urticaria (708.8), Urticaria, unspecified (708.9), Pruritus and related conditions (698), Pruritus of genital organs (698.1), Other specified pruritic conditions (698.8), Unspecified pruritic disorder (698.9)					
Respiratory compromise	Acute respiratory failure (518.81), acute respiratory distress (518.82), dyspnea/respiratory distress (786, 786.00–786.09), stridor (786.1), extrinsic asthma (493.0) or unspecified asthma (493.9)					
Reduced blood pressure	Hypotension (458, 458.0, 458.2, 458.8, 458.9), or syncope/collapse (780.2)					
Gastrointestinal symptoms	Allergic gastroenteritis/colitis (558.3), nausea/vomiting (787.0), nausea with vomiting (787.01), vomiting alone (787.03), or abdominal pain (789.0).					
Algorithm	Skin-mucosal tissue involvement was found in conjunction with shock caused by anesthesia (995.4)					
		Respiratory compromise was found in conjunction with shock caused by anesthesia (995.4) or angioneurotic edema (995.1)				
	Reduced blood pressure was found in conjunction with skin-mucosal involvement, angioneurotic edema (995.1), the toxic effect of venom (989.5), or an Ecode indicating the cause of poisoning was a venomous animal/plant (E905, E905.3, E905.5, E905.8, E905.9)					

Table E2. ICD-9-CM diagnostic codes and algorithm used for anaphylaxis

Reduced blood pressure and gastrointestinal symptoms were found in conjunction with an unspecified adverse
effect caused by the correct administration of a drug, medicinal, and biologic substance (995.2) or an
 unspecified allergic reaction (995.3)
Reduced blood pressure and skin-mucosal tissue involvement were found in conjunction with an unspecified
adverse effect caused by the correct administration of a drug, medicinal, and biologic substance (995.2) or an
 unspecified allergic reaction (995.3)
Respiratory compromise and reduced blood pressure were found in conjunction with an unspecified adverse
effect caused by the correct administration of a drug, medicinal, and biologic substance (995.2) or an
unspecified allergic reaction (995.3)
Respiratory compromise and gastrointestinal symptoms were found in conjunction with an unspecified adverse
effect caused by the correct administration of a drug, medicinal, and biologic substance (995.2); an unspecified
allergic reaction (995.3); skin-mucosal involvement; the toxic effect of venom (989.5); or an Ecode indicating
the cause of poisoning was a venomous animal/plant (E905, E905.3, E905.5, E905.8, E905.9)
Respiratory compromise and skin-mucosal tissue involvement were found in conjunction with an unspecified
adverse effect caused by the correct administration of a drug, medicinal, and biologic substance (995.2); an
unspecified allergic reaction (995.3); the toxic effect of venom (989.5); or an Ecode indicating the cause of
poisoning was a venomous animal/plant (E905, E905.3, E905.5, E905.8, E905.9)
Gastrointestinal symptoms and skin-mucosal tissue involvement were found in conjunction with an unspecified
adverse effect caused by the correct administration of a drug, medicinal, and biologic substance (995.2); an
unspecified allergic reaction (995.3); the toxic effect of venom (989.5); or an Ecode indicating the cause of
poisoning was a venomous animal/plant (E905, E905.3, E905.5, E905.8, E905.9)

Table E3. Characteristics of anaphylaxis-related ED visits among younger adults (18-64 years) and older adults (≥65 years) in
the US, 2006-2014 National Emergency Department Sample

		Adults (18-64 years)				Older Adults (≥65 years)			
	Unweighted Frequency	Weigl	nted Free	quency	Unweighted Frequency	Weig	nted Freq	uency	
Characteristics		Frequency	%	95% CI		Frequency	%	95% CI	
Overall number	81,169	365,509	100		20,989	93,795	100		
Sex									
Male	32,676	147,311	40	40-41	8,200	36,801	39	39-40	
Female	48,479	218,130	60	59-60	12,786	56,973	61	60-61	
Missing	14				<10				
Epinephrine Use	5,354	24,100	7	6-7	418	1,933	2	1.8-2.3	
Number of ED procedures				5					
0	74,311	333,408	91	90-92	19,757	88,143	94	93-95	
1	2,668	12,224	2	3-4	493	2,203	2	2-3	
2 or more	4,190	19,878	5	4-6	739	3,415	4	3-4	
Any chronic condition	43,933	196,606	54	53-55	18,194	81,230	87	86-87	
Type of Trigger									
Drug	14,968	67,158	18	18-19	7,428	33,013	35	34-36	
Venom	5,647	25,142	7	6-7	1,267	5,711	6	5.7-6.5	
Food	14,900	68,103	19	18-19	1,304	5,920	6	6.0-6.7	
Other/multiple	45,654	205,107	56	56-57	10,990	49,152	52	52-53	

Abbreviations: US, United States; CI, confidence interval; ED, emergency department.

