

Original Article

The Effect of Geographical and Climatic Factors on the Distribution of *Phlebotomus papatasi* (Diptera: Psychodidae) in Golestan Province, an Endemic Focus of Zoonotic Cutaneous Leishmaniasis in Iran, 2014

Seyed Hamid Hosseini^{1,2}, Ehsan Allah-Kalteh³; *Aiuob Sofizadeh⁴

¹School of Health, Tehran University of Medical Sciences, Tehran, Iran

²Vector borne Diseases Research Center, North Khorasan University of Medical Sciences, Bojnurd, Iran

³Health Management and Social Development Research Center, Golestan University of Medical Sciences, Gorgan, Iran

⁴Infectious Disease Research Center, Golestan University of Medical Sciences, Gorgan, Iran

*Corresponding author: Dr Aiuob Sofizadeh, E-mail: a_sofizadeh@yahoo.com

(Received 9 Feb 2018; accepted 21 Jun 2021)

Abstract

Background: *Phlebotomus papatasi* is known as the main vector of zoonotic cutaneous leishmaniasis. This study aimed to investigate the effect of geographical and bioclimatic factors on the *Ph. papatasi* distribution.

Methods: A total of 34 villages were selected, and sampling was performed three times using 120 sticky traps in each selected village. All the collected species were mounted and identified their species. The densities of *Ph. papatasi* were measured in all the villages and entered into ArcMap as a point layer. The required bioclimatic and environmental variables were extracted from the global climate database and The normalized difference vegetation index was obtained from the MODIS satellite imagery, also, all variables entered into ArcMap as raster layers, so The numerical value of each independent variable in the cell where the selected village is located in this, was extracted using spatial analyst tools and the value to point submenu. All the data were finally entered into IBM SPSS, and the relationship was examined between the number of collected *Ph. papatasi* and the independent variables using Spearman's correlation test.

Results: A total of 1773 specimens of *Ph. papatasi* were collected. The findings of this study showed that max temperature of warmest month, temperature annual range, temperature seasonality, mean diurnal range, precipitation seasonality, mean temperature of driest and warmest quarter were positively associated with the density of *Ph. papatasi*.

Conclusion: Air temperature and precipitation were shown as the most significant factors in the distribution of *Ph. papatasi*.

Keywords: Ecology; Phlebotomine sand fly; GIS

Introduction

Leishmaniasis is one of the most important vector-borne parasitic diseases and appears as a significant health problem in Iran. Cutaneous and visceral leishmaniasis are two types of the disease in Iran. The cutaneous leishmaniasis (CL) is also prevalent in two forms of anthroponotic cutaneous leishmaniasis (ACL) and zoonotic cutaneous leishmaniasis (ZCL) in the country (1). Different studies conducted in Iran showed that vectors for ACL and ZCL were *Phlebotomus sergenti* and *Phlebotomus papatasi*, respectively (1-3).

In Golestan Province as one of the most important ZCL foci in Iran (4-8), *Leishmania major* is the agent and wild rodents such as *Rhombomys opimus* and *Meriones libycus* are reservoir hosts of ZCL (9-11). In a wide range of similar studies conducted in the province, *Ph. papatasi* was shown as the main ZCL vector and the infection range of *Ph. papatasi* to *L. major* was reported as 10% (11-12). Further, various studies conducted in the province showed the CL incidence equal to 31.7 per 100,000 people. among them, the two counties of Gonbad-e Kavus and

Maraveh Tappeh located in the northeast of the province had the highest incidence rates of 153 and 117 per 100,000 people (13). During 2010–2017, 360–1766 people from all age and gender groups in the province were infected with CL annually (14). Compared to the other regions of the province, the vectors and reservoirs of the disease are more abundant in the northeastern region of the province and the disease has higher incidence. These differences could be reasons for the geographical and climatic susceptibility of the region to a higher prevalence of the disease (13, 15).

