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# Trends in inpatient admissions and emergency department visits for heart failure in adults with versus without diabetes in the USA, 2006–2017

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#### ABSTRACT

**Introduction** Heart failure (HF) is a major contributor to cardiovascular morbidity and mortality in people with diabetes. In this study, we estimated trends in the incidence of HF inpatient admissions and emergency department (ED) visits by diabetes status.

**Research design and methods** Population-based agestandardized HF rates in adults with and without diabetes were estimated from the 2006–2017 National Inpatient Sample, Nationwide ED Sample and year-matched National Health Interview Survey, and stratified by age and sex. Trends were assessed using Joinpoint.

**Results** HF inpatient admissions did not change in adults with diabetes between 2006 and 2013 (from 53.9 to 50.4 per 1000 persons; annual percent change (APC): -0.3 (95% Cl -2.5 to 1.9) but increased from 50.4 to 62.3 between 2013 and 2017 (APC: 4.8 (95% Cl 0.3 to 9.6)). In adults without diabetes, inpatient admissions initially declined (from 14.8 in 2006 to 12.9 in 2014; APC -2.3 (95% Cl -3.2 to -1.2)) and then plateaued. Patterns were similar in men and women, but relative increases were greatest in young adults with diabetes. HF-related ED visits increased overall, in men and women, and in all age groups, but increases were greater in adults with (vs without) diabetes.

**Conclusions** Causes of increased HF rates in hospital settings are unknown, and more detailed data are needed to investigate the aetiology and determine prevention strategies, particularly among adults with diabetes and especially young adults with diabetes.

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#### **INTRODUCTION**

People with diabetes are at increased risk for cardiovascular disease (CVD) and associated complications.<sup>1</sup> Although diabetes has become an increasingly common disease, estimated to affect 463 million people worldwide<sup>2</sup> and more than 34 million in the USA,<sup>3</sup> CVD and related mortality in people with diabetes has fallen dramatically in most high-income countries since the 1980s likely due to advances in treatment and better management of risk factors.<sup>4 5</sup> However, the

#### SIGNIFICANCE OF THIS STUDY

## WHAT IS ALREADY KNOWN ABOUT THIS SUBJECT?

- ⇒ Heart failure (HF) is a major contributor to cardiovascular morbidity and mortality in people with diabetes.
- ⇒ Whether HF-related hospitalizations among adults with versus without diabetes has changed over time remains unknown.

#### WHAT ARE THE NEW FINDINGS?

- ⇒ Rates of HF-related inpatient admissions and ED visits are three to five times higher in adults with versus without diabetes, and this excess risk has increased over time.
- ⇒ Though absolute rates remain lowest in the youngest age groups, the greatest relative increases in HF-related inpatient admissions and ED visits were observed in young adults with diabetes.
- ⇒ Increases in HF-related utilization among adults with diabetes was observed in both inpatient and ED settings, suggesting broader underlying causes rather than a shift in treatment setting.

#### HOW MIGHT THESE RESULTS CHANGE THE FOCUS OF RESEARCH OR CLINICAL PRACTICE?

- ⇒ Combined with current evidence from clinical trials, findings of this study support the use of intensive and focused prevention and management of diabetes, including the use of SGL2 inhibitors, to reduce the incidence of HF hospitalizations in people with diabetes.
- ⇒ Future research should focus on the drivers of increases in HF hospitalizations, especially among young people with diabetes.

reported declines in CVD among people with diabetes (both incidence and mortality) often do not include heart failure (HF) as an outcome, despite increasing recognition that HF is a major contributor to CVD

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In a 2015 paper, Shah *et al*<sup> $\theta$ </sup> demonstrated that HF is more likely to be an initial manifestation of CVD in people with type 2 diabetes compared with myocardial infarction, stroke and coronary disease. Despite the relative importance of HF in diabetes, few studies have comprehensively examined whether rates of HF in people with diabetes (vs without diabetes) has changed over time. In the USA, one recent study demonstrated that HF inpatient admissions, defined as the primary reason for hospital admission, increased 3.6% per year between 2013 and 2015 following a period of decline.<sup>12</sup> However, to understand the underlying drivers of changes in HF rates and develop subsequent interventions, a comparison with people without diabetes is needed. Such comparisons in atherosclerotic CVD (eg, myocardial infarction and coronary artery disease) have led to narrowing the gap by reducing the excess risk in diabetes populations.<sup>13 I4</sup> Furthermore, a more comprehensive approach to understanding the overall HF burden is necessary to inform healthcare planning and resource allocation. This includes consideration of multiple settings in which HF care is likely to occur, as well as consideration of HF as both a primary and contributory cause for hospitalization.

Using nationally representative USA data, we estimated secular trends in the incidence of HF-related inpatient admissions and ED visits among adults with diabetes versus adults without diabetes between 2006 and 2017.

#### **METHODOLOGY**

#### The National Inpatient Sample (NIS) and the Nationwide Emergency Department Sample (NEDS)

We analyzed annual data (2006–2017) from the Agency for Healthcare Research and Quality's NIS and NEDS.<sup>15</sup> NIS and NEDS, the largest all-payer inpatient and ED databases in the USA, includes 7 million and 30 million unweighted annual visits, respectively.<sup>15</sup> Both data sets approximate a 20% stratified sample of discharges and can be weighted to provide nationally representative estimates. Rehabilitation and long-term acute care hospitals are excluded from NIS. Both NIS and NEDS include International Classification of Diseases Clinical Modification (ICD-CM) diagnostic codes as well as patient demographics, hospital characteristics, payment sources, patient disposition and total charges. Both NIS and NEDS data represent hospital discharges, not individual persons, and therefore our analysis does not account for multiple admissions per person.

