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Prevalence of Hidradenitis Suppurativa: A Meta-Analysis of Global

Hidradenitis Suppurativa Atlas Studies

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National Psoriasis Foundation, and CHORD COUSIN Collaboration (C3). A. Garg receives research grants from Abbvie, UCB, and National Psoriasis Foundation. He is co-copyright holder of the HS-IGA and HiSQOL instruments. G.B.E. Jemec has received honoraria from AbbVie, Chemocentryx, Coloplast, Incyte, Inflarx, Novartis, Pierre Fabre and UCB for participation on advisory boards, and grants from Abbvie, Astra-Zeneca, Inflarx, Janssen-Cilag, Leo Pharma, Novartis, Regeneron and Sanofi, for participation as an investigator, and received speaker honoraria from AbbVie, Boehringer-Ingelheim, Galderma and MSD. He has also received unrestricted departmental grants from Abbvie, Leo Pharma and Novartis. **Key points** Question: What is the global prevalence of Hidradenitis Suppurativa, and how does it vary by age, sex, geographical location, BMI, smoking status, GDP, and HDI? **Findings:** The sample included 22,743 participants, identifying 247 HS patients. The prevalence estimates showed considerable variations. The overall global prevalence of HS was 0.99% (95% CI 0.67 to 1.46%). Female sex was the only factor observed to be associated with HS prevalence. **Meaning**: This study found significant global variation in HS prevalence, estimating it at $\sim 1\%$. Further studies are needed to investigate the underlying causes of the substantial global variations in the HS prevalence.

209 210 **ABSTRACT** 211 **Objective** To estimate the global prevalence of Hidradenitis Suppurativa and study the differences 212 in prevalence by age, sex, geographical location, Body Mass Index (BMI kg/m²), smoking status, 213 214 Gross Domestic Product (GDP), and Human Development Index (HDI). **Design:** A proportional meta-analysis based on a series of prospectively planned, descriptive cross-215 sectional studies conducted as part of the Global Hidradenitis Suppurativa Atlas (GHiSA) Global 216 217 Prevalence Study. **Setting:** Multicenter study conducted across outpatient settings, excluding dermatology, in 23 218 219 countries spanning six continents. 220 **Participants:** A total of 22,743 healthy adults were recruited. Participants were screened using an HS-specific questionnaire, and those screening positive underwent examination by an HS-trained 221 physician to confirm diagnosis. Approximately ten percent of the screen-negatives were also 222 randomly selected for an examination to rule out disease. All studies conducted in accordance with 223 the GHiSA method that had finalized the data collection prior to 05.19.2023 were included in the 224 225 present proportional meta-analysis. 226 Main outcome and Measures: Pooled global prevalence of HS estimated using a random-effects 227 model. Prevalence was analyzed in relation to contextual factors with a meta-regression. 228 **Results:** The sample included 22,743 participants, identifying 247 HS patients. While the prevalence estimates showed considerable inconsistency ($I^2 > 75\%$, $\tau^2 = 0.747$), the overall random-229 230 effects global prevalence of HS was 0.99% (95% CI 0.67 to 1.46%). Female sex was the only factor observed to be associated with the prevalence estimates (% female, $\beta = 1.02$ [95% CI: 1.01 to 1.03]). 231 **Conclusions and Relevance:** An estimated global prevalence between 0.7 and 1.5% surpasses 232 previous global estimates. Substantial global variations in HS prevalence were observed. Female 233

Introduction

Hidradenitis Suppurativa (HS) is an inflammatory skin disease with a strong negative impact on patients' lives and largely no global epidemiological data. ^{1,2} HS is associated with increased mortality, singular psychosocial burden, and clinically important comorbidities ³⁻⁹. While Jfri and colleagues ¹ previously estimated HS prevalence, uncertanties remain due to study differences and limited global data. The Global Hidradenitis Suppurativa Atlas (GHiSA) has systematically gathered comparable HS prevalence data through the *Global Prevalence Study* (GPS). Our objective was to estimate the global HS prevalence across diverse populations and to analyze prevalence variations by age, sex, geographical location, Body Mass Index (BMI kg/m²), smoking status, Gross Domestic Product (GDP), and Human Development Index (HDI).

Methods

This study is a proportional meta-analysis derived from GHiSA GPS, a series of cross-sectional studies based on the GHiSA methodology. ¹⁰ Briefly explained, healthy adults accompanying a patient receiving care at an outpatient setting, excluding dermatology, were invited to complete a screening questionnaire. ^{10,11} Participants who screened positive were examined by an HS-trained physician to verify the disease. Approximately ten percent of the screen negative participants were randomly selected to be examined to assess the diagnostic accuracy. Only studies following the GHiSA methodology that had finalized the data collection prior to 05.19.2023 were included. A separate ethical approval was not needed for this pooled analysis.