Leishmaniasis is naturally dependent on environmental factors and climate change (16), and the environment plays a significant role in the transmission of the disease (17). Due to different species of agents and vectors, environmental changes can have various effects on the transmission of leishmaniasis in different regions of the world (18). Hence, environmental factors and climate change may significantly affect the growth, development, and distribution of *Ph. papatasi* as the main ZCL vector. Temperature and relative humidity are known as the most significant factors associated with the distribution of *Ph. papatasi* (19), and the climate change that has taken place in recent years has become a critical factor in the distribution of this species (17). It is also believed that the wide distribution of this species in the nature is dependent on environmental conditions and many of these conditions are measurable (20). Therefore, in recent years, many studies have been carried out on finding effective factors in the distribution of *Ph. papatasi* and several factors have been considered as essential factors in the distribution of this species. In a study conducted in Golestan Province (21), slope, altitude, annual mean temperature, and the normalized difference vegetation index (NDVI) were introduced as the most significant factors in the distribution of *Ph. papatasi*. Another study conducted in Iran (22) also reported the mean temperature of the wettest quarter, slope, precipitation seasonality, and the precipitation of

the wettest quarter as major factors associated with the distribution of this species. A study conducted in Eastern Mediterranean (23) revealed that land cover, mean temperature of coldest quarter, max temperature of warmest month, and min temperature of coldest month were the most critical factors for the distribution of *Ph. papatasi*. Further, in a study conducted in Libya (20), the altitude from the sea level was reported as a major factor associated with the distribution of *Ph. papatasi*. Given the significance of *Ph. papatasi* in the transmission of CL in Golestan Province, as well as the climate diversity of different regions in the province, the present study was carried out to examine the effect of different geographical and climatic factors (mentioned in the materials and methods section) on the distribution of *Ph. papatasi*.

Materials and Methods

Study area

Golestan Province is one of the 31 provinces of Iran, located in the north-east of the country, south of the Caspian Sea (53°57'–56°23' E, 36°30'–38°08' N) and makes approximately 1.3% of Iran's total area with a landmass of 20437.74 square km. This province is located between the three provinces of Mazandaran, Semnan and North Khorasan, is bordered by Turkmenistan from the north. The province is connected to the Caspian Sea from the east and to Alborz Mountains from the south. Weather conditions are largely diverse in various regions of the province. As such, one can experience mountainous climates in the south part of the province, arid and semi-arid climate in the north part, and mild Mediterranean climate in the western and central regions. In 2019, the province had maximum and minimum air temperatures of 40 and 20 mm, respectively, maximum and minimum relative humidity of 70 and 90%, respectively, and a rainfall amount of 333ml. It can be concluded that a wide range of suitable cli-

matic and geographical conditions are available for the development of various insect species in the province (24).

Sand fly collection

This analytical cross-sectional study was carried out from July to September 2014 by performing a three-time capture in a total of 34 villages (2–4 villages in each county). Sticky paper traps coated with Castor oil were used to collect sand flies. For each village, 60 indoor and 60 outdoor sticky paper traps were installed before the sunset and collected the following morning before the sunrise. All the collected sand flies were placed in acetone for two minutes and stored in 70% ethanol before transferring to the laboratory. In the laboratory, microscopic slides of the specimens were provided and the specimens were mounted in Puri's medium. Species of all the sand flies were determined using relevant morphological keys (25-26). The data were analyzed using Spearman's correlation test in IBM SPSS version 22.0.

Study variables

In this study, we used 22 geographical and climatic variable including: Alt: altitude from sea level (m) (Alt), Slope: slope in degrees obtained from altitude (%), BIO1: annual mean temperature (°C), BIO2: mean diurnal range (monthly mean (max temp-min temp)) (°C), BIO3: isothermality (BIO2/BIO7) ($\times 100$), BIO4: temperature seasonality (standard deviation $\times 100$), BIO5: max temperature of warmest month (°C), BIO6: min temperature of coldest month (°C), BIO7: temperature annual range (BIO5-BIO6) (°C), BIO8: mean temperature of wettest quarter (°C), BIO9: mean temperature of driest quarter (°C), BIO10: mean temperature of warmest quarter (°C), BIO11: mean temperature of coldest quarter (°C), BIO12: annual precipitation (mm), BIO13: precipitation of wettest month (mm), BIO14: precipitation of driest month (mm), BIO15:), BIO16: precipitation of wettest quarter (mm), BIO17: precip-

itation of driest quarter (mm), BIO18: precipitation of warmest quarter (mm), BIO19: precipitation of coldest quarter (mm), NDVI: normalized differentiated vegetation index.