A hospitalization was considered to be related to HF if at least one ICD-9-CM diagnosis code 428.x between January 2000 and September 2015, or ICD-10-CM diagnosis code I50.x between October 2015 and December 2017, appeared in NIS or NEDS data. This approach is aimed to better capture the overall burden of HF by including HF listed as the primary or contributory cause of the hospitalization. In a sensitivity analysis, we defined

HF as the primary cause of hospital admission in NIS and NEDS between January 2006 and September 2015. This analysis was restricted to September 2015 and earlier due to known coding changes implemented in October 2015 that impacted the likelihood of HF being listed as the primary cause of hospital admissions in later years.<sup>16</sup> The 2015 population data (from National Health Interview Survey (NHIS)) were weighted by 0.75 to reflect that only three-quarters of the numerator data was used.<sup>17</sup> To avoid double-counting, we excluded ED visits where the disposition was admission to the hospital because these HF events were accounted for in the inpatient data. Each HF-related admission was considered to be related to diabetes if any of the listed diagnoses also included a diabetes code (ICD-9-CM: 250 .x, 357.2, 366.41; ICD-10-CM: E10, E11 and E13). Comorbidities, adapted from the Charlson Comorbidity Index, among hospitalized patients with HF and with or without diabetes were defined using ICD-9-CM and ICD-10-CM, as appropriate (see online supplemental table 1).

#### **The National Health Interview Survey**

Using annual data (2006–2017) from the NHIS, we estimated the number of persons aged  $\geq 18$  years with and without diabetes.<sup>18</sup> The NHIS is a household-based survey of the health of the civilian, non-institutionalized USA population.<sup>18</sup> We defined adults with diabetes if the sample adult responded yes to the question, 'other than during pregnancy, have you ever been told by a doctor or other health professional that you have diabetes or sugar diabetes?'. This survey does not distinguish between diabetes types; but given that type 2 diabetes accounts for 90%–95% of all diabetes cases,<sup>19</sup> we consider the results of this study to be generalizable to people with diagnosed type 2 diabetes. Data from the NHIS were weighted to make estimates representative of the demographic characteristics of the US civilian non-institutionalized population.

#### **Statistical analysis**

We reported the crude weighted number of patients with HF at the start (2006), middle (2011) and end (2017) of the study period, stratified by diabetes status, and age group, sex, location (urban, micropolitan and rural), household income (quartiles), USA region (northeast, midwest, south and west) and comorbidities for both NIS and NEDS. The weighted results estimate the number of inpatient admissions and non-admission ED visits in the USA due to HF.

Annual rates were calculated as the number of HF hospitalizations with and without diabetes (as determined from NIS and NEDS), divided by the number of persons with and without diabetes (as determined from NHIS). We reported age-standardized rates of HF per 1000 adults with diabetes and per 1000 adults without diabetes. Age (grouped into 18–44, 45–64, 65–74 and ≥75 years) and sex-specific rates were also calculated. Rates were age standardized using the 2000 USA standard population. Excess risk between diabetes and non-diabetes populations was estimated as rate ratios

		Emergen	cy depan	tment (NED	3)**			Inpatient (	(SIN				
	Diabetes	2006		2011		2017		2006		2011		2017	
	status	z	%	z	%	z	%	z	%	z	%	z	%
Weighted total, N		863 816		1 215 499		2 291 311		4 199 309		4 328 315		5 076 844	
Demographic characteristics	(0												
Age group (years)													
18-44	Yes	16 745	5.7	27 349	5.9	56 166	5.8	44 483	2.8	50 329	2.7	68 740	2.9
	No	47 556	8.3	65 266	8.7	120 347	9.1	102 984	3.9	96 885	3.9	124 950	4.6
45-64	Yes	97 036	33.3	164 710	35.5	360 344	37	414 267	26.4	507 741	27.1	672 170	28.2
	No	134 910	23.6	202 597	27	397 474	30.2	468 952	17.8	469 112	19.1	581 875	21.6
65-74	Yes	70 292	24.1	111 663	24.1	252 654	25.9	412 325	26.2	499 910	26.7	690 205	29
	No	95 300	16.7	122 115	16.2	240 782	18.3	479 143	18.2	435 137	17.7	556 425	20.7
≥75	Yes	107 552	36.9	160 043	34.5	305 444	31.3	701 007	44.6	815 249	43.5	952 585	40
	No	294 425	51.5	361 756	48.1	558 100	42.4	1 576 148	60	1 453 952	59.2	1 429 894	53.1
Gender													
Men	Yes	122 793	42.1	205 775	44.4	457 279	46.9	724 625	46.1	908 406	48.5	1 227 515	51.5
	No	249 204	43.6	329 884	43.9	621 073	47.2	1 213 043	46.2	1 143 371	46.6	1 333 365	49.5
Women	Yes	168 800	57.9	257 662	55.6	517 247	53.1	847 414	53.9	964 767	51.5	1 156 150	48.5
	No	322 796	56.4	421 466	56.1	695 466	52.8	1 414 070	53.8	1 311 565	53.4	1 359 705	50.5
Location													
Urban	Yes	217 555	74.8	357 555	77.2	790 845	81.3	1 234 506	78.7	1 469 309	79.9	1 955 570	82.3
	No	422 590	74.1	571 866	76.3	1 054 555	80.4	2 033 634	77.6	1 905 172	79.5	2 193 210	81.7
Micropolitan	Yes	43 821	15.1	63 141	13.6	106 273	10.9	191 267	12.2	210 987	11.5	235 854	9.9
	No	81 791	14.3	101 643	13.6	146 877	11.2	333 365	12.7	276 922	11.6	271 774	10.1
Rural	Yes	29 566	10.2	42 292	9.1	75 427	7.8	143 331	9.1	159 368	8.7	185 925	7.8
	No	65 664	11.5	76 464	10.2	110 611	8.4	254 262	9.7	213 847	8.9	218 230	8.1
Household income													
First quartile (poorest)	Yes	102 414	35.8	172 413	37.9	383 475	40	515 942	33.6	624 720	33.9	826 755	35.2
	No	187 940	33.6	258 795	35.1	470 234	36.4	773 507	30.1	730 924	30.3	819 735	31
Second quartile	Yes	85 225	29.8	125 087	27.5	268 279	28	406 474	26.5	466 148	25.3	639 255	27.2
	No	160 980	28.8	198 215	26.9	355 320	27.5	670 920	26.1	604 367	25.1	714 885	27
Third quartile	Yes	63 322	22.2	98 071	21.5	186 367	19.4	340 853	22.2	447 474	24.3	514 840	21.9
	No	125 866	22.5	165 483	22.4	268 241	20.7	598 751	23.3	593 410	24.6	608 900	23
												5	Continued

