# The incidence of psoriasis in Chile: an analysis of the National Waiting List Repository

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# **Summary**

**Background.** Psoriasis is a serious and chronic noncommunicable disease. However, the fundamental measure of disease occurrence, the incidence, has been scarcely reported globally. There are no previous studies of psoriasis incidence in Latin America.

**Aim.** To estimate the incidence rates of psoriasis in Chile during 2016 and 2017 using an administrative database, the Waiting List Repository.

**Methods.** We examined referrals of psoriasis at onset, made by physicians to dermatologists, evaluated the agreement of diagnosis, and estimated the incidence of the disease considering the eligible population at risk.

**Results.** In most cases, the referrals corresponded to incident cases of psoriasis (73.3%; 95% CI: 66.6-79.2). The national incidence rates of psoriasis were 22.1 (95% CI: 21.1-23.1) and 22.7 (95% CI: 21.8-23.6) per  $100\ 000$  person-years in 2016 and 2017, respectively. The most common type of psoriasis was the late-onset type. We observed a high variation in the figures throughout the country, with a range from 0.75 (95% CI: 0.3-1.5) per  $100\ 000$  person-years in the Metropolitan region to 164.9 (95% CI: 138.6-195.1) per  $100\ 000$  person-years in the Aysen region.

**Conclusion.** We describe for the first time the incidence of psoriasis in a Latin American country. Our findings could potentially guide collaborations to improve our global understanding of psoriasis in Latin America.

## Introduction

Psoriasis is a common chronic inflammatory skin disease that entails both a physical and psychological burden, and is associated with multiple comorbidities. The available epidemiological information on psoriasis

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varies depending on the geographical region assessed. Only 19% of the countries of the world report information on the epidemiology of psoriasis, according to the latest available systematic analysis. The majority of information comes from the USA and Europe, and focuses on disease prevalence.

Incidence studies of psoriasis are limited in number and there are no established epidemiological criteria for the definition of a new psoriasis case. To our knowledge, only 10 countries have reported on the incidence of psoriasis at a population level, with studies coming from Canada, Denmark, Germany, Italy, Israel, Russia, Taiwan, the Netherlands, the UK and the USA. The incidence rates reported in these studies range between 22 and 521 cases per 100 000 person-years. This broad range is due in part to different criteria being used, and the data show significant geographical variations.

A recent systematic analysis reported an estimated prevalence of psoriasis of 0.87% (95% uncertainty interval 0.29%–2.41%) for high-income southern Latin America. There are no population-level reports of the incidence of psoriasis from the region, a finding that was also corroborated by a regional systematic review.

The aim of the present study was to examine the incidence of psoriasis in Chile. We show the development and validation of an algorithm that detects new cases of psoriasis, and then demonstrate its use by estimating annual incidence rates of psoriasis in Chile during 2016–2017, using an administrative database, the Waiting List Repository (WLR). This database records the first contact of a patient requiring specialist care for a determined problem within the public healthcare system. A detailed description of the Chilean healthcare system, the database and its processing can be found in Data S1 (Appendix; section headed 'The Chilean Public Healthcare System').

## Methods

As de-identified data were provided, the study did not require ethics committee approval as the information is public.

## Study design

We performed an open cohort study using the WLR. Access to the WLR was obtained by request through the Chilean transparency law to the 29 health services of the country from January 2017 to December 2018.

In Chile, primary care physicians must refer all patients with psoriasis (even those with mild disease) according to the requirement expressed in the national medical curriculum; this requirement is assessed by a mandatory test before PCPs are allowed to provide medical care in the country. Therefore, our definition of an incident case was the record of the first referral due to psoriasis in the WLR. To validate this selection of cases, namely, to determine if the referral corresponded to the first diagnosis of psoriasis, we examined the commentaries made by the dermatologists after care was provided, and the positive predictive value (PPV) was calculated. A comprehensive description of

the validation process can be found in Data S1 (Appendix; 'Validation') and Table S1.

The population at risk of developing the disease was defined as a dynamic population, taking into account beneficiaries of the public health system at mid-year.