Bioclimatic variables (n=19), environmental variables including altitude and slope, and NDVI were independent variables. The bioclimatic variables were obtained from the WorldClim global climate database (<http://www.worldclim.org/current>) at a spatial resolution of 1km². These variables were derived from long-term (1950–2000) monthly rainfall and temperature values for the development of significant biological variables. The environmental variables such as altitude and slope were obtained from a digital elevation model, and NDVI was obtained from MODIS satellite images in August, 2014. This study evaluated the relationship between the number of collected *Ph. papatasi* as the dependent variable and the independent variables. The data from the sampled villages were entered into ArcMap as a point layer, and each of the independent variables was entered into ArcMap as a raster layer. Then, each village layer was activated separately with an independent variable layer, by using spatial analyst tools and the value to point submenu, the numerical value of each independent variable in a cell, where the selected village was located, was extracted and entered into SPSS 22.0. Eventually, due to the non-normal distribution of densities of *Ph. papatasi* and the lack of linear regression assumptions, the relationship between the densities of *Ph. papatasi* with the bioclimatic variables, the environmental variables, and NDVI was examined using Spearman's correlation test.

Results

In this study, a total of 1773 *Ph. papatasi* were collected from 34 villages of Golestan Province, with the highest frequency in Gonbad-e-Kavus and Maraveh Tappeh counties (Table 1). The findings show that max temperature of warmest month (°C), temperature an-

nual range (BIO5-BIO6) (°C), temperature seasonality (\pm SD \times 100), mean diurnal range (average min/max temp) (°C), precipitation seasonality (coefficient of variation), mean temperature of driest quarter (°C), and mean temperature

of warmest quarter (°C) were positively associated with the density of *Ph. papatasi*, Moreover, the density of *Ph. papatasi* showed a positive inverse correlation with NDVI, but no correlation with altitude and slope (Table 2).

Table 1. Densities of *Phlebotomus papatasi* in different counties of Golestan Province, 2014

County	Number of selected villages	Number of collected <i>Ph. papatasi</i>	Number of collected <i>Ph. papatasi</i> for 60 traps
Maraveh Tapeh	3	396	132
Gonbad-e Kavus	4	564	141
Aqqala	2	114	57
Ramiyan	2	42	21
Gomishan	2	64	32
Aliabad-e Katul	3	156	52
Azadshahr	3	138	46
Gorgan	2	14	7
Kalaleh	3	261	87
Kordkuy	2	2	1
Bandar-e Gaz	2	0	0
Bandar-e torkman	2	10	5
Galikesh	2	6	3
Minudasht	2	6	3
Total	34	1773	52.1

Table 2. Bioclimatic and environmental variables with direct and positive correlation with densities of *Phlebotomus papatasi*

Correlation	Variables	Mean \pm SD*	Correlation Coefficient	P
Direct and positive	Number of collected <i>Ph. papatasi</i>	50.08\pm54.43	r= 0.747	P= 0.00
	Max temperature of warmest month (°C)	34.06\pm1.16		
	Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43	r= 0.529	P= 0.002
	Temperature annual range (max temp of warmest month–min temp of coldest month) (°C)	32.01\pm2.17		
	Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43	r= 0.523	P= 0.002
	Temperature seasonality (standard deviation \times 100)	7.66\pm0.56		
	Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43	r= 0.466	P= 0.006
	Mean diurnal range (mean of monthly (max temp–min temp)) (°C)	11.52\pm0.87		
	Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43	r= 0.465	P= 0.006
	Precipitation seasonality (coefficient of variation)	57.45\pm8.91		
	Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43	r= 0.449	P= 0.009
	Mean temperature of driest quarter (°C)	26.81\pm1.08		
	Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43	r= 0.387	P= 0.026
	Mean temperature of warmest quarter(°C)	26.97\pm0.87		
Inverse	Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43	r= - 0.345	P= 0.045
	NDVI	0.36\pm0.11		
	Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43	r= - 0.459	P= 0.007
	Annual precipitation (mm)	348.42\pm80.26		
Number of. collected <i>Ph. papatasi</i>	50.08\pm54.43			

Table 2. Continued ...

