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### EPIDEMIOLOGY OF CUTANEOUS LEISHMANIASIS IN DISTRICT KARAK, KHYBER PAKHTUNKHWA, PAKISTAN

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#### ABSTRACT

Cutaneous Leishmaniasis is a vector-borne infection caused by flagellated parasitic protozoans in the genus *Leishmania*. The bite of an infected female sandfly from the genera *Phlebotomus* or *Lutzomyia* can spread the infectious promastigote stage to humans. In many regions of Pakistan, especially Khyber Pakhtunkhwa, CL is regarded as one of the main public health issues. The goal of the current study was to look at the clinical and demographic trends of CL in Pakistan's Karak District, Khyber Pakhtunkhwa. Between March 2022 and April 2023, the survey was conducted with both local residents and temporarily displaced people (TDPs) living in 14 distinct villages within the district. There is total no588 people that were analyzed, 35 were TDPs and 553 were members of the local population. In the local population, the general prevalence of CL was 7.80%, whereas in TDPs, it was 7.00%. According to the age-wise distribution, children ages 1 to 10 had the highest occurrence in both categories, suggesting that younger people are more susceptible. Males had a much higher incidence than females, according to a gender-wise analysis. This could be because males are more likely to be exposed to sandfly bites outside. A comparison of the villages showed that Panos had the highest prevalence (11.00%) while Alamsheri had the lowest frequency (1.40%). The findings suggest that CL remains endemic in the Karak District, with notable variation in prevalence across different age groups, genders, and localities. These

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results highlight the need for targeted awareness programs, preventive measures, and vector control strategies to reduce disease transmission. Moreover, timely diagnosis and effective treatment facilities are required, particularly for vulnerable groups such as children and displaced population. These results indicate that cutaneous Leishmaniasis is as much common in male individual as compared to female. Suggesting that male individual are more exposed to the fly.

## 1. Introduction

Leishmaniasis is a serious global health concern caused by intracellular protozoan parasites of the genus *Leishmania*, belonging to the family Trypanosomatidae. Recognized as a neglected tropical disease (NTD), it disproportionately affects populations in resource-limited regions. Although *Leishmania* parasites were first identified in the late 1980s (Tarr et al., 1988), the disease continues to pose a major public health threat due to its complex epidemiology and inadequate control strategies. The parasites reside and proliferate within host macrophages, resulting in three primary clinical manifestations such as cutaneous leishmaniasis (CL), which affects the skin and mucocutaneous leishmaniasis which targets mucous membrane and which visceral leishmaniasis (VL) at invades internal organs (Hotez et al., 2006).

The two main species that cause CL in Pakistan are *Leishmania major* and *Leishmania tropica* (Alexander et al., 1998). Asia, Europe, and South America are among the regions where CL is endemic. Because of its constantly increasing incidence—more than one million new cases are recorded each year—the World Health Organization (WHO) has ranked it as one of the most urgent global health challenges (WHO, 2001).

Leishmaniasis is a serious and unmanaged public health issue in Pakistan, especially in regions that border Afghanistan and in cities that house sizable refugee populations (Kassi et al., 2008). The growing incidence of CL is caused by a number of environmental factors, including urbanization, poor sanitation, and sleeping outside, as well as personal risk factors, such as HIV infection, malnourishment, genetic predisposition, and low socioeconomic status (Rijal et al., 2010). Furthermore, it is thought that the spatial distribution and dynamics of transmission of vector-borne illnesses such as leishmaniasis are impacted by climate change (Patz et al., 2000). Numerous anatomical locations, including the lips, genitalia (chancriform), fingertips (paronychia), pulp of fingers (whitlow), eyelids, scalp, and palms/soles, can be impacted by cutaneous leishmaniasis (Ul Bari, 2006). In Pakistan, there are two epidemiological types of CL: anthroponotic CL (ACL), where humans are the only reservoir, and zoonotic CL (ZCL), which involves animal reservoirs (Desjeux, 2004). The main cause of ACL, which is prevalent across the nation, particularly in the province of Khyber Pakhtunkhwa (KP), is *Leishmania tropica* (Amtul et al., 2001 & Hussan et al., 2017). In endemic areas, lesions are locally known as "Kandahar sores," "tropical sores," or "Baghdad boils" (Soomro et al., 2009).

There have been reports of autochthonous transmission of ACL in a number of Pakistani regions, which are frequently characterized by a greater frequency among children and family clustering (Kolaczinski et al., 2004). *Leishmania donovani* is the main cause of the disease's more severe visceral form, known as VL (Tanoli et al., 2005). Both *L. donovani* and *L. infantum* are agents of VL, and *Phlebotomus alexandri* is the vector for both of them (Azizi et al., 2006).