	Hospital	alization Cardiac arrest		/Intubation	Death	
Risk Factors	OR 95% CI		OR	95% CI	OR 95% C	
Year	0.92	0.89-0.94	0.99	0.96-1.02	1.03	0.97-1.09
Age						
65-74 years	1.00 (referent)		1.00 (referent)		1.00 (referent)	
75-84 years	1.43	1.27-1.62	1.34	1.16-1.55	1.29	0.96-1.75
≥85 years	2.12	1.72-2.62	1.46	1.18-1.80	2.81	1.97-4.01
Sex						
Male	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Female	0.83	0.75-0.93	0.83	0.72-0.96	0.98	0.74-1.31
Primary Health Insurance						
Private	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Public	1.37	1.15-1.62	1.27	0.99-1.63	1.22	0.65-2.27
Self-pay	1.39	0.74-2.64	1.32	0.58-3.03	Not calculated	
Other	1.13	0.61-2.10	1.10	0.41-2.94	1.77	0.36-8.61
Median household income quartile			·			
1 (lowest)	1.38	1.14-1.66	1.59	1.28-1.98	1.18	0.77-1.80
2	1.18	0.99-1.40	1.40	1.14-1.72	1.38	0.94-2.02
3	1.16	0.98-1.37	1.24	1.00-1.52	1.17	0.78-1.75
4 (highest)	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Season of Discharge						
First quarter (Jan - Mar)	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Second quarter (Apr - Jun)	1.00	0.86-1.17	0.83	0.68-1.01	0.77	0.52-1.15
Third quarter (Jul - Sep)	0.93	0.80-1.07	0.80	0.67-0.96	0.86	0.59-1.27
Fourth quarter (Oct - Dec)	1.06	0.90-1.25	0.99	0.82-1.19	0.96	0.66-1.41
Hospital Region						
Northeast	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Midwest	0.61	0.49-0.77	0.77	0.61-0.97	0.55	0.35-0.87
South	0.52	0.41-0.64	0.85	0.69-1.05	0.56	0.38-0.84
West	0.64	0.51-0.81	1.06	0.84-1.35	0.67	0.42-1.08
Hospital Status						
Non-Metropolitan	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Metropolitan, Non-teaching	1.77	1.48-2.12	1.68	1.30-2.18	1.79	0.98-3.28
Metropolitan Teaching	1.98	1.65-2.37	2.15	1.68-2.75	2.42	1.34-4.40
Weekend presentation						

Table E4. Clinical risk factors for severe anaphylaxis (hospitalization, cardiac arrest/intubation, or death) among US olderadults – Method I only, 2006-2014 National Emergency Department Sample

No	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Yes	0.94	0.83-1.07	0.96	0.80-1.14	1.02	0.72-1.43
Type of Trigger						
Food	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Drug	3.75	3.10-4.54	2.39	1.80-3.18	5.60	2.47-12.70
Venom	0.52	0.29-0.94	1.60	0.70-3.65	Not calculated	
Other	1.95	1.68-2.28	2.09	1.62-2.69	5.23	2.37-11.54
Epinephrine use in ED						
Yes	1.00 (referent)		1.00 (referent)		1.00 (referent)	
No	11.59	7.12-18.88	5.21	2.08-13.10	2.07	0.50-8.68
Any chronic condition						
No	1.00 (referent)		1.00 (referent)		1.00 (referent)	
Yes	15.77	12.70-19.58	19.96	10.69-37.26	17.96	4.13-78.13

Note: Number of chronic conditions had to be collapsed due to unstable estimates for the 2 or more group