Various data were extracted from the relevant studies and included information on geographical location, risk factors (age, sex, median BMI, BMI >30, and smoking status), prevalence estimates (including 95% confidence intervals), and data on HDI ¹² and GDP. ¹³

An initial descriptive analysis was conducted. A random-effects model was applied to pool the prevalence estimates with 95% confidence intervals, due to the assumption that there was no true effect size. Heterogeneity between studies was assessed using both the I^2 statistic and τ^2 . Additionally, 95% prediction intervals (PIs) were calculated. ¹⁴ By default, Wald-type CIs, based on a standard normal distribution, were calculated. The proportions were log-transformed using the logit method ¹⁵. Finally, a meta-regression was applied to explore the association of various factors and any heterogeneity between the included countries. The statistical analyses were performed using the R software (version 4.2.2) with the package *metafor* ¹⁶.

Results

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Out of the initial 74 countries invited to participate, a total of 51 countries were excluded due to lack of response (n=13), failure to initiate (n=12), obtain ethical approval (n=10), or finalize the study (n=14). Additionally, two countries were excluded for non-adherence to the study protocol. Finally, data from a total of 23 countries (n=25 studies), ¹⁷⁻⁴¹ spanning six continents, were included in this pooled analysis.

322 Study characteristics

This pooled analysis evaluated data from a cohort of 22,743 participants, of which 247 HS patients were identified. A summary of the study characteristics is presented in **Table 1**.

<u>Prevalence</u>

As illustrated in Figure 1, the overall random-effect pooled global prevalence of HS was 0.99%
 (95% CI 0.67 to 1.46; 25 studies). As anticipated, prevalence estimates showed considerable
 heterogeneity; I²>75%, τ² = 0.747 and 95% PI ranging from 0.18 – 5.39The prevalence estimates

ranged from 0.13% observed in the sample from Bangladesh ¹⁹ to 4.07% in the population from Saudi Arabia. ³⁶ High prevalence estimates (pragmatically defined as those higher than 0.99%) were seen in eight countries (Figure 1). **Supplementary Table 1** offers a subgroup analysis by continent and region. No statistically significant difference in continent, region and prevalence were identified. Finally, a statistically significant association between sex (% female) and prevalence was identified (β =1.02, 1.01 to 1.03). No other significant associations between demographic/risk factors were found. The result of

of HS increases by a factor 1.02 (i.e., 2%). See Table 3 for more information.

the full meta-regression can be found in Supplementary Table 2.

Discussion

Prospectively designed pooled prevalence studies have proven effective in minimizing the impact of measurement errors and biases. ⁴² The present analysis, carefully planned in advance and conducted through the GHiSA GPS initiative, presents the first and largest attempt to report of the global prevalence of HS. The findings reveal a global prevalence of 0.99% (95% 0.67, 1.46). The prevalence surpasses previous estimates (0.4%, 95% CI, 0.26, 0.63) reported by Jfri and colleagues ¹, derived from a meta-analysis including studies from Western Europe, Scandinavia, The US, and Australia. The variance in these estimates was attributed to clinical and statistical heterogeneity across studies. Similarly, our analysis identified considerable heterogeneity within our sample; however, due to the consistent and distinct methodological approach employed through the GHiSA GPS, this variance doesn't necessarily imply inconsistency, but rather that the proportions or populations are different.

The subgroup analysis by continent and region found no statistically significant overall difference in prevalence. The subgroup analysis by continent and region found no statistically significant overall difference in prevalence in prevalence. Most regions were only represented by one study, leading to

wide confidence intervals. Observed variations in prevalence across regions and continents has been found in other skin diseases such as eczema ⁴³ and psoriasis. ⁴⁴ These variations may be indicative of unidentified environmental and socioeconomic factors possibly influencing the prevalence patterns.

The meta-regression identified that only one variable, sex (% female) affected the prevalence. The

The meta-regression identified that only one variable, sex (% female) affected the prevalence. This aligns with current literature. ^{45,46} In contrast to prevailing literature, our study did not establish associations between BMI, BMI > 30, smoking and HS prevalence. ⁴⁶⁻⁴⁸ Disease severity was not assessed in this analysis and given that HS patients were identified among healthy adults, it is plausible that the majority exhibited mild disease. Additionally, our study focused solely on cigarette smoking and did not encompass other globally used tobacco alternatives. ⁴⁹ Investigations into the association of BMI, tobacco alternatives, and disease severity are warranted. Finally, no associations emerged between HS prevalence and HDI and GDP, potentially due to various factors, including the limitations of using these metrics alone, especially in regions with significant disparities in income distributions.