When female sand flies inject promastigotes into the host's skin while feeding on blood, the transmission cycle starts (Gawade et al., 2012). Neutrophils and other phagocytic cells

quickly absorb these promastigotes. Promastigotes change into amastigotes and then the replicative form that propels the course of disease within the phagolysosomes of macrophages (Sacks et al, 2001). About 30 species of blood-feeding sand flies from the genera *Phlebotomus* and *Lutzomyia* are known to transmit leishmaniasis out of the 700 species of blood-feeding sand flies that are known to exist worldwide (Ali et al., 2016). There are roughly 37 species of sand flies known to exist in Pakistan (Kakarsulemankhel, 2009).

Leishmaniasis treatment is still difficult since it necessitates the long-term use of costly and toxic medications. There are currently no approved vaccinations for human use, despite the fact that vaccine development has been a priority for more than a century (Selvapandiyan et al., 2006). Results from early attempts to create a vaccine utilizing entire dead Parasite were mixed (Handman, 2001). Clinical examination serves as the primary basis for CL diagnosis, which is then verified by microscopic inspection of lesion samples (WHO, 2010). Particularly in cases of post-kala-azar dermal leishmaniasis (PKDL), lesions that manifest as hypopigmented macules, papules, or nodules may go untreated because there are no systemic symptoms (Mondal et al., 2010).

Remarkably, those who recover from visceral and cutaneous leishmaniasis frequently develop protective immunity against recurrent infections (Evans et al., 2012). Leishmanization, a technique that involves purposeful exposure to live parasites to provide long-lasting immunity following lesion healing, has historically been used to capitalize on this immunological response (Khamesipour et al., 2005). A robust cellular immune response that includes efficient parasite removal and macrophage activation is the main mediator of host resistance. The establishment of protective immunity depends on both innate and adaptive immunological components, including dendritic cells and antigen-specific CD4<sup>+</sup> and CD8<sup>+</sup> T cells (Stanley & Engwerda, 2007).

## 2. Material and Methodologies

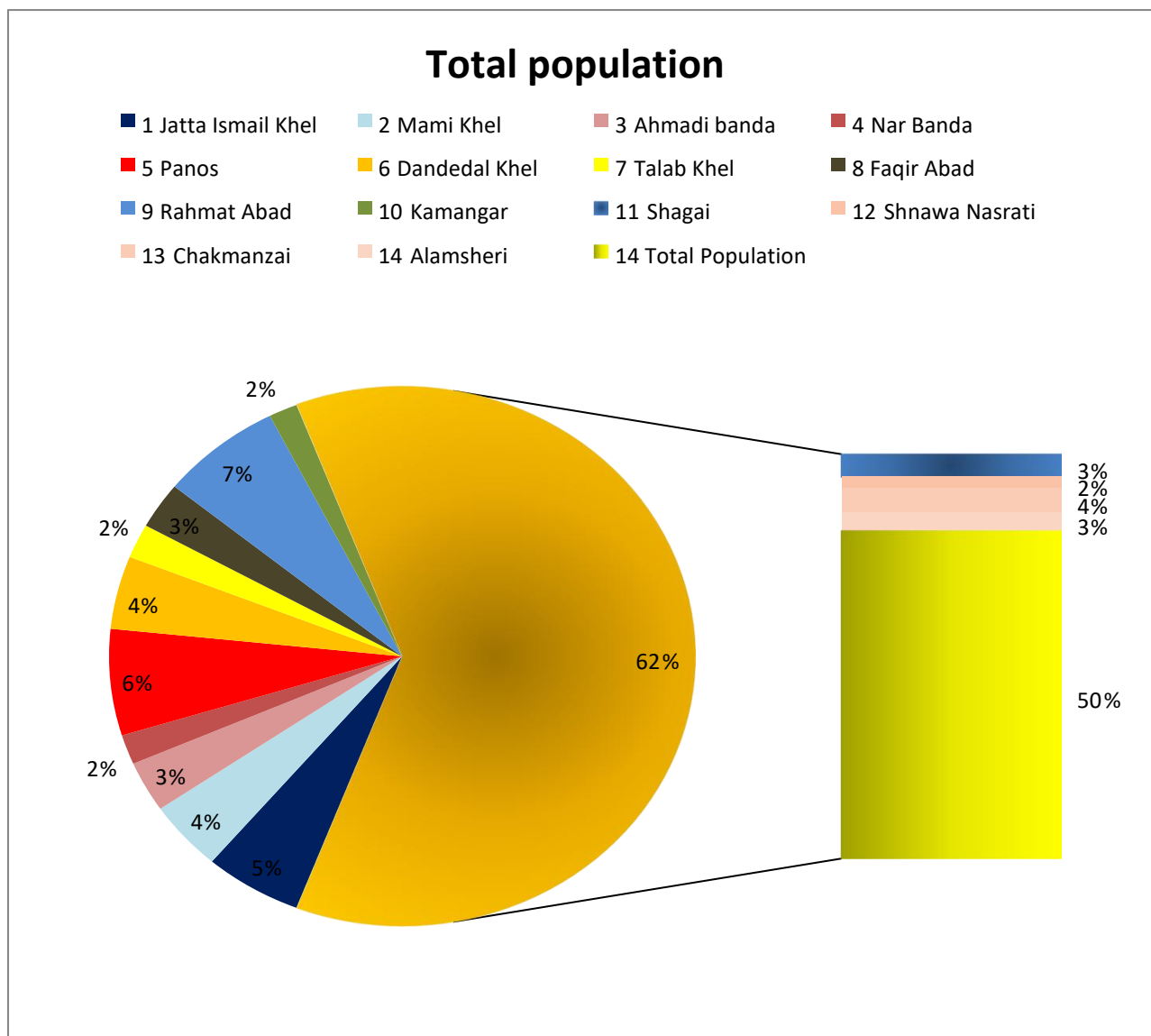
### 2.1. Study Area:

The study was carried out from March 2022 to April 2023 at 14 different villages of District Karak including small number of Tdps residing in the district. The studied areas are given in tabulated form. The population data provided by the population department of District karak.

**Table 2.1: Village Wise Detail of Studied Area**

S.No	Village Name	Total Population
1	Jatta Ismail Khel	6083
2	Mami Khel	4757
3	Ahmadi Banda	3448
4	Nar Banda	2017
5	Panos	7215

6	Dandedal Khel	<b>4961</b>
7	Talab Khel	<b>2320</b>
8	Faqir Abad	<b>3155</b>
9	Rahmat Abad	<b>7587</b>
10	Kamangar	<b>1792</b>
11	Shagai	<b>3910</b>
12	Shnawa Nasrati	<b>1915</b>
13	Chakmanzai	<b>4238</b>
14	Alamsheri	<b>3227</b>
	Total Population	<b>56625</b>



**Chart 2.2: Village Wise Detail of Studied Area**

## 2.2.Sampling:

The study of cutaneous leishmeniasis in various villages was based upon the data provided by the woman & children hospital karak city leishmeniasis center /opd. From the information supplied, we paid the patients a visit at their residence. On a questionnaire, other details were gathered, such as name, sex, age, occupation, type of lesion (active, scarred, dry, or wet), number of lesions, lesion location, travel history to an endemic area, house type (brick or mud),

Presence of animal and treatment profile.

In all, 7500 local residents (7000) and tdps (n=500) who lived in the area under study were surveyed. House to house survey was carried out by a team comprising of research supervisor and research student. Information were obtained from head of each house from randomly selected house in east west and north south directions. All individuals of a selected endemic village were eligible, while only active lesions individual were selected for the study.

### 2.3. Sampling design

A team of research project supervisors and students from the Zoology Department at GDC Takht-e-Nasrati Karak, along with a medical technologist and a representative from the leishmaniasis clinic of the civil hospital district of Karak, conducted a door-to-door survey in each of the 14 villages in the district that had previously been tabulated and a small number of idps living in the village. Patients with leishmaniasis were given questionnaires that asked about their age, sex, the number of lesions they had, and whether or not they used bed nets. Additionally, the heads of the households were questioned about their work, length of stay, number of animals (dogs, cattle), home type (concrete or mud), and hygienic conditions. Additionally, heads of households were asked if any members had active skin lesions or scars from CL or other causes. For additional care, several probable CL patients were sent to Leishmaniasis center. Information on each case was documented based on the quantity and location of lesions and scars, as well as factors that contributed to the infection, such as travel history, contact with an infected person, and sleeping outside. In order to determine the prevalence, people having lesions were regarded as cases of active CL.