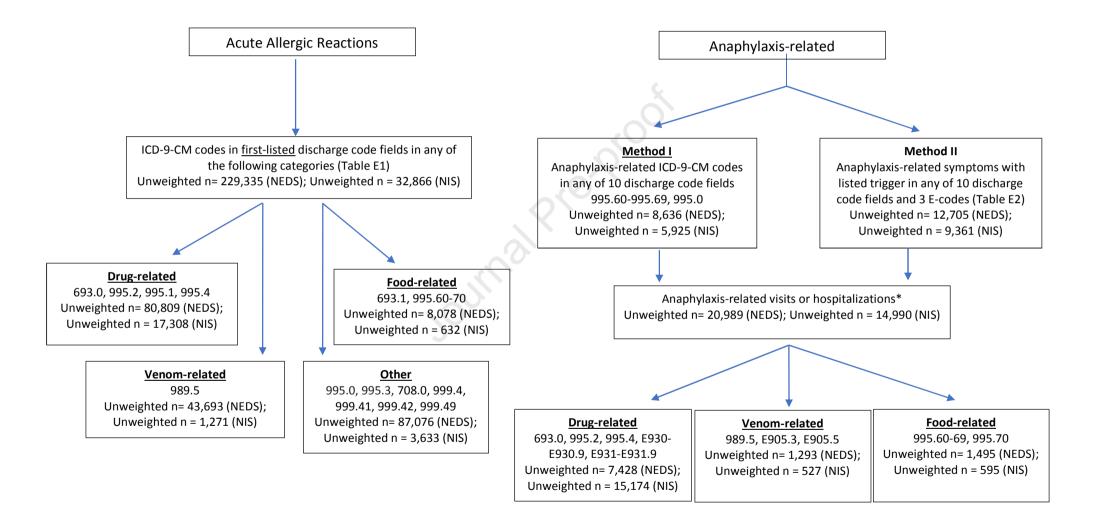
Survey weights were used to construct this model. Unweighted n = 8,273 (due to missing data); weighted n = 37,092.

	Cardiac arres	Death		
Risk Factors	OR	95% CI	OR	95% CI
Year	1.00	0.97-1.02	0.98	0.93-1.04
Age				
65-74 years	1.00 (referent)		1.00 (referent)	
75-84 years	1.12	0.96-1.30	1.30	0.95-1.77
≥85 years	0.89	0.70-1.12	2.13	1.47-3.09
Sex				
Male	1.00 (referent)	50	1.00 (referent)	
Female	0.90	0.79-1.03	0.93	0.71-1.22
Race/Ethnicity				
White	1.00 (referent)		1.00 (referent)	
Black	1.50	1.17-1.92	1.21	0.76-1.95
Hispanic	1.11	0.84-1.45	0.86	0.49-1.52
Asian	1.03	0.70-1.51	0.96	0.45-2.06
Other	1.24	0.84-1.82	2.01	1.09-3.71
Primary Health Insurance				
Private	1.00 (referent)		1.00 (referent)	
Public	0.86	0.68-1.07	1.40	0.80-2.45
Self-pay	0.78	0.32-1.89	1.78	0.29-10.92
Other	1.30	0.61-2.79	3.11	0.97-9.97
Median household income quartile				
1 (lowest)	1.13	0.92-1.39	1.07	0.71-1.61
2	1.04	0.85-1.26	1.05	0.71-1.55
3	0.98	0.81-1.18	1.02	0.70-1.51
4 (highest)	1.00 (referent)		1.00 (referent)	
Season of Discharge				
First quarter (Jan - Mar)	1.00 (referent)		1.00 (referent)	
Second quarter (Apr - Jun)	1.10	0.91-1.33	1.16	0.79-1.71
Third quarter (Jul - Sep)	0.97	0.80-1.16	0.89	0.60-1.31
Fourth quarter (Oct - Dec)	1.00	0.83-1.21	1.10	0.75-1.61
Hospital Region				
Northeast	1.00 (referent)		1.00 (referent)	

 Table E5. Clinical risk factors for severe anaphylaxis (cardiac arrest/intubation or death) among US older adults – Method I only, 2006-2014 National Inpatient Sample

Midwest	1.03	0.84-1.28	0.76	0.49-1.19
South	0.98	0.82-1.18	0.93	0.64-1.36
West	1.14	0.93-1.40	1.08	0.72-1.63
Hospital Status				
Non-Metropolitan	1.00 (referent)		1.00 (referent)	
Metropolitan, Non-teaching	1.10	0.87-1.41	1.03	0.59-1.80
Metropolitan Teaching	1.40	1.10-1.78	1.05	0.61-1.82
Weekend presentation				
No	1.00 (referent)		1.00 (referent)	
Yes	0.99	0.83-1.17	1.17	0.83-1.65
Type of Trigger			D'	
Food	1.00 (referent)		1.00 (referent)	
Drug	0.90	0.69-1.19	1.56	0.69-3.50
Venom	1.51	0.35-6.45	Not calculated	
Other	1.15	0.90-1.47	2.88	1.38-6.05
Elixhauser Index Score	1.06	1.05-1.07	1.08	1.06-1.10
Any chronic condition				
No	1.00 (referent)		1.00 (referent)	
Yes	1.17	1.03-1.32	1.11	0.85-1.45
Comorbidities* (referent: no for each condition)				
Alcohol	2.64	1.65-4.22	1.81	0.78-4.24
Congestive Heart Failure	0.81	0.66-0.99	0.73	0.50-1.06
Chronic Pulmonary Disease	1.21	1.05-1.40	0.91	0.68-1.22
Diabetes	1.11	0.95-1.29	1.15	0.84-1.56
Hypertension	1.03	0.88-1.19	0.84	0.63-1.11
Liver Disease	0.45	0.24-0.86	0.79	0.28-2.21
Obesity	1.58	1.26-1.97	1.57	1.01-2.46
Oncologic	0.62	0.49-0.78	0.64	0.42-0.98
Renal Failure	0.76	0.63-0.93	0.92	0.64-1.32
Rheumatoid arthritis/collagen vascular diseases	0.94	0.67-1.32	1.36	0.74-2.50