## Epidemiology/Health services research

Table 1 Continued													
		Emergend	sy depar	tment (NED	s)**			Inpatient ()	AIS)				
	Diabetes	2006		2011		2017		2006		2011		2017	
	status	Z	%	Z	%	z	%	z	%	Z	%	N	%
Fourth quartile (wealthiest)	Yes	34 886	12.2	59 563	13.1	121 124	12.6	272 696	17.8	302 516	16.4	365 980	15.6
	No	85 056	15.2	114 868	15.6	199 544	15.4	527 656	20.5	480 756	20	504 725	19.1
US region													
Northeast	Yes	33 609	11.5	46 079	9.9	86 556	8.9	320 429	20.4	354 641	18.9	426 350	17.9
	No	81 856	14.3	87 542	11.6	144 018	10.9	501 525	19.1	507 939	20.7	513 079	19.1
Midwest	Yes	85 067	29.2	122 662	26.4	242 622	24.9	393 679	25	474 545	25.3	583 045	24.5
	No	156 589	27.4	193 037	25.7	328 833	25	658 381	25.1	611 503	24.9	644 774	23.9
South	Yes	129 239	44.3	211 335	45.6	462 321	47.4	618 281	39.3	733 689	39.2	971 856	40.8
	No	234 413	41	322 829	42.9	578 959	44	1 061 978	40.4	938 078	38.2	1 059 016	39.3
West	Yes	43 709	15	83 689	18	183 109	18.8	239 693	15.2	310 355	16.6	402 449	16.9
	No	99 333	17.4	148 326	19.7	264 894	20.1	405 343	15.4	397 565	16.2	476 274	17.7
Comorbidities													
Myocardial Infarction	Yes	247 859	16.6	418 179	15.3	629 091	21.1	272 138	17.3	396 389	21.2	568 300	23.8
	No	383 379	15.1	300 754	16.5	619 893	17.5	433 155	16.5	455 611	18.6	548 890	20.4
Peripheral Vascular Disease	Yes	144 798	9.7	218 434	8	440 161	14.8	195 980	12.5	318 289	17	565 005	23.7
	No	166 635	6.6	186 437	10.2	522 821	14.7	216 342	8.2	285 756	11.6	501 535	18.6
Cerebrovascular Disease	Yes	98 752	6.6	187 399	6.8	123 358	4.1	120 054	7.6	184 883	9.9	133 380	5.6
	No	163 958	6.5	125 599	6.9	139 687	3.9	197 503	7.5	228 489	9.3	150 715	5.6
Dementia	Yes	99 882	6.7	299 327	10.9	263 721	8.9	105 888	6.7	177 371	9.5	248 990	10.4
	No	241 362	9.5	140 117	7.7	391 736	1	251 099	9.6	322 289	13.1	362 215	13.4
Chronic Pulmonary Disease	Yes	532 654	35.7	987 082	36	1 178 523	39.6	574 059	36.5	740773	39.5	972 615	40.8
	No	921 924	36.3	672 799	36.9	1 349 228	38.1	1 016 015	38.7	951 659	38.8	1 101 715	40.9
Rheumatic Disease	Yes	27 135	1.8	86 688	3.2	78 835	2.6	30 915	0	53 319	2.8	70 870	ო
	No	64 962	2.6	39 071	2.1	128 802	3.6	76 293	2.9	100 400	4.1	119 205	4.4
Peptic Ulcer Disease	Yes	18 072	1.2	35 223	1.3	33 329	1.1	21 700	1.4	29 251	1.6	35 165	1.5
	No	36 891	1.5	19 024	-	42 069	1.2	44 371	1.7	42 480	1.7	45 205	1.7
Liver Disease Mild	Yes	34 623	2.3	80 354	2.9	125 905	4.2	39 761	2.5	78 917	4.2	119 895	5
	No	67 995	2.7	50 306	2.8	128 975	3.6	77 314	2.9	102 382	4.2	123 860	4.6
Renal Disease Mild to	Yes	244 008	16.3	576 139	21	1 076 815	36.2	281 640	17.9	717 912	38.3	1 039 750	43.6
Moderate	No	263 803	10.4	551 260	30.3	855 003	24.1	308 784	11.8	657 730	26.8	837 925	31.1
												0	ontinued