### 3. Results:

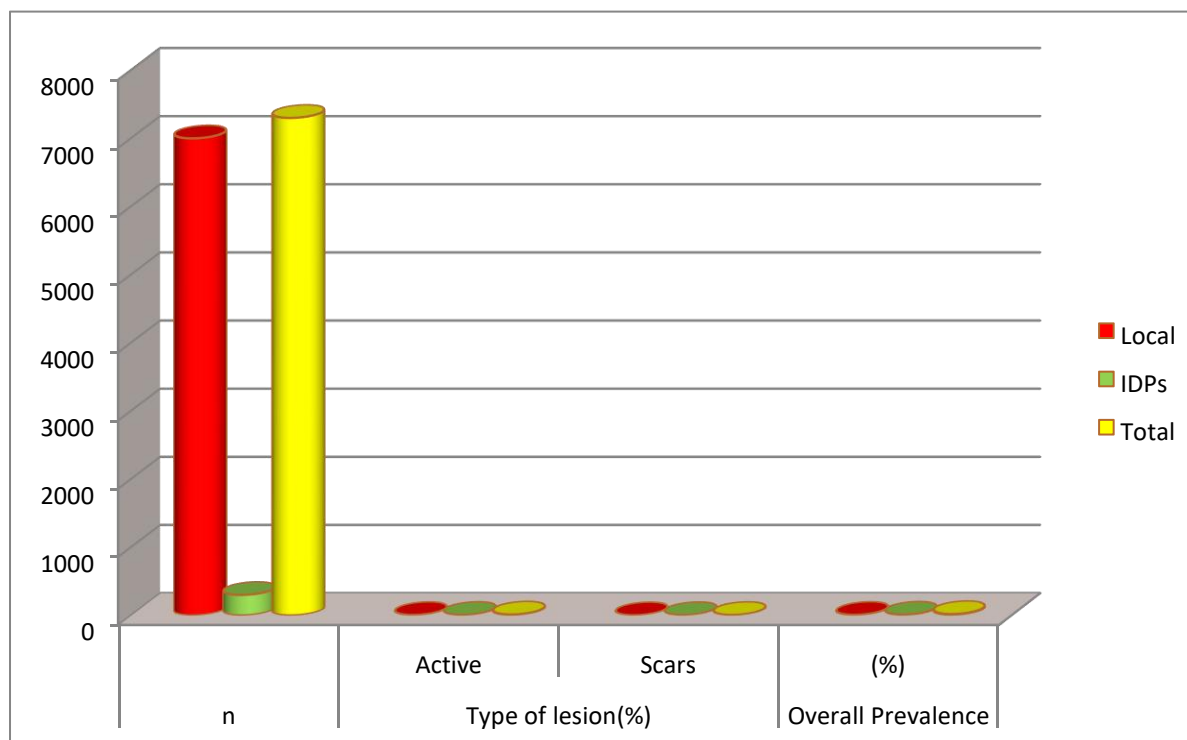
There are total 500 Tdps and 7,500 members of the local people were surveyed in 14 distinct villages in the District of Karak. Patients were given questionnaires to complete in order to collect data for research (Table No.1). From these localities, a total of 355 tdps cases (both active scar cl lesions) and 553 local residents were reported. There were 256/553 (46.29%) females and 297/553 (53.70%) males among them. A recorded 14.8% overall prevalence, which included 7.00% in tdps and 7.80% in the local population

**Table 3.1: Shows that overall high prevalence was recorded in the local Population.**

Population	N	Type Of Lesion (%)		Overall Prevalence (%)
		Active	Scars	
Local	7000	5.80	2.00	7.80

<b>IDPS</b>	500	4.06	2.46	7.00
<b>Total</b>	7500	12.80	6.00	14.80

**Chart 3.2: Showing Overall Prevalence of Cutaneous Leishmaniasis Patients**



**Table 3.3: Shows the Prevalence of Cutaneous Leishmaniasis in Local Population Of District Karak Kpk Pakistan.**

Location	Total Population	Population Surveyed	Positive Cases	Active Cases	Scars	Total Prevalence (%)
Jatta Ismailkhel	6083	824	101	71	30	8.61
Mami Khel	4757	512	47	32	15	9.2
Ahmadibanda	3448	260	21	14	7	8.1
Nar Banda	2017	158	09	06	3	5.7

Panos	7215	1272	140	11 2	28	11.0
Dandedalkhe	4961	550	32	18	14	5.8
Talab Khel	2320	304	23	17	6	7.6
Faqir Abad	3155	454	41	35	6	9.0
Rahmat Abad	7587	960	94	77	17	9.8
Kamangar	1792	72	04	3	1	5.6
Shagai	3910	250	9	7	2	3.6
Shawanasrati	1915	130	4	4	0	3.1
Chakmanzia	4238	764	18	11	7	2.4
Alamsheri	3227	706	10	08	2	1.4
Total	5665	726	553	45	2	7.0%
				5.8%	2.0%	