#### Statistical analysis

Crude and direct standardized incidence rates were calculated, and adjusted by age and sex with respect to the most recently observed population, FONASA-2017 (see Data S1). 95% CI of each incidence rate was computed as previously reported.<sup>6</sup>

## Results

The PPV of the algorithm was 73.3% (95% CI: 66.6–79.2). The flowchart representing the steps required to process the database and to validate the referrals as incident cases can be seen in Fig. 1. The detailed figures and characteristics of the study population are presented in Data S1 (Appendix; 'Characteristics of the study population'), Figure S1, and Tables S2 and S3.

#### Standardized incidence rates of psoriasis

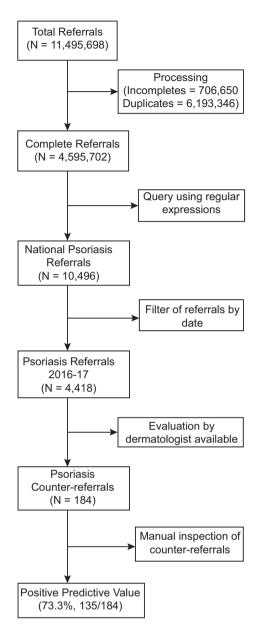
The national standardized annual incidence rates of psoriasis were 22.1 (95% CI: 21.1–23.1) and 22.7 (95% CI: 21.8–23.6) per 100 000 person-years for 2016 and 2017, respectively.

## Incidence rates by age

Cases, population and estimates were stratified by age groups (Table 1). The incidence rates of psoriasis were similar for both years, with the minimum incidence found in the demographic age group 0–5 years: 6.4 (95% CI: 4.5–8.9) and 4.9 (95% CI: 3.4–6.9) per 100 000 person-years for 2016 and 2017, respectively. The maximum incidence rate was found in the age group 55–65 years, corresponding to 37.1 (95% CI: 33.5–41.1) and 39.4 (95% CI: 35.9–43.1) cases per 100 000 person-years for 2016 and 2017, respectively. The incidence then dropped for the age group > 75 years, corresponding to 14.6 (95% CI: 11.5–18.5) and 16.5 (95% CI: 13.4–20.1) cases per 100 000 person-years, for 2016 and 2017.

#### Incidence rates by sex

Cases, population and estimates were also stratified by sex (Table 1) and were found to vary differently



**Figure 1** Flowchart of the validation algorithm. Raw data requested through the Chilean transparency law were termed 'total referrals', and data after processing were termed 'complete referrals'. Within this data subset, a query using regular expressions was performed, obtaining the national psoriasis referrals. Referrals were filtered by the time window of this study, thus obtaining psoriasis referrals for 2016–17. Data were also available for specialist evaluation of the referral (i.e. the counterreferral) for one of the largest health services in the country, the South-East Metropolitan Health Service of Santiago, and with these data we computed the positive predictive value, with 135 of 184 cases established as incident cases for 2017.

throughout life depending on sex (Fig. 2). In women there were two peaks of incidence, one during child-hood and adolescence (5–15 years) and a second in

middle age (from 45–65 years), whereas the incidence for men showed a single later peak (56–75 years). For both sexes, the incidence rates decreased from the age of 75 years.

# Incidence rates by region

To examine the occurrence of psoriasis throughout Chile, we plotted the incidence rates on a choropleth map (Fig. 3a). The rates fluctuated from a minimum of 0.75 (95% CI: 0.3–1.5) cases per 100 000 person-years in the Metropolitan South Health Service (Santiago; central zone, 33.4°S), to a maximum of 164.9 (95% CI: 138.6–195.1) cases per 100 000 person-years in the Aysen Health Service (below southern region; 46°S). The median of the observed figures across the health services was 20.9, with 75% of the observations having an incidence of < 49 cases per 100 000 person-years (Fig. 3b).