Limitations

The limitations of this study include the inherent constraints of large multi-center/country studies, including differences in language, culture, and healthcare systems, all potentially affecting the pooled prevalence estimate and its interpretation. Furthermore, selection bias needs to be considered, as a stratified sampling was not conducted. Additionally, countries were recruited via International League of Dermatological Societies member societies; only those that responded and had resources to participate were included, introducing potential non-response bias. As only 23 countries were included, a substantial amount of global data is therefore still missing. Moreover, ecological fallacy is possible for the meta-regressions exploring individual patient characteristics based on aggregated data. The strengths of this study include the rigorous and similar approach

379 employed to estimate the global prevalence of HS within the included studies. This study also successfully included numerous previously overlooked countries, and this review therefore provides 380 381 the most robust and up-to-date global HS prevalence estimates available. Conclusion 382 In conclusion, we estimated the global prevalence of HS to 0.99% (95% CI 0.67 1.46%). Proportion 383 384 of female sex was the only factor that was associated with the prevalence. Substantial global variations in HS prevalence were observed. Further studies are needed to investigate the underlying 385 386 causes of the substantial global variations in the HS prevalence. 387 Acknowledgments 388 389 Section for Biostatistics and Evidence-Based Research, the Parker Institute is grateful for the support provided by the Oak Foundation. 390 We further gratefully acknowledge the time and effort taken by the volunteers who participated in 391 392 all the countries. 393 394 **Funding** 395 DB has been supported by a research gran by Region Siælland – Medicinsporet. 396 Section for Biostatistics and Evidence-Based Research, the Parker Institute is supported by a core 397 grant from the Oak Foundation (OCAY-18-774-OFIL). 398 399

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 Table I: Study characteristics of Hidradenitis Suppurativa (HS) patients globally

Country	Continen t	Region	Cases with HS (n)	Total screened (n)	Prevalence* [95% CI]	Female (%)	Age in years	Median BMI**	BMI >30** (%)	Smoking (%)	GDP*** (\$)	HDI ****
Algeria ¹⁷	Africa	Northern Africa	11	1,434	0.77 [0.43 to 1.37]	90.9	34.0	26.2	18.2	0.0	163,044.4 4	0.745
Australia 18	Oceania	Australia/Ne w Zealand	9	1,002	0.90 [0.50 to 1.70]	66.7	30.0	41.8	100.0	44.4	1,552,667 .36	0.951
Bangladesh 19	Asia	Southern Asia	3	2,377	0.13 [0.04 to 0.37]	0.0	45.0	28.0	0.0	-	416,264.9 4	0.661
Chile ²⁰	Latin America and the Caribbean	South America	12	500	2.40 [1.40 to 4.10]	100.0	35.5	30.9	58.3	25.0	317,058.5 1	0.855
China ²¹	Asia	Eastern Asia	2	552	0.36 [0.10 to 1.31]	50.0	28.5	26.1	0.0	50.0	17,734,06 2.65	0.768
France ²²	Europe	Western Europe	18	525	3.43 [2.20 to 5.40]	55.6	36.0	24.1	16.7	44.4	2,957,879 .76	0.903
Ghana ^{23,24}	Africa	Western Africa	14	1,988	0.70 [0.42 to 1.20]	64.3	34.0	27.2	35.7	0.0	77,594.28	0.632
Greece ²⁵	Europe	Southern Europe	1	553	0.18 [0.03 - 1.02]	0.0	51.0	32.6	100.0	0.0	214,873.8 8	0.887
Greenland ²⁶	Northern America	Northern America	16	506	3.16 [1.95 to 5.10]	87.5	40.5	29.1	43.8	93.8	3,076.02	-
Indonesia ²⁷	Asia	South- Eastern Asia	14	3,237	0.43 [0.26 - 0.72]	50.0	29.0	23.0	7.1	28.6	1,186,092 .99	0.705
Iran ²⁸	Asia	Southern Asia	3	990	0.30 [0.10 to 0.89]	33.3	28.0	27.2	33.3	-	359,713.1 5	0.774
Malaysia ²⁹	Asia	South- Eastern Asia	7	500	1.40 [0.68 to 2.90]	42.9	33.0	23.5	0.0	14.3	372,980.9 6	0.803
Nigeria 30,31	Africa	Western Africa	51	1,700	3.00 [2.30 to 3.90]	66.7	33.0	26.6	25.5	3.9	440,833.5 8	0.535
North Macedonia 32	Europe	Southern Europe	5	597	0.84 [0.36 to 1.95]	40.0	36.0	28.1	40.0	60.0	13,825.05	0.770
Oman ³³	Asia	Western Asia	10	484	2.07 [1.10 to 3.80]	80.0	29.5	26.9	10.0	20.0	88,191.98	0.816
Papua New Guinea 34	Oceania	Melanesia	2	520	0.38 [0.11 to 1.40]	0.0	23.5	24.0	0.0	50.0	26,594.31	0.558
Poland ³⁵	Europe	Eastern Europe	15	932	1.61 [0.90 to 2.50]	33.3	38.0	28.7	33.3	-	679,444.8 3	0.876
Saudi Arabia 36	Asia	Western Asia	28	688	4.07 [2.83 to 5.82]	60.7	34.5	27.4	35.7	7.1	833,541.2 4	0.875
Serbia ³⁷	Europe	Southern Europe	4	490	0.82 [0.32 - 2.10]	50.0	40.5	25.7	25.0	75.0	63,082.05	0.802
South Africa 38	Africa	Southern Africa	9	500	1.80 [0.95 to 3.40]	88.9	36.0	27.3	22.2	33.3	419,015.0 2	0.713
Sri Lanka ³⁹	Asia	Southern Asia	2	993	0.20 [0.06 to 0.73]	50.0	26.5	29.4	0.0	50.0	88,927.26	0.782
The Netherlands 40	Europe	Western Europe	4	663	0.60 [0.32 to 1.54]	100.0	41.0	28.2	25.0	75.0	1,012,846 .76	0.941
Türkiye 41	Asia	Western Asia	7	1,012	0.69 [0.33 to 1.40]	57.1	44.0	25.8	14.3	57.9	819,035.1 8	0.838