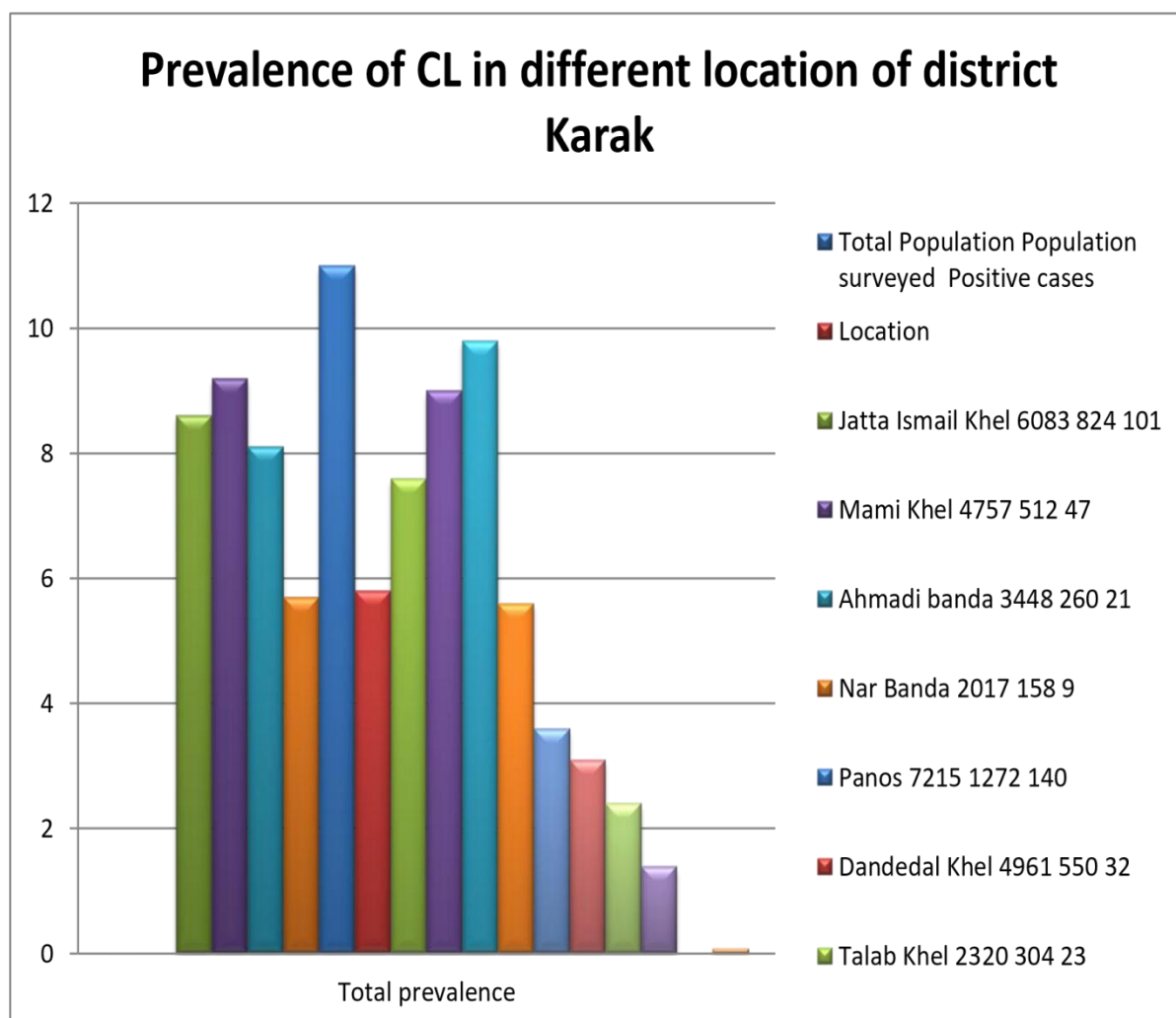
As we See from the above table 3.3 data that highest prevalence was recorded in village “Panos” (11.0%) in lowest in “Alamsheri” (1.4%).

**Chart 3.4: Showing Overall Prevalence “CL” Patient in District Karak**

### 3.1.Age Wise Distribution Of CL In District Karak

The Patients were separated into age groups ranging from 1 year to >80 years in order to examine the susceptibility of CL with age. The young age group (1–10 years old) had the