Figure E1. Flow diagram for acute allergic reactions and anaphylaxis-related ED visits and hospitalizations in the U.S. from 2006-2014, National Emergency Department Sample (NEDS) and National Inpatient Sample (NIS)



*We also included an "other" category for anaphylaxis-related visits which included visits with multiple triggers or those visits that were not included in the above categories.

Figure E2. Rate of emergency department visits for acute allergic reactions per 100,000 population among all adults (≥18 years) in the US, 2006-2014 National Emergency Department Sample

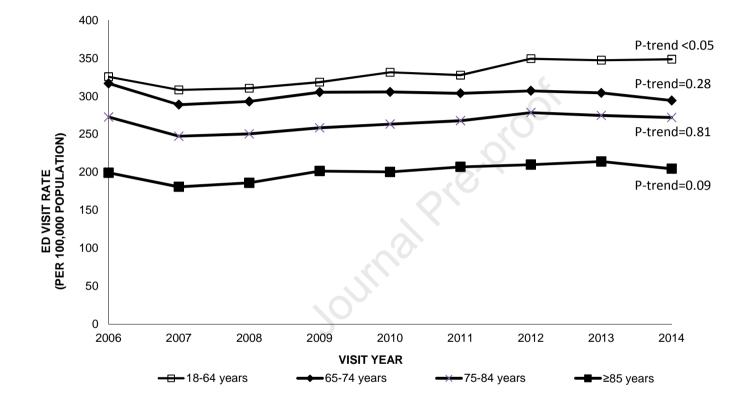
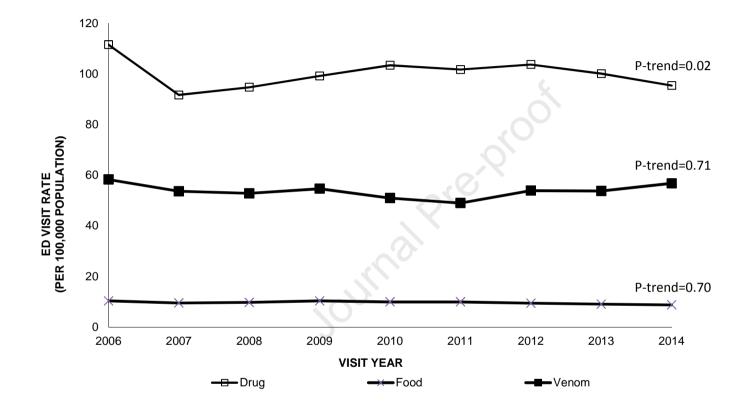


Figure E3. Rate of emergency department visits for acute allergic reactions per 100,000 population among older adults (≥65 years) in the US, 2006-2014 National Emergency Department Sample



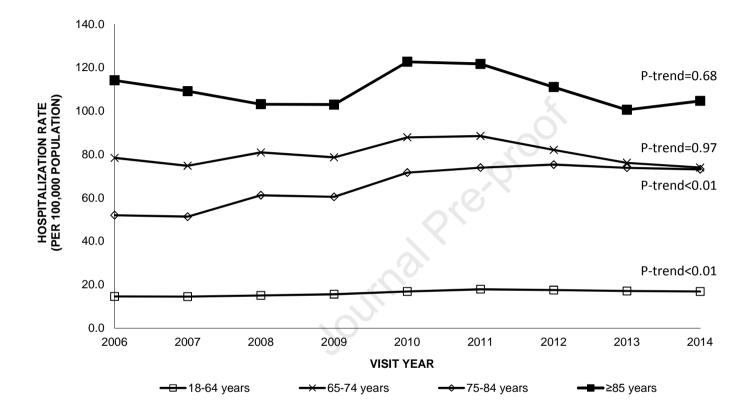


Figure E4. Rate of hospitalizations per 100,000 population for acute allergic reactions among all adults (≥18 years) in the US, 2006-2014 National Inpatient Sample