#### Discussion

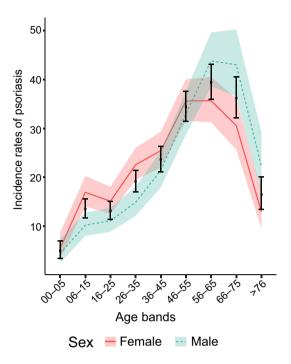
We report the first population-based cohort study examining the incidence of psoriasis in Chile, and indeed the first in Latin America. We used the WLR to perform the study. Unusually, and different from European results, we observed that most new cases of psoriasis corresponded to the late-onset type (occurring for the first time in people aged > 40 years), with earlier presentation in women and a slight predominance of new male cases in adulthood and old age. We also found geographical variation in the incidence of psoriasis across the country. We believe these variations can be explained by the different levels of complexity in the Chilean healthcare system, i.e. the different capabilities of the 29 health services to adequately manage referrals to dermatology specialists. As in other countries, the distribution of dermatologists is not uniform in Chile and is concentrated in large conurbations.7

A strength of this study is that it was conducted both on a national scale and in individual geographical areas. This high coverage of patients allowed us to depict the geographical heterogeneity in psoriasis incidence in Chile. Another strength was our use of WLR referrals to detect incident cases. This novel method allowed the use of semistructured data as cases were detected by searching for character strings rather than structured diagnostic codes. According to a recent systematic review, case detection using regular expressions is as good as that with rule-based or probabilistic algorithms, with the advantage that

 Table 1
 Cases, population at risk and standardized incidence rates of psoriasis in Chile by sex and age groups during 2016 and 2017.

		Females			Males			Total		
Year	Age group, years	Cases	Population	Rates <sup>a</sup>	Cases	Population	Rates <sup>a</sup>	Cases	Population	Rates <sup>a</sup>
2016	0–5	18	283 119	6.36 (3.77–10.05)	19	292 799	6.49 (3.91–10.13)	37	575 918	6.42 (4.52–8.86)
	6–15	97	604 166	16.06 (13.02–19.59)	74	630 304	11.74 (9.22–14.74)	171	1 234 471	13.85 (11.85–16.09)
	16–25	129	694 412	18.58 (15.51–22.07)	77	651 947	11.81 (9.32–14.76)	506	1 346 360	15.3 (13.28–17.54)
	26–35	139	707 114	19.66 (16.53–23.21)	89	558 925	12.17 (9.45–15.42)	207	1 266 040	16.35 (14.2–18.74)
	36-45	160	996 889	25.04 (21.31–29.24)	144	510 687	28.2 (23.78–33.2)	304	1 149 653	26.44 (23.55–29.59)
	46–55	225	671 469	33.51 (29.27–38.18)	190	566 988	33.51 (28.91–38.63)	415	1 238 456	33.51 (30.36–36.89)
	29–92	194	549 084	35.33 (30.53-40.67)	183	465 528	39.31 (33.82–45.44)	377	1 014 612	37.16 (33.5–41.1)
	99–75	16	385 615	23.6 (19–28.97)	113	313 298	36.07 (29.73–43.36)	204	698 914	29.19 (25.32–33.48)
	> 75	40	309 648	12.92 (9.23–17.59)	33	187 352	17.61 (12.12–24.74)	73	497 000	14.69 (11.51–18.47)
	Total	1093	4 843 594	22.45 (21.14–23.82)	901	4 177 828	21.88 (20.47–23.36)	1994	9 021 422	22.11 (21.15–23.11)
2017	0–5	18	327 163	5.5 (3.26–8.7)	15	337 766	4.44 (2.49–7.32)	33	664 928	4.96 (3.42–6.97)
	6–15	119	703 262	16.92 (14.02–20.25)	75	733 812	10.22 (8.04–12.81)	194	1 437 074	13.5 (11.67–15.54)
	16–25	121	800 246	15.12 (12.55–18.07)	84	756 794	11.1 (8.85–13.74)	202	1 557 040	13.17 (11.43–15.1)
	26–35	192	851 014	22.56 (19.48–25.99)	102	685 162	14.89 (12.14–18.07)	294	1 536 176	19.14 (17.01–21.46)
	36–45	191	751 090	25.43 (21.95–29.3)	131	611 941	21.41 (17.9–25.4)	322	1 363 030	23.62 (21.11–26.35)
	46–55	278	780 939	35.6 (31.54-40.04)	220	666 854	32.99 (28.78–37.65)	498	1 447 792	34.4 (31.44–37.56)
	29–95	235	659 038	35.66 (31.24-40.52)	248	566 468	43.78 (38.5–49.58)	483	1 225 506	39.41 (35.98–43.09)
	66–75	137	448 116	30.57 (25.67–36.14)	159	369 742	43 (36.58–50.23)	596	817 858	36.19 (32.19–40.56)
	> 75	48	371 264	12.93 (9.53–17.14)	21	229 514	22.22 (16.54–29.22)	66	622 009	16.48 (13.39–20.06)
	Total	1339	5 692 133	23.33 (22.09–24.62)	1085	4 958 052	22.19 (20.89–23.56)	2424	10 650 184	22.72 (21.82–23.64)

 $^{\rm a} \text{Cases}$  per 100 000 person-years (95% CI).



