

Diagnosed Prevalence and Incidence of Vitiligo in the United States: Analysis of Employer-Sponsored Insurance Claims

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TO THE EDITOR

Vitiligo is an autoimmune disease characterized by areas of skin depigmentation caused by the immune system targeting melanocytes (Ezzedine et al., 2015). The onset of the disease can occur at any age and is often associated with adverse psychosocial effects as well as comorbid autoimmune conditions (Ahmed et al., 2013; Dahir and Thomsen, 2018; Gill et al., 2016; Nguyen et al., 2016; Rzepecki et al., 2018; Sheth et al., 2013; Wu and Cohen, 2019). Estimates of the point prevalence of vitiligo vary across published sources, with a recent analysis of insurance claims reporting a diagnosed prevalence of 0.05%, whereas a meta-analysis gave estimates of 0.2% in population-based studies and 1.8% in hospital-based studies (Lim et al., 2017; Zhang et al., 2016). Published data on the incidence of vitiligo are scant. Given this variation in reported prevalence and the paucity of data on incidence, we conducted a retrospective, observational study using the IBM MarketScan Commercial and Medicare Supplemental database to estimate the annual diagnosed prevalence and incidence of vitiligo among the employer-sponsored insurance (ESI) population in the United States of America (US). Annual incidence and prevalence trends were compared over 5 years. The MarketScan Commercial Claims and Encounters database contains de-identified data from active employees, early retirees, COBRA continuers, and dependents insured by employer-sponsored plans (i.e., persons eligible for Medicare). not The Medicare Supplemental database contains Medicare-eligible retirees with commercially available Medicare Advantage supplemental plans.

Our analysis identified prevalent cases of vitiligo as patients with ≥ 1 inpatient or outpatient claim for vitiligo (International Classification of Disease–Clinical Modification 9/10: 709.01, L80, or H02.731-H02.739) between January 1, 2013, and December 31, 2017. Continuous health plan enrollment was assessed for each year, and patients with a claim for vitiligo from previous calendar years were carried over to the year of analysis if continuous enrollment criteria were met. This cohort of patients served as the numerator for calculating annual diagnosed prevalence referred to as prevalence hereon. The denominator for calculating the annual diagnosed prevalence was the total number of insured members with continuous enrollment in each calendar year. Diagnosed incident cases of vitiligo, referred to as incidence hereon, were identified as patients with ≥ 1 inpatient or outpatient claim for vitiligo during the same time period used to calculate prevalence and no evidence of vitiligo in the previous 12 months. The denominator for calculating the annual incidence was the total number of insured members with continuous enrollment in each calendar year and no evidence of vitiligo in the 12 months before the first claim for vitiligo. Estimates of annual prevalence and incidence were calculated for the overall cohort as well as for adult (aged \geq 18 years) and pediatric (aged <18 years) patients. A sensitivity analysis of both prevalence and incidence was

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conducted to account for the potential misdiagnosis of vitiligo. It used a more restricted criteria for vitiligo that included patients with ≥ 1 inpatient claim or ≥ 2 outpatient claims for vitiligo. Among patients with ≥ 2 outpatient claims for vitiligo, the first outpatient claim was required to be a diagnosis for vitiligo. The second outpatient claim could have been (i) a diagnosis of vitiligo that was ≥ 30 days apart from the first vitiligo diagnosis claim or (ii) a phototherapy procedure (Current Procedural Terminology codes: 96912, 96900, 96910) that may have occurred within \leq 30 days of the first vitiligo diagnosis claim. All estimates were weighted to 2018 national estimates of the US ESI population using MarketScan Commercial Insurance weights that were constructed using the public-use microdata sample of the American Community Survey by the US Census Bureau (US Census Bureau, 2020). The proportion of patients with a first claim for vitiligo diagnosis from a primary care provider or dermatologist was also investigated. All statistical analyses were conducted using SAS v9.4 software (Cary, NC). All study data were de-identified and fully compliant with Health Insurance Portability and Accountability Act Regulations (45 Code of Federal Regulations 164.514e); therefore, approval from an institutional review board was not required, and informed consent was not obtained.