		Emergend	cy depar	tment (NEDS	**(			Inpatient (N	lis)				
	Diabetes	2006		2011		2017		2006		2011		2017	
	status	N	%	Z	%	N	%	Z	%	Z	%	N	%
Hemiplegia or Paraplegia	Yes	10 333	0.7	30 742	1.1	37 755	1.3	12 234	0.8	23 801	1.3	43 430	1.8
	No	22 070	0.9	15 487	0.8	46 474	1.3	25 058	-	37 340	1.5	54 355	2
Any malignancy	Yes	51 584	3.5	139 644	5.1	111 149	3.7	68 008	4.3	96 629	5.2	119 180	5
	No	125 689	4.9	66 860	3.7	168 915	4.8	171 192	6.5	172 419	7	187 905	7
Liver Disease moderate to	Yes	7 801	0.5	20 780	0.8	34 597	1.2	10 484	0.7	21739	1.2	37 535	1.6
severe	No	14 861	0.6	13 485	0.7	33 431	0.9	17 231	0.7	25 950	<del>.</del> .	37 905	1.4
Renal disease severe	Yes	328 819	22	164 424	9	418 838	14.1	371 670	23.6	238 246	12.7	335 555	14.1
	No	356 566	14	197 091	10.8	212 217	9	407 246	15.5	154 810	6.3	164 155	6.1
HIV/AIDS	Yes	1 971	0.1	9 117	0.3	7 158	0.2	2 378	0.2	3 267	0.2	5 435	0.2
	No	8 966	0.4	3 013	0.2	16 791	0.5	9 239	0.4	8 588	0.3	11 295	0.4
Metastatic solid tumor	Yes	15 407	-	40 665	1.5	36 266	1.2	20 809	1.3	25 695	1.4	39 840	1.7
	No	42 913	1.7	17 100	0.9	58 244	1.6	59 113	2.2	51 215	2.1	67 090	2.5
*HF defined as any diagnosis; **ex HF heart failure NEDS Nationwid	cludes those v e Emergency I	who were ad	mitted to Sample: N	hospital. IIS National Inr	atient S	alum							

(RRs). We used SAS-callable SUDAAN (RTI International) to account for the complex sampling design in NIS, NEDS and NHIS, and the Taylor series linearization was used to estimate the variance of the ratio of the numerator and denominator. The delta method was used to compute SEs and 95% CIs for rates and RRs accounting for the weighted design of NIS, NEDS and NHIS.<sup>20</sup>

In 2012, the NIS sampling design was changed, which has implications for trend analyses. Per NIS guidelines, we used NIS-provided trend weights for the years preceding 2012 and the discharge weights beginning in 2012 to make the discharge outcome consistent with the new sampling design.<sup>21</sup>

Joinpoint regression was used to examine trends over time.<sup>22</sup> This software uses permutation tests to identify points where linear trends change significantly in either direction or magnitude and calculates an annual percentage change (APC) for each time period identified. A maximum of two joinpoints were specified. A p value of <0.05 was established as statistical significance.

#### **RESULTS**

Characteristics of adults with HF in inpatient (NIS) and ED (NEDS) settings in 2006, 2011 and 2017, and by diabetes status, are described in table 1. In brief, among HF-related inpatient admissions and ED visits between 2006 and 2017, there was an increase in the proportion of men, middle-age adults (aged 45-64 and 65-74 years), adults residing in urban settings, adults reporting low-income households and adults living in the West. This was broadly true for people with and without diabetes. In addition, the proportion of HF hospitalizations, both in ED and inpatient settings, with comorbidities increased in people with and without diabetes with a few exceptions: HF-related hospitalizations with cerebrovascular disease and severe renal disease decreased over time in people with and without diabetes, and the proportion of HF-related hospitalizations with peptic ulcer disease, HIV, malignancy or metastatic solid tumor did not change over time in people with or without diabetes. The increasing proportion of most comorbidities was, generally, higher in people with as compared without diabetes.

#### National Inpatient Sample

In 2017, rates of HF-related inpatient admissions were more than five times as high in adults with versus without diabetes (RR: 5.1 (95% CI 4.7 to 5.5)), a significant increase from 3.6 (95% CI 3.3 to 4.0) in 2006 (table 2). Overall, between 2006 and 2013, rates of HF-related inpatient admissions did not change among adults with diabetes, and then increased sharply between 2013 and 2017 from 50.4 to 62.3 per 1000 persons (APC: 4.8 (95% CI 0.3 to 9.6) (figure 1A and table 2). Among adults without diabetes, the opposite was observed: between 2006 and 2014, rates declined from 14.8 to 11.7 (APC -2.3 (95% CI -3.2 to -1.2) and plateaued thereafter. Similar patterns were observed in both men and women.

By age, differences were noted (figure 2A and table 2). First, the excess risk associated with diabetes decreased with increasing age. For example, the 2017 RR was 20.2 (95% CI 16.9 to 23.5) versus 2.8 (95% CI 2.5 to 3.2) for those aged 18–44 and ≥75 years, respectively. Second, in adults aged 18-44 years with and without diabetes, rates of HF-related inpatient admissions increased similarly such that there was no significant change in the excess risk associated with diabetes over time. Third, among adults aged 45-64 and 65-74 years with and without diabetes, HF rates increased after a period of decline and the excess risk associated with diabetes increased. Last, among adults aged ≥75 years, rates of HF-related inpatient admissions declined throughout the study period in adults without, but not with, diabetes and the excess risk associated with diabetes increased (from RR of 2.0 to 2.8; APC 2.4 (95% CI 0.6 to 4.2)).

#### **Nationwide Emergency Department Sample**

In 2017, rates of HF-related ED visits were more than five times as high in adults with versus without diabetes (RR: 5.2 (95% CI 4.5 to 5.9)), a significant increase from 3.7 (95% CI 3.2 to 4.1) in 2006 (table 3). Overall, between 2006 and 2017, rates of HF-related ED visits increased in adults with (from 11.5 to 43.6 per 1000 persons) and without (from 3.1 to 5.9 per 1000 persons) diabetes (figure 1B and table 3). However, the rate of increase was greater in adults with diabetes, leading to an increase in the excess risk of HF-related ED visits associated with diabetes over.