**Figure 2** Incidence rates of psoriasis by age of onset and sex in Chile, 2017. Continuous red line indicates females, dashed blue line represents males, bars represent both sexes (upper and lower bounds of the CI intervals for the national estimates of psoriasis incidence rates and shaded regions correspond to the CIs of the incidence rates.

implementation is simple and understandable.<sup>8</sup> The WLR method differs from previous work calculating incidence rates of psoriasis using administrative data<sup>2,3,9,10</sup> by the fact that the referral is an explicit representation of a new episode of disease. A final strength was the high value of the PPV that we found during the validation. Our results are similar to those presented by Basarab *et al.*,<sup>11</sup> who reported that the accuracy of a diagnosis of psoriasis made by general practitioners is 78%.

One of the main limitations of this study relates to the data-generating process. We believe that the psoriasis incidence rates estimated using this repository demonstrate a phenomenon known as the inverse care law.<sup>12</sup> This is a public health problem, and means that even though the most deprived populations have greater health problems (both in severity and frequency), they usually receive less care in comparison with wealthier populations. Although it is stipulated in Chile's professional regulations that all cases of psoriasis should be referred,<sup>5</sup> it seems unlikely that this actually happens. For example, we observed that in the Metropolitan Region (approximately 40% of the

Chilean population), the psoriasis incidence rates were higher in the east side of Santiago, in areas where average deprivation is lower or they have easier access to dermatologists (Fig. 3a). Consequently, the figures presented here should be interpreted with caution, as our estimates refer only to the population who consulted healthcare professionals, thus there is a potential issue of underestimating the true incidence of the disease. This is a limitation common to the majority of studies using population-based databases. As noted previously, 13 the computation of incidence from first contact with healthcare is accurate if the disease is severe and the data from the healthcare systems are centralized. However, Chile has a mixed healthcare system and patients can access dermatologists directly through the private system, which may contribute to a bias toward underestimation.

Our findings add to the existing literature from other regions that have used routinely collected data for the estimation of incidence of psoriasis at a national level, such as in North America, Europe and Asia. 2,3,6,9,10,14-20 The only previous work that studied the incidence of psoriasis in Latin America at a population level was a survey performed in the 1960s by Convit, and Farber and Nall, in Bolivia, Ecuador, Peru and Venezuela; they did not find any cases in the region. 21,22 A more recent cross-sectional survey performed in Brazil reported a psoriasis prevalence of 1.3% (95% CI: 1.1%-1.5%).<sup>23</sup> The range of our incidence rates across Chile is very wide, with the lower range being similar to the disease frequency in China<sup>10</sup> and the upper range being similar to that in the UK.6 The median was similar to a German study3 of 2012, which reported rates of 17-26 and 19-26 cases per 100 000 person-years for males and females, respectively; however, these figures are at variance with other estimates from Germany, which reported a rate of 521.1 per 100 000 person-years.<sup>2</sup> This discordance demonstrates that even for the same country. using the same database and assessing across very similar periods of observation, the estimates can vary widely depending on the methodology applied.

As previously reported in the literature, the incidence rates of psoriasis in our study showed evidence of different disease onset by age group and sex,<sup>24</sup> with the highest rates observed in age group of 46–65 years in women, and 56–75 years in men, which is similar to results from Canada, Denmark and the USA.<sup>9,15,17</sup> Overall, our study showed that the most common type of psoriasis was the late-onset type. We also observed a bimodal presentation in women, which was less pronounced than in other studies, but we did

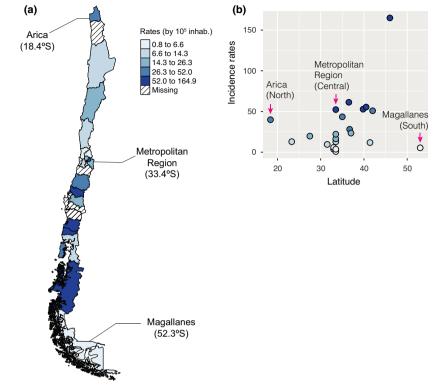


Figure 3 (a,b) Incidence rates of psoriasis in Chile in 2017, stratified by health service. (a) Choropleth map of the health services in Chile. The blue scale represents the value of the incidence rates of psoriasis, with darker tones indicating higher rates. The scale is categorized into five quantiles of data distribution.