**Chart Showing “CL” Patients In Different Location Of District Karak.**

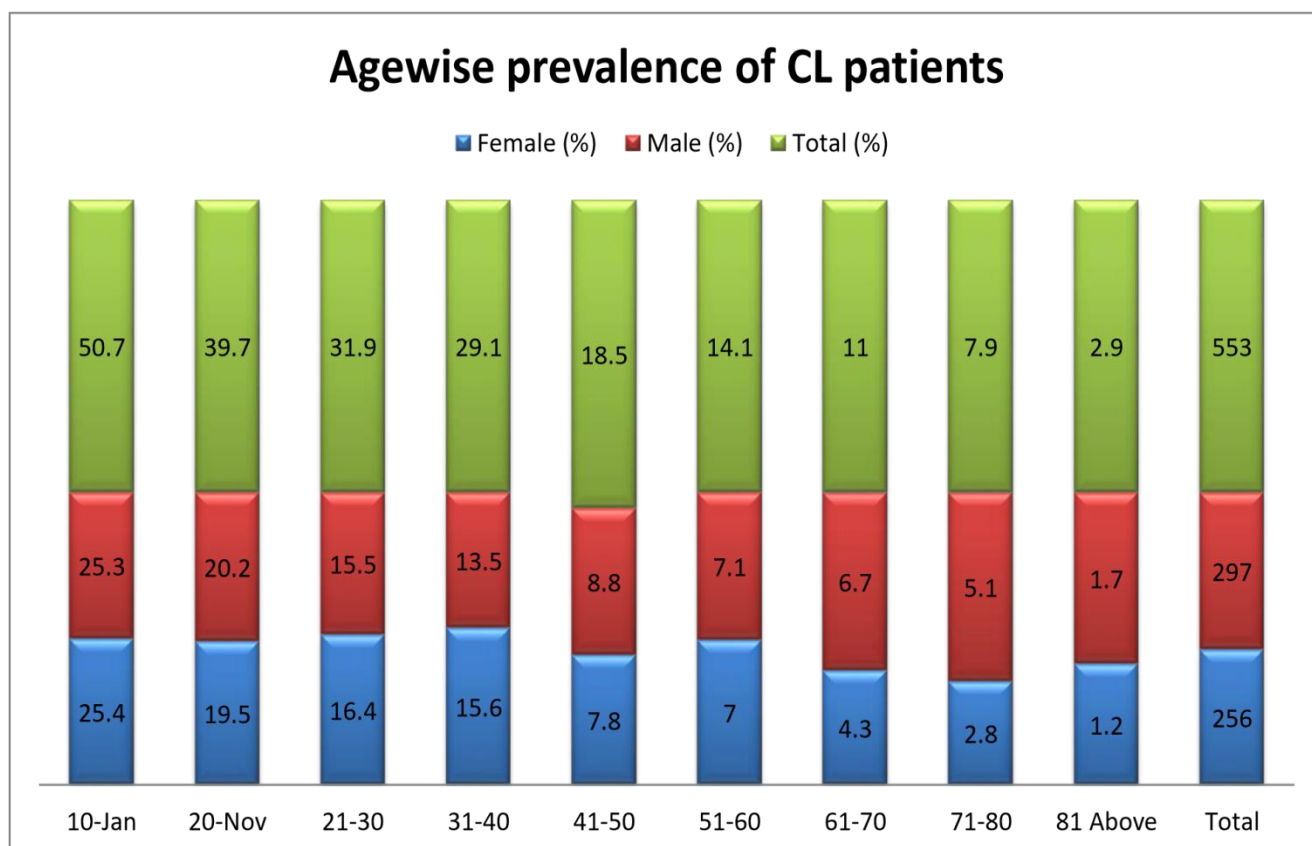


highest prevalence (55.9%) in local people, whereas the 11–20 age group had the highest frequency (57.4%) in TDPS. However, in both local and TDPS, people older than 80 years were less vulnerable. Additionally, as shown in table 04, the prevalence rate was higher in males in both local and TDPS. The percentage of patients in the local and IDPS populations by age and gender from March 2022 to April 2023.

**Table 3.5: Shows the Age Wise Distribution of CL in District Karak.**

		Local Population			Temporarily Displaced People(TDPS)		
S No.	Age In Year	Female (%)	Male (%)	Total (%)	Female (%)	Male (%)	Total
1	1-10	25.4	25.3	50.7	16.7	11.1	27.8
2	11-20	19.5	20.2	39.7	33.3	4.1	57.4
3	21-30	16.4	15.5	31.9	12.1	5.6	17.7
4	31-40	15.6	13.5	29.1	10.6	16.7	27.3
5	41-50	7.8	8.8	18.5	7.6	13.0	20.5
6	51-60	7.0	7.1	14.1	6.1	11.1	17.2
7	61-70	4.3	6.7	11.0	6.1	3.7	9.8
8	71-80	2.8	5.1	7.9	4.5	93	13.8
9	81abov	1.2	1.7	2.9	3.0	5.6	8.6
	Total	256	297	553	20	15	35