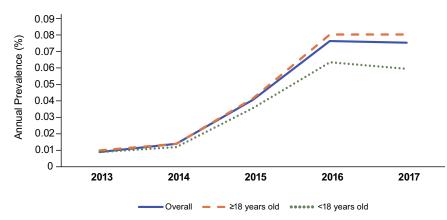
Our results showed that there were 107,612 patients with vitiligo in the prevalence cohort in 2017, up from 11,239 patients in 2013, with annual prevalence increasing from 0.009% to 0.076% during the 5-year observation period (Figure 1 and Table 1). Stratification into adult and pediatric patients showed similar trends to the overall cohort. In 2017, there were 85,821 newly diagnosed cases of vitiligo in the incidence cohort, which represented an increase in annual incidence from 0.007% to 0.061% during the 5-year

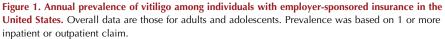
Abbreviations: ESI, employer-sponsored insurance; US, United States of America

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observation period (Figure 2 and Table 2). Similar to prevalence, stratification into adult and pediatric patients showed similar trends to the overall cohort. Results from the sensitivity analysis were consistent with the primary analysis in showing an increase in prevalence and incidence over time for the overall cohort as well as for the adult and pediatric subgroups. Approximately 50% of the first claims for vitiligo diagnosis were initiated by a dermatologist in each of the 5 years from 2013 to 2017, and approximately 10% were from primary care physicians.

Although healthcare claims are commonly used to estimate disease burden, including prevalence and incidence, our retrospective, observational analysis has some limitations that are well recognized with the use of claims data for such estimation (Broder et al., 2017; Konrad et al., 2019). First, insurance claim data are collected for payment purposes and may have inherent limitations when used for research purposes, such as International Classification of Disease-Clinical Modification codes not allowing the identification of segmental or nonsegmental subtypes of vitiligo. The International Classification of Disease-Clinical Modification 9/10 codes of vitiligo assume that the clinician followed formal diagnostic criteria. If the formal diagnostic criteria are not met, a misdiagnosis could result. The other International Classification of Disease-Clinical Modification codes used do not include other areas of the body that vitiligo commonly affects, such as the neck, hands, and skin creases. This methodology for identifying subjects with vitiligo has not been validated. Second, our results may underestimate the real-world prevalence and incidence of vitiligo due to the minimal or absent healthcare-seeking behavior of some patients with this condition, which may in turn be influenced by the absence of approved pharmacologic treatments and restrictions on the insurance coverage for non-pharmacologic treatments, such as phototherapy. Indeed, we note our diagnosed prevalence estimates are lower than those of population-based studies, including a recent US-based study that estimated the diagnosed prevalence of vitiligo at 0.46%-0.77% (Gandhi et al., 2022), but comparable to a previous claims study that used the MarketScan database, Milliman Medicare Part D Claims database, and Milliman Medicaid Consolidated Health Cost Guidelines Sources Database and reported vitiligo prevalence at 0.05%. Third, data were limited to US enrollees with commercial health insurance and therefore may not be generalizable to other populations, such as those solely insured through government funded programs, such as Medicare, Medicaid, and Veterans Affairs, the uninsured, and those outside the US. Fourth, there was potential for inaccurate classification of the diagnosis of vitiligo when using data obtained from a single insurance claim, although the trends we observed were consistent in the sensitivity analysis. Fifth, race is not captured in IBM MarketScan, so we were unable to determine the variation in prevalence or incidence by race. Sixth, incident cases may be overestimated, as our study included a preindex observation period of 12 months with no diagnosis of vitiligo to identify incident cases, but it is possible that some of the cases identified may have had a previous claim that predated the observation period. In addition, new plan enrollees may have had a vitiligo diagnosis over the past several years but may be identified as an incident case in the current plan by virtue of their first observed claim in the current plan. Finally, the differences in vitiligo prevalence observed over time within this study could be due to a change in the underlying health plans, including the enrolled patient population that comprised the MarketScan database. The shifts in underlying plan characteristics may explain the changes observed. The increases in prevalence and incidence over time observed in this study may also be supported by the increases in prevalence and incidence estimates observed for autoimmune conditions like thyroid disease that are comorbid with vitiligo (Leese et al., 2008). Future studies may investigate trends in vitiligo-related comorbidities over time as explanatory empirical evidence. Nevertheless, our study strengths include the evaluation of annual trends for the incidence and prevalence of vitiligo over 5 years and addresses earlier literature gaps on the nonavailability of such trends. The study used a large database of commercially insured US populations. Large databases created by healthcare operations are frequently updated, cover large populations, and are well suited for estimating prevalence and incidence, as well as analyzing trends over time. In conclusion, there were 107,612