 Baseline characteristics of identified HS patients, (n=247). The table includes patient characteristics of patients from 23 countries. *Percent ** Body Mass Index (kg/m2), ***Gross Domestic Product (GDP) per capita, denominated in US dollars (figures from 2021) (71), **** Human Development Index (HDI): Human Development Reports 2021/2022 (72), n= numbers.

Figure I: Global prevalence of Hidradenitis Suppurativa. Forest plot from a meta-analysis of GHiSA studies reporting of prevalence of Hidradenitis Suppurativa (back-converted from logits)

Country	Cases	Total				Estimate [95% CI]
Algeria	11	1434	⊢ ■			0.77 [0.43, 1.38]
Australia	9	1002	⊢			0.90 [0.47, 1.72]
Bangladesh	3	2377	■ ⊣			0.13 [0.04, 0.39]
Chile	12	500	⊢			2.40 [1.37, 4.18]
China	2	552	⊢ ■			0.36 [0.09, 1.44]
France	18	525	⊢	——		3.43 [2.17, 5.38]
Ghana	14	1988	⊢≡			0.70 [0.42, 1.19]
Greece	1	553	l 1			0.18 [0.03, 1.27]
Greenland	16	506	⊢ ■	—		3.16 [1.95, 5.10]
Indonesia	14	3237	H ≡ →			0.43 [0.26, 0.73]
Iran	3	990	⊢ ■────			0.30 [0.10, 0.94]
Malaysia	7	500	⊢ ■			1.40 [0.67, 2.91]
Nigeria	51	1700	⊢			3.00 [2.29, 3.93]
North Macedonia	5	597	⊢ ■			0.84 [0.35, 2.00]
Oman	10	484	⊢			2.07 [1.12, 3.80]
Papau New Guniea	2	520	⊢ ■			0.38 [0.10, 1.52]
Poland	15	932	⊢			1.61 [0.97, 2.65]
Saudi Arabia	28	688	⊢		-	4.07 [2.82, 5.83]
Serbia	4	490	⊢ ■			0.82 [0.31, 2.15]
South Africa	9	500	-			1.80 [0.94, 3.42]
Sri Lanka	2	993	⊦= 1			0.20 [0.05, 0.80]
The Netherlands	4	663	⊢ ■			0.60 [0.23, 1.60]
Türkiye	7	1012	⊢■			0.69 [0.33, 1.44]
Random effects model	247	22743				0.99 [0.67, 1.46]
Prediction interval						[0.18, 5.39]
Heterogeneity, Q = 171	.89. df = 22	$2. p < 0.001: l^2 = 88$	3.2%	,		[0.10, 0.00]
ristorogonomy, & - 171	.55, 41 22	., p . 0.001, 1 = 00				
			0 1 2 3 4	5	6	
			Prevalence (%)			

Forest plot (back converted from logits) illustrating the random-effects and fixed effect-model derived from a comprehensive meta-analysis of GHiSA studies reporting a prevalence of HS. The plot includes the summary measure, offering a precise representation of the pooled results. A total of 23 countries were included. CI: confidence interval ([prediction intervals], I2 index: percentage of variability not attributed to sampling error.