**Chart 3.6: Showing Age Wise Distribution Among Patient**  
**3. Chart Showing Age Distribution Among Patient**



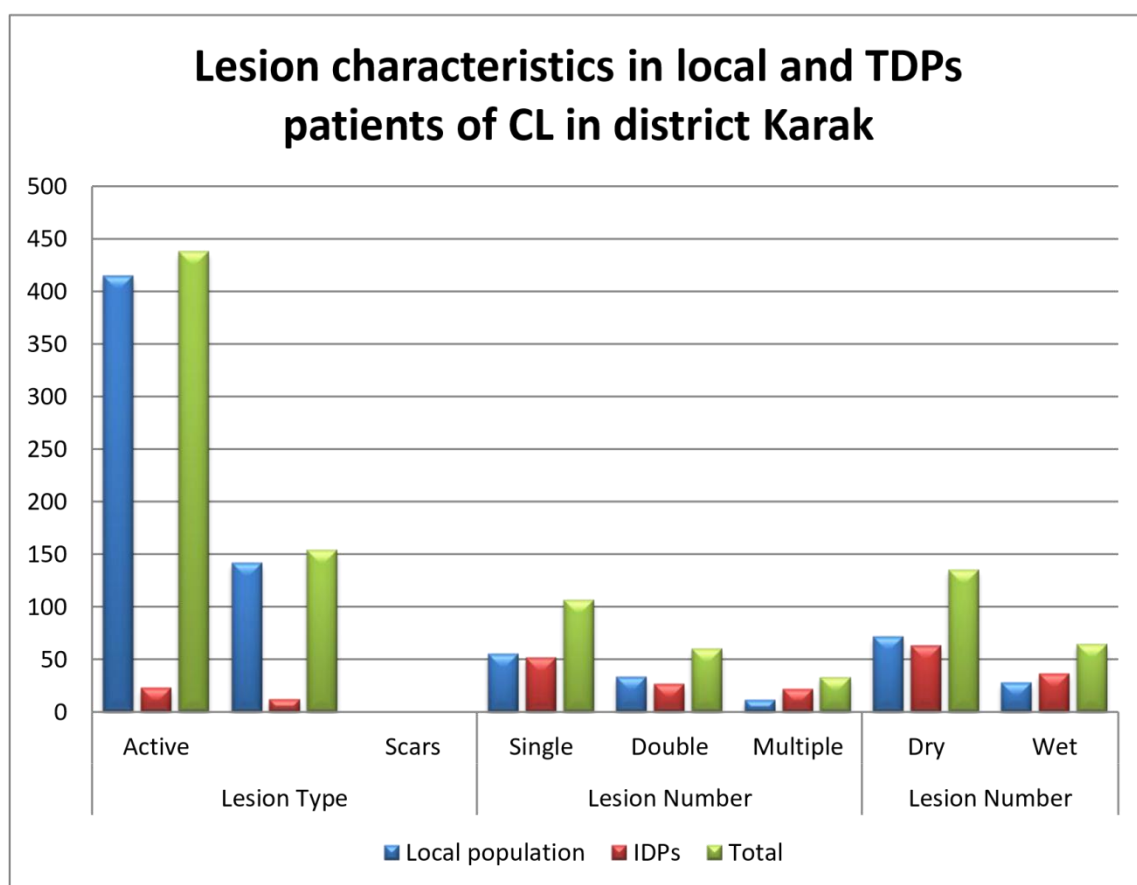
### 3.2. Lesion Characteristics in Local and Tdps Patients Of CI in District Karak.

Furthermore patients were also analysed on the basis of different characteristics of lesion like active (Dry and Wet) and scar, number of lesions and position of lesion (table no.5).it was observed that, in both types of population, the highest number (55.15%) of patients were having single active or scar cl lesion while majority of patients exhibited active dry form of lesion

**Table 3.7: Shows Lesion Characterstics in Local and Tdps Patient of CI District Karak**

Population	Lesion Type	Lesion Number	Lesion Number
n			

	Active	Scars %	Single %	Double %	Multiple %	Dry %	Wet %
<b>Local Population</b>	415	142	55.15	33.45	11.39	71.97	28.02
<b>IDPs</b>	23	12	51.67	26.67	21.67	63.33	36.67
<b>Total</b>	438	154	106.82	60.12	33.06	135.30	64.69



**Chart 3.8: Shows Lesion Characteristics in Local and Tdps Patient of CL in District Karak.**

The most noticeable site of the lesion was on exposed body parts, such as the face, hands, and feet. Factors such as house type (cemented or non-cemented), roof type, presence of animals in or near the house, education profile of the infected population, knowledge of the disease, and access to treatment were also investigated, along with a few other unusual clinical presentations and some individuals (not identified separately in the results)