patients diagnosed with vitiligo and 85,821 newly diagnosed cases among the US ESI population in 2017. Our estimate of the annual diagnosed prevalence of vitiligo in 2017 (0.076%) was similar to the estimate reported by a previous analysis of insurance claims data (Lim et al., 2017). The annual diagnosed prevalence and incidence of vitiligo in the US both increased more than 5-fold in the 5-year period between 2013 and 2017. The cause of this significant increase warrants further investigation, including consideration of potential factors such as better access to medical care, greater awareness and better tools for vitiligo diagnosis, trends in associated comorbidities, and last but not the least, evolving diversity of the US

Table 1. Annual Prevalence of Vitiligo Among Individuals with Employer-Sponsored Insurance in the United States

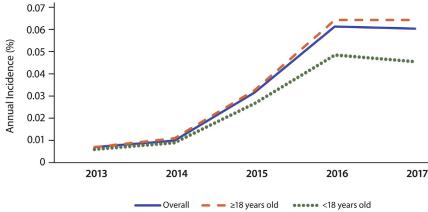
	Year					
Population and Variable	2013	2014	2015	2016	2017	
All ages						
Patients meeting vitiligo definition and continuously enrolled in a health plan (numerator), n	11,239	16,091	51,048	98,697	107,612	
Patients with continuous health plan enrollment (denominator), n	119,443,347	118,266,837	125,870,232	128,795,066	140,998,102	
Annual prevalence, %	0.009	0.014	0.041	0.077	0.076	
Sensitivity analysis numerator, n	4,244	6,216	18,876	28,277	31,935	
Sensitivity analysis annual prevalence, %	0.004	0.005	0.015	0.022	0.023	
Age ≥ 18 years						
Patients meeting vitiligo definition and continuously enrolled in a health plan (numerator), n	8,927	12,840	41,165	80,895	88,289	
Patients with continuous health plan enrollment (denominator), n	92,846,379	92,036,794	98,202,751	100,177,023	108,927,783	
Annual prevalence, %	0.010	0.014	0.042	0.081	0.081	
Sensitivity analysis numerator, n	3,214	4,646	14,288	22,166	25,170	
Sensitivity analysis annual prevalence, %	0.004	0.005	0.015	0.022	0.023	
Age <18 years						
Patients meeting vitiligo definition and continuously enrolled in a health plan (numerator), n	2,312	3,251	9,883	17,802	19,323	
Patients with continuous health plan enrollment (denominator), n	26,596,968	26,230,043	27,667,481	27,718,043	32,070,319	
Annual prevalence, %	0.009	0.012	0.036	0.064	0.060	
Sensitivity analysis numerator, n	1,030	1,571	4,588	6,110	6,765	
Sensitivity analysis annual prevalence, %	0.004	0.006	0.017	0.022	0.021	

population that may have contributed to the underlying genetic variation and population lifestyle changes that resulted in a higher incidence and prevalence over time.

Summary

Vitiligo is an autoimmune disease characterized by areas of skin depigmentation. Estimates of the point prevalence of vitiligo vary between 0.05% (insurance claims), 0.2% (population-based studies), and 1.8% (hospital-based studies). Data on the incidence of vitiligo are scant. We performed an observational, retrospective database study to estimate the annual diagnosed prevalence and incidence of vitiligo among the ESI population in the US. Data from the IBM MarketScan database were used to identify patients with an \geq 1 inpatient or outpatient claim for vitiligo from January 1, 2013, through December 31, 2017. Estimates of the annual diagnosed prevalence and incidence were calculated for the overall cohort and adult (aged \geq 18 years) and

pediatric (aged <18 years) patients. There were 107,612 patients diagnosed with vitiligo and 85,821 newly diagnosed cases among the ESI population in the US in 2017. Our estimate of the annual diagnosed prevalence of vitiligo in 2017 (0.076%) was similar to the estimate reported by a previous analysis of insurance claims data. The annual diagnosed prevalence and incidence of vitiligo both increased more than 5-fold in the 5-year period between 2013 and 2017. The cause of this significant increase warrants further investigation.