(b) Psoriasis incidence rates in Chile showing variations between health services located in the same administrative region and along the geographical latitude spectrum.

not see a bimodal onset in men. As summarized in a recent systematic review, our study was concordant with an earlier onset of psoriasis in women than in men, and an increasing trend of incidence until 70 years, after which it decreased.<sup>24</sup>

## Conclusion

We provide for the first time a core measure of epidemiological information about psoriasis not previously available in Latin America. Specifically, we have proposed a novel methodology that uses an information resource, the WLR, not previously considered as a means to identify new cases of psoriasis at the population level. We believe the variations observed could be due to (i) awareness of the disease, meaning that better medical care could produce more referrals to dermatologists in a more timely manner; (ii) administrative and economic reasons, related to the number of available physicians in the geographical zones, and that wealthy patients can self-refer directly to a dermatologist; (iii) environmental factors, such as latitude, solar irradiance and pollution; and (iv) genetic admixtures of the populations, as psoriasis has an important genetic component in its aetiopathogenesis. Our findings have important consequences for health policy in the region, as they show that incidence rates vary throughout the country, with areas in which psoriasis is as frequent as in Europe and the USA.

## Conflict of interest

CL has participated in clinical trials sponsored by Pfizer and Novartis. CD has served as speaker and principal investigator to Lilly, Pfizer, AbbVie, Janssen, Novartis and Amgen, ITM has served as advisor and/ or received speaking fees and/or participated in clinical trials sponsored by AbbVie, Almirall, Amgen, BMS, Celgene, Eli Lilly, LEO Pharma, Janssen-Cilag, MSD, Novartis, Pfizer, Pierre Fabre, Roche, Sanofi and UCB. CEMG reports receiving honorariums or research grants from AbbVie, Almirall, Celgene, Eli Lilly, Galderma, Janssen, LEO Pharma, Novartis, Pfizer, Sandoz, Sanofi and UCB Pharma. DMA reports research grants from AbbVie, Almirall, Celgene, Eli Lilly, Novartis, UCB and the Leo Foundation. The remaining authors state they have no conflict of interest. The Global Psoriasis Atlas (GPA) project is delivered by the academic project staff based at the University of Manchester and University Medical Center Hamburg-Eppendorf. The GPA is a collaboration between three leading international organizations in the world of dermatology: the International Federation of Psoriasis Associations (IFPA), The International League of Dermatological Societies (ILDS) and the International Psoriasis Council (IPC). The GPA has been supported by grants and sponsorship from the Leo Foundation, AbbVie, Eli Lilly UK, Novartis Pharma AG, UCB and Almirall. All decisions concerning analysis, interpretation and publication are made independently of any industrial contribution. DMA and CEMG are funded in part by the National Institute for Health Research (NIHR) Manchester Biomedical Research Centre. JD and FV are funded by CMM-Basal AFB170001, supported by Cost Center 570111 - CIMT-CORFO and Project U-INICIA VID 2019 UI-004/19. SH is funded by ICM P09-015-F. DAAD 57220037 and 57168868. CORFO 16CTTS-66390. Fondecyt 1181823, **FONDEF** 19I10334 and ANID COVID0733. JMT is an employee of USZ and holds a 'Filling the GAP' scholarship.

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# **Supporting Information**

Additional Supporting Information may be found in the online version of this article:

**Figure S1**. Population pyramids of the study, database and source population in 2017.

Data S1. Appendix.

**Table S1.** Misclassification bias analysis of referrals of psoriasis using the inverse matrix method. Note that as the predictive values of the classification schemes are modified, the estimates of the algorithm are biased towards or away of the null.

**Table S2**. Study population (WLR), database population (FONASA), and source population (national) during the time window of the study.

**Table S3**. Regression analysis of genetic ancestry and incidence rates of psoriasis in Chile.