Increases in HF-related ED visits were observed across all age groups and in adults with and without diabetes (figure 2B and table 3). For all age groups, excluding 65–74 years, the excess risk associated with diabetes did not significantly change over time, indicating increasing rates of HF ED visits were similar in adults with and without diabetes. However, among adults aged 65–74 years, the HF rate increase was greater in adults with diabetes, leading to an increase in the excess risk associated with diabetes over time (from RR of 3.3 to 4.4; APC 2.0 (95% CI 0.7 to 3.3)).

#### Sensitivity analyses

In a sensitivity analysis, we examined trends in HF inpatient admissions and ED visits between 2006 and 2015 where HF was defined as the primary reason for the admission (online supplemental tables 2 and 3). Overall, in 2015 rates of primary inpatient HF admissions and HF ED visits were 4.7 (95% CI 4.4 to 5.1) and 3.2 (95% CI 2.8 to 3.5) times as high in adults with versus without diabetes, respectively.

Though absolute rates were substantially lower when HF was defined as the primary (vs any) reason for admission, inpatient patterns were similar insofar as the excess risk associated with diabetes, particularly among younger adults, increased over time (online supplemental table 2).

		Age-standardized inpa	atient admission rate (pe	er 1000 people) (95% CI)-	Trend 1		Trend 2‡	
		2006	2011	2017	Year	APC (95% CI)	Year	APC (95% CI)
Total	Diabetes	53.9 (49.4 to 58.3)	54.6 (50.9 to 58.4)	62.3 (58.2 to 66.4)	2006-2013	-0.3 (-2.5 to 1.9)	2013-2017	4.8 (0.3 to 9.6)§
population	No diabetes	14.8 (13.7 to 15.8)	12.9 (12.2 to 13.6)	12.2 (11.6 to 12.9)	2006-2014	-2.3 (-3.2 to -1.2)§	2014-2017	1.4 (-2.6 to 5.6)
	RR	3.6 (3.3 to 4.0)	4.2 (3.9 to 4.6)	5.1 (4.7 to 5.5)	2006-2017	2.8 (1.9 to 3.7)§	n/a	
Sex								
Men	Diabetes	54.6 (48.9 to 60.3)	54.8 (50.2 to 59.4)	62.1 (57.2 to 67.1)	2006-2009	-4.7 (-13.4 to 4.9)	2009-2017	3.3 (1.6 to 4.9)§
	No diabetes	16.1 (14.8 to 17.4)	13.9 (13.0 to 14.9)	13.7 (12.9 to 14.6)	2006-2013	-2.4 (-3.7 to -1.2)§	2013-2017	1.2 (-1.3 to 3.7)
	RR	3.4 (2.9 to 3.8)	3.9 (3.5 to 4.4)	4.5 (4.0 to 5.0)	2006-2017	3.1 (1.9 to 4.3)§	n/a	
Women	Diabetes	53.0 (48.0 to 58.1)	54.6 (50.0 to 59.2)	62.5 (57.5 to 67.6)	2006-2013	-1.1 (-3.4 to 1.3)	2013-2017	5.2 (0.2 to 10.5)§
	No diabetes	13.7 (12.6 to 14.7)	12.0 (11.3 to 12.8)	11.0 (10.4 to 11.6)	2006-2017	-1.7 (-2.4 to -1.1)§	n/a	
	RR	3.9 (3.4 to 4.4)	4.5 (4.1 to 5.0)	5.7 (5.1 to 6.3)	2006-2017	2.8 (1.7 to 3.8)§	n/a	
Age group	(years)							
18-44	Diabetes	15.2 (12.8 to 17.7)	18.6 (15.9 to 21.2)	22.8 (19.4 to 26.2)	2006-2017	3.9 (1.9 to 6.0)§	n/a	
	No diabetes	1.0 (0.9 to 1.0)	0.9 (0.8 to 1.0)	1.1 (1.1 to 1.2)	2006-2008	-9.4 (-24.2 to 8.1)	2008-2017	3.1 (1.5 to 4.7)§
	RR	15.9 (13.0 to 18.7)	20.7 (17.2 to 24.3)	20.2 (16.9 to 23.5)	2006-2017	2.2 (-0.1 to 4.4)	n/a	
45-64	Diabetes	53.4 (47.7 to 59.2)	52.3 (47.7 to 57.0)	63.4 (57.8 to 69.0)	2006-2008	-10.3 (-23.5 to 5.1)	2008-2017	3.8 (2.5 to 5.1)§
	No diabetes	7.1 (6.6 to 7.5)	6.6 (6.2 to 7.0)	8.0 (7.5 to 8.4)	2006-2012	-1.3 (-3.0 to 0.4)	2012-2017	4.7 (3.1 to 6.3)§
	RR	7.6 (6.6 to 8.5)	7.9 (7.1 to 8.8)	7.9 (7.1 to 8.8)	2006-2017	0.9 (0.2 to 1.7)§	n/a	
65–74	Diabetes	118.9 (104.1 to 133.6)	102.9 (93.6 to 112.2)	122.8 (11.8 to 133.8)	2006–2015	–1.7 (–2.8 to –0.6)§	2015-2017	11.4 (–0.7 to 24.8)
	No diabetes	30.7 (28.1 to 33.3)	25.6 (23.8 to 27.4)	23.4 (22.0 to 24.8)	2006-2013	-4.2 (-5.3 to -3.0)§	2013-2017	1.3 (-1.0 to 3.6)
	RR	3.9 (3.3 to 4.4)	4.0 (3.6 to 4.5)	5.3 (4.7 to 5.8)	2006-2017	2.0 (0.6 to 3.3)§	n/a	
575	Diabetes	236.4 (204.3 to 268.5)	245.6 (218.8 to 272.4)	249.6 (221.3 to 277.9)	2006-2017	0.1 (-1.4 to 1.6)	n/a	
	No diabetes	115.7 (105.8 to 125.6)	100.6 (93.5 to 107.7)	88.0 (82.2 to 93.8)	2006-2017	-2.2 (-2.8 to -1.7)§	n/a	
	RR	2.0 (1.7 to 2.4)	2.4 (2.1 to 2.8)	2.8 (2.5 to 3.2)	2006-2017	2.4 (0.6 to 4.2)§	n/a	
Data source: *RR is rate in	s: National Center	for Health Statistics, National	Il Health Interview Survey and	I Agency for Healthcare Rese	arch and Quality	, National Inpatient Sam	ıple.	
†Years 2006, †Indicates th	, 2011 and 2017 a	ire displayed in this table for e	ease of presentation. Rates a	nd 95% Cl for other years are	e presented in fiç	jures 1 and 2.		
Sp <sub>trend</sub> <0.05.	oretada chanada	an failure had failure an a	econd trand identified					
	hai nai Itaga ni mi	טר, חר, ווכמו ומוועו כ, ווימ, יוט ט	בכחות וובוות ותבויווובת.					