Data availability statement

De-identified claims data from the IBM MarketScan Commercial and Medicare Supplemental database were used for study purposes. Minimal data sets necessary to interpret and or replicate data in this paper are available upon request to the corresponding author.

\sim Overall \sim \sim \geq 18 years old \sim $<$ 18 years old	0002-4321-2467
Figure 2. Annual incidence of vitiligo among individuals with employer-sponsored insurance in the	Keisha Maugh 7233-7487
United States. Overall data are those for adults and adolescents. Incidence was based on 1 or more	Amit G. Pand
inpatient or outpatient claim.	916-2651

ORCIDs

Markqayne	e Ray: <mark>h</mark>	http://orcid.org/0000-0001-86
91-3756		
Kavita	Gandhi	: http://orcid.org/0000-
0002-4521	-2467	
Keisha <i>I</i>		http://orcid.org/0000-0001-
7233-7487		
	Pandya:	http://orcid.org/0000-0002-3
916-2651		

Prevalence and Incidence of Vitiligo in the US

Table 2. Annual Incidence of Vitil	go Among Individuals wit	th Employer-Sponsored Insura	nce in the United States

	Year				
Population and Variable	2013	2014	2015	2016	2017
All ages					
Patients meeting vitiligo definition and continuously enrolled in a health plan (numerator), n	8,319	11,896	39,940	78,834	85,821
Patients with continuous health plan enrollment (denominator), n	119,434,412	118,255,211	125,842,664	127,844,265	140,901,558
Annual incidence, %	0.007	0.010	0.032	0.062	0.061
Sensitivity analysis numerator, n	2,362	3,500	11,620	17,390	20,446
Sensitivity analysis annual incidence, %	0.002	0.003	0.009	0.014	0.015
Age ≥ 18 years					
Patients meeting vitiligo definition and continuously enrolled in a health plan (numerator), n	6,727	9,644	32,571	65,294	70,952
Patients with continuous health plan enrollment (denominator), n	92,839,434	92,027,575	98,180,722	100,136,233	108,849,402
Annual incidence, %	0.007	0.011	0.033	0.065	0.065
Sensitivity analysis numerator, n	1,765	2,585	8,748	13,698	16,116
Sensitivity analysis annual incidence, %	0.002	0.003	0.009	0.014	0.015
Age <18 years					
Patients meeting vitiligo definition and continuously enrolled in a health plan (numerator), n	1,592	2,252	7,369	13,540	14,869
Patients with continuous health plan enrollment (denominator), n	26,594,977	26,227,636	27,661,942	27,708,032	32,052,157
Annual incidence, %	0.006	0.009	0.027	0.049	0.046
Sensitivity analysis numerator, n	597	915	2,873	3,692	4,331
Sensitivity analysis annual incidence, %	0.002	0.004	0.010	0.013	0.014

CONFLICT OF INTEREST

MR and KG were employees of Pfizer at the time of study conduct and analyses; therefore, MR and KG may hold stock and/or stock options with Pfizer. KG is currently an employee of Janssen Pharmaceuticals and holds stock options of Johnson and Johnson. KM is an employee of STATinMED Research; STATinMED Research received payment from Pfizer for participation in this study. AGP declares acting as a consultant for AbbVie, Arcutis, Avita, Chromaderm, Immune Tolerance Network, Incyte, Pfizer, Twi, Viela Bio, and Villaris and holds stock or stock options in Tara Medical and Zerigo Health.

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AUTHOR CONTRIBUTIONS

Conceptualization: MR, KG, KM, AGP; Data Curation: MR, KG, KM; Writing – Original draft Preparation: MR, KG, KM, AGP; Writing – Review and Editing: MR, KG, KM, AGP

Markqayne Ray^{1,5,*}, Kavita Gandhi^{1,5}, Keshia Maughn² and Amit G. Pandya^{3,4}

¹Patient & Health Impact, Pfizer Inc, Collegeville, Pennsylvania, USA; ²STATinMED Research, Plano, Texas, USA; ³Department of Dermatology, Palo Alto Foundation Medical Group, Sunnyvale, California, USA; and ⁴Department of Dermatology, University of Texas Southwestern Medical Center, Dallas, Texas, USA

⁵Affiliation when this study was conducted.

*Corresponding author

e-mail: markqayne@gmail.com

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