Characteristics	Geographical distribution*	Estimate (95%CI)	$ au^2$	I ² (%)	p-value	Prediction intervals
Overall (n=23)		0.99 (0.67–1.46)	0.747	88.2	-	0.18 - 5.39
Continent (n=23)			0.801	87.2	0.474	
Africa	4 (17.4)	1.33 (0.54–3.25)				0.19 - 8.86
Asia	9 (39.1)	0.68 (0.36-1.31)				0.11 - 4.30
Europe	6 (26.1)	1.04 (0.46–2.31)				0.15 - 6.76
Latin America and the Caribbean	1 (4.3)	2.40 (0.39–13.47)				0.19 – 23.87
Northern America	1 (4.3)	3.16 (0.52–16.82)				0.26 - 29.07
Oceania	2 (8.8)	0.63 (0.15-2.61)				0.07 - 5.80
Region (n=23)		,	0.619	83.2	0.171	
Australia/New Zealand	1 (4.3)	0.90 (0.17-4.62)				0.09 - 8.12
Eastern Asia	1 (4.3)	0.36 (0.05-2.82)				0.03 - 4.60
Eastern Europe	1 (4.3)	1.61 (0.32–7.67)				0.17 - 13.32
Melanesia	1 (4.3)	0.38 (0.05-2.98)				0.03 - 4.88
Northern Africa	1 (4.3)	0.77 (0.15-3.88)				0.08 - 6.90
Northern America	1 (4.3)	3.16 (0.64–14.17)				0.35 - 23.43
South-Eastern Asia	2 (8.7)	0.76 (0.23-2.42)				0.11 - 5.05
South America	1 (4.3)	2.40 (0.47–11.30)				0.26 – 19.00
Southern Africa	1 (4.3)	1.80 (0.34-8.93)				0.19-15.18
Southern Asia	3 (13.0)	0.20 (0.06-0.61)				0.03 - 1.32
Southern Europe	3 (13.0)	0.60 (0.20-1.86)				0.09 - 3.96
Western Africa	2 (8.7)	1.50 (0.49–4.50)				0.22 - 9.35
Western Asia	3 (13.0)	1.89 (0.74–4.76)				0.31 – 10.55
Western Europe	2 (8.7)	1.61 (0.49-5.21)				0.23 - 10.43

A subgroup analysis showcasing the prevalence of HS across various geographical locations. The table includes $\tau 2$, I2, and prediction intervals, providing insights into the in-between study variance and variability in effect sizes due to heterogeneity. p-values illustrating any significant difference between prevalence in various regions are also included. *Data are no (%) of included countries pr continent/region. CI: confidence interval.

Supplementary Table 2: Meta-regression of sociodemographic/risk factors and prevalence of HS

619						
Characteristics	No of	Summary*	Estimate (95%CI)	τ^2	I^2	p-value
	studies				(%)	
Female, % (range)	23	55.6 (0.0–100.0)	β 1.02 (1.01 to 1.03)	0.549	84.7	0.0034
Age in years, median (range)	23	34.5 (29.8–39.3)	β 1.00 (0.93 to 1.07)	0.786	88.8	0.9899
BMI median	23	27.2 (26.0–28.5)	β 0.99 (0.89 to 1.10)	0.795	88.9	0.8905
BMI > 30, %	23	25.0 (0.0-100.0)	β 1.01 (0.99 to 1.02)	0.753	88.4	0.4437
Current smoking, % (range)	20	38.9 (0.0–93.8)	β 1.00 (0.98 to 1.01)	0.648	86.7	0.6661
GDP **	23	372981	β 1.00 (1.00 to 1.00)	0.765	88.7	0.5160
(US \$ millions) 2021		(88560-826288)				
HDI 2021	22	0.792 (0.721–0.870)	β 5.01(0.14 to 183.51)	0.736	86.4	0.3800
620	•					

A meta-regression analysis illustrating the relationship between the prevalence of Hidradenitis Suppurativa (HS) and various demographic and risk factors. The estimate β indicates the proportional increase (or decrease) in odds of having HS pr one-unit increase in the sample. For % females, 1 unit increase in the proportion of females in the sample, indicates that the prevalence of HS increases by a factor 1.02, i.e. 2%. *Data are median percentages (range of percentages), or medians (range of medians) for aggregated data. BMI: Body Mass Index (kg/m2) GDP: Gross

Domestic Product in US \$ (millions), HDI: Human Development Index