Figure 1 Age-standardized inpatient admission (NIS (A)) and ED visit (NEDS (B)) rates for HF in people with (i) versus without diabetes (ii) in the USA between 2006 and 2017. HF, heart failure; NEDS, Nationwide Emergency Department Sample; NIS, National Inpatient Sample.

This was driven by continued declines in HF rates among people without diabetes throughout the study period, while HF rates among people with diabetes plateaued from approximately 2010 onwards. For HF-related ED visits defined as the primary cause, the excess risk associated with diabetes also increased over time driven by increases in HF rates among people with, but not without, diabetes in the latter study period (online supplemental table 3).

#### DISCUSSION

In this study, we provide the first comprehensive summary of trends of HF-related inpatient admissions and nonadmitted ED visits in the USA among adults with and without diabetes and note several important findings. First, rates of HF-related inpatient admissions and ED visits were three to five times higher in adults with versus without diabetes, and this excess risk has increased over time. Second, while absolute rates remained lowest in the youngest age groups, the greatest relative increases in HF-related inpatient admissions and ED visits were observed in young adults with diabetes. Third, increases in HF-related utilization among adults with diabetes was observed in both inpatient and ED settings, suggesting broader underlying causes rather than a shift in treatment setting.

Our results are consistent with the few studies that have reported changes in HF incidence over time. In the USA, a NIS-based study reported a 3.6% annual decline in HF inpatient admissions among adults  $\geq$ 35 years with diabetes between 1998 and 2014.<sup>13</sup> This decline was likely driven by significant decreases in the earlier period (ie, 1998–2006) and explains why we, in contrast, observed a non-significant decline in HF-related inpatient admissions from 2006 to 2013. Another study, also using the NIS, reported an overall 38.9% decline in primary HF admissions in people with diabetes between 1995 and 2015.<sup>12</sup> This decline also appeared to be driven by reductions in



Figure 2 Age-specific inpatient admission (NIS (A)) and ED visit (NEDS (B)) rates for HF in people with (i) versus without diabetes (ii) in the USA between 2006 and 2017. HF, heart failure; NEDS, Nationwide Emergency Department Sample; NIS, National Inpatient Sample.

the earlier study period as non-significant increases were observed between 2013 and 2015.<sup>12</sup> In Spain, a significant 5.4% annual increase in HF hospitalizations was observed between 1997 and 2010 in patients with diabetes, broadly similar to findings in the current study.<sup>23</sup> However, the NIS-based studies and the Spanish study did not compare changes in HF incidence in people with versus without diabetes. This comparison is necessary to understand whether diabetes is an underlying cause of changing HF rates and to develop targeted interventions to reduce the HF burden in this subpopulation. Only one other study has compared rates of HF hospitalizations in people with versus without diabetes. In Sweden, a 29% decrease in HF hospitalization rates, defined as primary of contributory cause, among persons with type 2 diabetes was observed between 1999 and 2013, and this decline was greater than what was observed for people without type 2 diabetes.<sup>24</sup> Unfortunately, data beyond 2013 were not available, and thus, it remains to be elucidated whether the recent

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increase in HF hospitalizations seen in our US data is also occurring in other populations and settings.

The increasing rates of HF among people with diabetes, especially young adults with diabetes, are consistent with a recent resurgence of other diabetes-related complications in the USA.<sup>25</sup> Between 2010 and 2015, national data show increases in lower extremity amputations (LEAs)<sup>26</sup> and hyperglycaemic crises among adults with diabetes,<sup>27</sup> while long-term declines in end-stage renal disease, acute myocardial infarction (AMI) and stroke have stalled.<sup>25</sup> These trends appear to be driven by increases in young (aged 18-44 years) and middle-aged (aged 45-64 years) adults, among whom the risk of hyperglycaemic crisis, AMI, stroke and LEAs each increased by more than 25%between 2010 and 2015.<sup>25</sup> We add to this growing body of literature that increases in HF also disproportionally affect young people with diabetes at or around the same time. There are several possible reasons to explain this observed increase. First, we have observed a changing