#### 4. Discussion

Humans constitute a host reservoir for the cl parasite, human population mass plays a crucial role in the transmission of cl. As a result, it may be assumed that cl is more common in places with high population density. Due to the prevalence of both ACL and ZCL caused by *L. tropica* and *L. major*, Pakistan is particularly significant in terms of leishmaniasis. The seasonal recurrence of disease shows that physical factors have an impact on the transmission of CL. Variations in the weather and climate limit or worsen the conditions that are conducive to the spread of disease. For instance, higher humidity affects the growth and development of the leishmanial parasite by shortening its incubation time. The regions of our country that are close to the Afghan border, such as the towns and villages of Baluchistan province, Kurram, Mohmand, and the north Waziristan District of Khyber Pakhtunkhwa, cutaneous leishmaniasis is more prevalent. Our findings show that the most important factor raising the likelihood of cl incidence is the relocation of IDPS from the hyper endemic zone.

The goal of the current study was to determine the prevalence of cutaneous leishmaniasis in the Karak, KPK, Pakistan area between March 2022 and April 2023. Our study found that the prevalence of CL in local populations is higher (8.66%) than that of IDPS (7.00%). The official from the district Karak health department attributes this high prevalence to the movement of IDPS from the cl hyper endemic zone (north and south Waziristan) and the movement of animals from Baluchistan province to the district Karak for commercial purposes. Similar to the previous study from Peshawar. The overall prevalence rate of cutaneous leishmaniasis was higher in men, and sex-wise differences were also previously reported. In the 2019 study conducted in District Kohat, the prevalence was higher in men than in women and the study conducted and also showed that the disease is more common in men than in women. The results obtained and showed that 54 (90%) of the patients were male, while 10% were female, which is extremely high in men (UL bari et al., 2006). The other studies likewise demonstrate that CL affects both sexes to differing degrees at all ages and that the disease affects both sexes. We discovered an overall prevalence of 7.00% among IDPS resident in the local community, with an active lesion of 6.46% and scars of 2.20%. In comparison to the 2017 study conducted in the Kohat district of KPK, where the active lesion prevalence was 39.19% and the scars lesion prevalence was 1.26% (Hussain et al., 2017), and the study conducted in the refugee camp by Brokered al., where the active lesion prevalence was 2.7% and the scars lesion prevalence was 4.7%, the district has a 4.60% active lesion and 2.40% scars. The percentage of IDPS in the research region was also included, but our main goal was to ascertain the prevalence of leishmaniasis in the local population of District Karak. The majority of cases were found to have dry type lesions, with a small number having wet type lesions, which is similar to the study. The current study also found a very high prevalence of active lesions compared to scars lesions, and other studies in Afghanistan have also looked into high prevalence of scars lesions. Additionally, the findings were broken down by age group, with the highest prevalence (55.9%) occurring in the 1–10 age group. This is in contrast to the findings of the study carried out in the Dargai region Kpk, where the highest prevalence (34.32%) was examined in the 11–20 age group. We also concur with I Khan et al.'s 2013 study, which found that leishmaniasis was most common in patients aged 1–10 years. Our study also found that children aged  $\geq 10$  years had the highest rates of

leishmaniasis (55.9%), which may be due to social factors as the kids play outside without adequate body protection from sand flies, which are the leishmaniasis vector. The 61.11% of patients had a single type of lesion, 29.28% had a double lesion, and 9.61% had multiple lesions. In contrast to our study, a previous study conducted in Kohat revealed a higher prevalence percentage of patients with a single type of lesion (60.7%) double 96.75) and multiple lesions (86.67%) also found a lower percentage of single lesions (50.9%) and double lesions (24.6%) and 29.4% of patients had numerous (3–15) lesions. Our results exceeded those of a study in Karak, which found that 58.4% of patients had a single lesion, 29.2% had two, and 12.38% had more than two lesions. In contrast, our study found slightly fewer patients with more than two lesions.

## 5. Conclusion

Cutaneous leishmaniasis emerged as a serious health issue in a number of Karak district communities. Its prevalence in Karak is influenced by a number of environmental variables, including a poor sewage system, inadequate sanitation, particularly in rural areas, a lack of safe drinking water, a dense population, and deforestation. The community should be educated about sand flies and control initiatives should be started in order to reduce the vector population in Karak. Based on our findings, it was also determined that men are more likely than women to have CL. Men are more gregarious and spend more time outdoors in the evening and night, which exposes them to sand fly bites, which accounts for the high occurrence in men. The current study also found that the population of people under the age of fifteen had the highest rate of infection compared to the other groups.

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