Table 3 A	ge-standardized	ED visit rates and rate I	ratios (RRs)* for HF in ad	lults with versus withou	t diabetes in the	USA between 2006 a	Ind 2017	
		Age-standardized EC	) visit rate (per 1000 pe	ople) (95% CI)†	Trend 1‡		Trend 2‡	
		2006	2011	2017	Year	APC (95% CI)	Year	APC (95% CI)
Total	Diabetes	11.5 (10.3 to 12.7)	16.4 (14.9 to 17.9)	43.6 (27.3 to 33.9)	2006–2012	6.0 (3.4 to 8.7)§	2012-2017	12.6 (8.9 to 16.5)§
population	No diabetes	3.1 (2.9 to 3.4)	3.8 (3.6 to 4.1)	5.9 (5.4 to 6.4)	2006–2012	4.1 (2.7 to 5.5)§	2012-2017	8.7 (6.1 to 11.3)§
	RR	3.7 (3.2 to 4.1)	4.3 (3.8 to 4.8)	5.2 (4.5 to 5.9)	2006-2017	2.7 (1.9 to 3.5)§	n/a	
Sex								
Men	Diabetes	10.4 (9.2 to 11.7)	14.9 (13.3 to 16.6)	27.1 (23.8 to 30.4)	2006–2010	4.3 (-1.8 to 10.8)	2010-2017	11.8 (9.0 to 14.5)§
	No diabetes	3.2 (2.9 to 3.4)	3.8 (3.5 to 4.1)	6.2 (5.7 to 6.7)	2006-2017	4.1 (1.5 to 6.7)	2012-2017	9.5 (5.6 to 13.5)§
	RR	3.3 (2.8 to 3.8)	3.9 (3.4 to 4.4)	4.4 (3.7 to 5.0)	2006-2017	2.7 (1.5 to 4.0)§	n/a	
Women	Diabetes	12.5 (11.1 to 14.0)	17.9 (16.0 to 19.7)	34.2 (30.1 to 38.4)	2006-2013	6.1 (4.2 to 8.1)§	2013-2017	14.3 (9.1 to 19.6)§
	No diabetes	3.1 (2.8 to 3.4)	3.8 (3.6 to 4.1)	5.6 (5.2 to 6.1)	2006–2012	4.0 (2.6 to 5.4)§	2012-2017	7.8 (5.1 to 10.5)§
	RR	4.0 (3.5 to 4.6)	4.7 (4.1 to 5.2)	6.1 (5.2 to 7.0)	2006-2017	2.7 (1.8 to 3.7)§	n/a	
Age group								
18-44	Diabetes	5.7 (4.6 to 6.8)	10.1 (8.5 to 11.7)	18.6 (15.3 to 22.0)	2006-2012	10.6 (8.5 to 12.8)§	n/a	
	No diabetes	0.4 (0.4 to 0.5)	0.6 (0.5 to 0.7)	1.1 (1.0 to 1.2)	2006-2017	8.9 (7.6, to 10.2)§	n/a	
	RR	12.9 (10.1 to 15.8)	16.7 (13.7 to 19.8)	17.1 (13.4 to 20.8)	2006-2017	1.8 (-0.1 to 3.8)	n/a	
45-64	Diabetes	12.5 (11.0 to 14.1)	17.0 (15.1 to 18.8)	34.0 (29.8 to 38.2)	2006–2012	5.9 (3.6 to 8.3)§	2012-2017	14.7 (11.2 to 18.2)§
	No diabetes	2.0 (1.9 to 2.2)	2.8 (2.6 to 3.1)	5.5 (4.9 to 6.0)	2006–2012	7.0 (5.4 to 8.6)§	2012-2017	12.4 (10.3 to 14.5)§
	RR	6.2 (5.2 to 7.1)	6.0 (5.1 to 6.8)	6.2 (5.2 to 7.2)	2006-2017	0.3 (-0.6 to 1.2)	n/a	
65-74	Diabetes	20.3 (17.5 to 23.0)	23.0 (20.5 to 25.5)	45.0 (39.6 to 50.3)	2006–2014	3.9 (1.8 to 6.1)§	2014–2017	17.3 (7.5 to 28.0)§
	No diabetes	6.1 (5.6 to 6.7)	7.2 (6.6 to 7.8)	10.1 (9.3 to 11.0)	2006–2013	2.7 (0.6 to 4.8)§	2013-2017	9.1 (4.8 to 13.6)§
	RR	3.3 (2.8 to 3.9)	3.2 (2.8 to 3.6)	4.4 (3.8 to 5.1)	2006-2017	2.0 (0.7 to 3.3)§	n/a	
≥75	Diabetes	36.3 (31.0 to 41.5)	48.2 (42.1 to 54.3)	80.0 (69.3 to 90.8)	2006–2013	3.9 (1.0 to 7.0)§	2013-2017	13.4 (6.1 to 21.1)§
	No diabetes	21.6 (19.6 to 23.6)	25.0 (22.9 to 27.2)	34.3 (31.3 to 37.4)	2006-2017	4.5 (3.6 to 5.4)§	n/a	
	RR	1.7 (1.4 to 2.0)	1.9 (1.6 to 2.2)	2.3 (2.0 to 2.7)	2006-2017	1.7 (-0.0 to 3.6)	n/a	
*RR is rate ii †Years 2006 ‡Indicates th \$P <sub>tend</sub> <0.05; APC, annual	, 2011 and 2017 an , 2011 and 2017 an ne year in which trei sample time (from percentage change	<ul> <li>n-diabetes.</li> <li>e displayed in this table fo ads change significantly in RR of 3.7 to 5.2; APC 2.7</li> <li>3; ED, emergency departm</li> </ul>	r ease of presentation. Rate i either direction or magnitu (95% Cl 1.9 to 3.5)). Simila nent; HF, heart failure; n/a, n	is and 95% CI for other ye de. r patterns were observed i o second trend identified.	ars are presented i n both men and wc	n figures 1 and 2. omen.		

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profile of newly identified diabetes cases that are more obese and may have more poorly managed risk factors (eg, blood pressure and lipids) as compared with earlier years, particularly among younger adults.<sup>4</sup> Second, a longer average duration of diabetes may be leading to a shift in risk of complications. Third, the younger age group may include a larger relative proportion of type 1 diabetes who may be at increased risk for HF. However, accumulating evidence suggests that diabetes complication rates may be higher in young adults with type 2 diabetes as compared with type 1 diabetes.<sup>28</sup> Fourth, changes in healthcare policy such as the introduction of high-deductible health plans have led to reductions in early preventive care in people with diabetes.<sup>29 30</sup> Fourth, increased costs of insulin and other diabetes medications may have led patients to cut back on treatment to minimize costs, thus exposing them to increased risk for complications including HF.<sup>31</sup> Last, in 2012, the US Centers for Medicare and Medicaid Services implemented the Hospital Readmissions Reduction Program, which financially penalized hospitals with high 30-day readmission rates for HF.<sup>32</sup> The role of this policy in influencing HF trends in the current study is unclear as NIS and NEDS do not identify hospital readmission. Overall, it is most likely that a combination of these factors explains the increases in HF-related ED visits and hospitalization among US adults with diabetes.

The results of this study offer important implications for public health and healthcare practice. First, in this study, we show that diabetes is associated with an almost fivefold increased risk for HF-related inpatient and non-admission ED visits. The continued increase in the prevalence of diabetes is likely to increase the number of people with HF in the future and will have important implications for both outpatient and hospital burdens, pharmacotherapies and resource allocation. Second, we hypothesize that increasing risk for HF may lead to an increase in subsequent HF-related mortality with some early evidence to support this hypothesis. For example, Cheng *et al*<sup> $\beta$ 3</sup> reported an increase in HF-related mortality among young US adults with diabetes between 1988 and 2015, despite mortality rates for several other CVDs declining in that time, and an Australian study reported no change in HF-related mortality despite declines for other CVD outcomes.<sup>34</sup> Third, improved awareness by healthcare providers that diabetes is an important risk factor for HF might stimulate more intensive and focused prevention and management opportunities. For example, post hoc analysis of the Steno-2 trial in Denmark demonstrated a reduction in HF hospitalizations among patients with diabetes receiving intensive (vs conventional) therapy.<sup>35</sup> Furthermore, emerging trial data of sodium-glucose cotransporters 2 (SGLT2) inhibitors show promising findings for HF. For example, randomized trials of SGLT2 inhibitors (vs placebo) have shown a pooled 31% reduction in HF hospitalizations in type 2 diabetes patients at high risk of CVD,<sup>36</sup> as well as improved outcomes among those with existing diabetes and HF.<sup>37</sup> Real-world studies, such as CVD-REAL (Comparative Effectiveness of Cardiovascular Outcomes in New Users of SGLT-2 Inhibitors), have also demonstrated the positive effects of SGLT-2 inhibitors in HF prevention in patients with type 2 diabetes, irrespective of atherosclerotic disease status.<sup>38,39</sup>

This is the largest study to explore rates of HF over time in USA adults with and without diabetes in two nationally representative patient datasets. Nonetheless, there are limitations to be considered. First, NIS and NEDS represent hospital discharges, not individual persons and therefore may include multiple hospital stays for some persons. This may lead to an increase in population-based rates, especially in certain subpopulations at higher risk for recurrence, including those with diabetes.<sup>40</sup> However, the primary objective of this study was to examine changes in HF admissions over time in people with versus without diabetes. To that end, and in the absence of contrary data, we assume that the risk of readmission in people with versus without diabetes remained constant during the study period and readmissions are, therefore, unlikely to impact our key conclusions. Second, because of the inability to differentiate diabetes type in the NHIS survey data, we were not able to report trends in HF by diabetes type. Therefore, all types of diabetes are included in the current analysis with the assumption that the vast majority ( $\sim 90\% - 95\%$ ) have type 2 diabetes.<sup>41</sup> In addition, the NHIS is self-reported and does not include undiagnosed diabetes and thus likely underestimates the number of people with diabetes in the population. Furthermore, the underlying characteristics of people with diagnosed diabetes could be changing over time. However, there have not been adequate data or studies to characterize such changes. Third, a shift from ICD-9-CM to ICD-10-CM in October 2015 may have affected our observed rates. However, observed changes in trends occurred before this period, and therefore, it is unlikely that this coding shift influenced the overall patterns that we observed in this study. Furthermore, coding changes do not explain differential increases in people with versus without diabetes and in younger versus older adults. Fourth, admissions for hypertensive heart disease with HF were not included in the current analysis. Fifth, NIS and NEDS do not report HF stages and we were unable to explore differential impacts of diabetes on HF stages, though this is an important future direction. Sixth, location (urban/ rural) and poverty status, although available in NHIS, were not categorized in the same way in NEDS and NIS, so these factors were excluded from rate calculations. In addition, the race/ethnicity variable in NIS was incomplete prior to 2012, and so trends were not calculated by race/ethnicity. Finally, this is a descriptive observational study designed to assess the relative burden of HF hospitalizations in people with versus without diabetes over time. Future studies with more appropriate datasets (ie, with individual level data) are needed to tease out the underlying mechanisms with which diabetes leads to an increase in HF hospitalization, particularly among young adults.

#### **CONCLUSIONS**

In this nationally representative study, we show that: (1) rates of HF-related inpatient admissions increased in adults with, but not without, diabetes and (2) rates of HF-related ED visits increased in adults with and without diabetes, but absolute and relative increases were greater in adults with diabetes; and (3) the greatest relative increases in HF-related inpatient admissions and non-admission ED visits was seen among young adults with diabetes. More detailed and subnational data analyses may help to investigate the aetiology and determine clinical and public health strategies to address these growing burdens.

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