Direct and negative	Precipitation of driest month (mm)	6.78±4.41	r= - 0.497	P= 0.003
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43		
	Precipitation of driest quarter (mm)	23.93±13.56	r= - 0.552	P= 0.001
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43		
	Precipitation of warmest quarter (mm)	25.66±13.29	r= - 0.590	P= 0.00
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= - 0.348	P= 0.047
	Precipitation of coldest quarter (mm)	120.6±22.66		
Without correlation	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= 0.018	P= 0.925
	Altitude	208.48±312.79		
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= - 0.174	P= 0.331
	Slope	89.78±0.37		
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= - 0.127	P= 0.481
	Annual mean temperature (°C)	17.20±1.15		
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= 0.123	P= 0.495
	Isothermality (BIO2/BIO7) (×100)	3.54±0.09		
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= - 0.244	P= 0.164
	Mean temperature of wettest quarter (°C)	25.16±21.75		
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= - 0.323	P= 0.067
	Mean temperature of coldest quarter (°C)	7.60±1.88		
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= - 0.285	P= 0.108
	Precipitation of wettest month (mm)	60.06±13.27		
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= - 0.191	P= 0.287
	Precipitation of wettest quarter (mm)	145.15±25.53		
	Number of. collected <i>Ph. papatasi</i>	50.08±54.43	r= - 0.340	P= 0.053
Min temperature of coldest month (°C)	2.05±1.93			

*: standard deviation

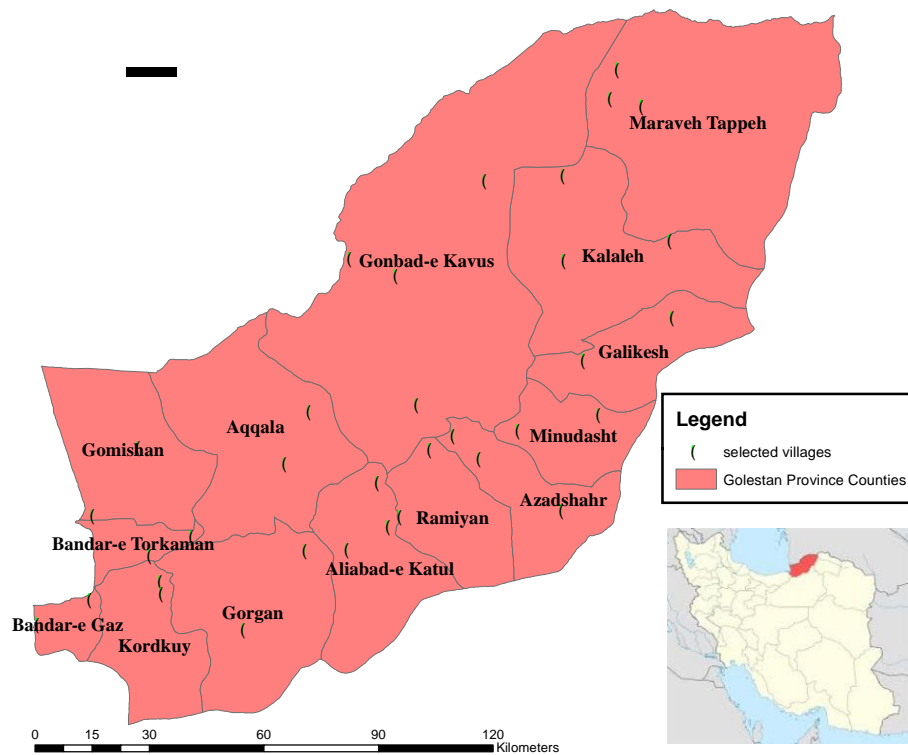


Fig. 1. Location of Golestan Province in Iran and collection sites for *Phlebotomus papatasi*

Discussion

The highest frequency of the collected *Ph. papatasi* was reported in Gonbad-e-Kavus and Maraveh Tappeh counties in accordance with results of a similar study conducted in Golestan Province, similarly reporting that the presence probability of this species was higher in these counties (21). These two counties are known as endemic focus for CL in Golestan Province (4-9). Moreover, the incidence of CL and the number of the rodent's active burrows (well-known reservoir hosts of ZCL) were higher in these regions than in the other regions of Golestan Province (7, 9, 15).

Temperature was shown as one of the main factors associated with the growth and development of insects such as *Ph. papatasi* (19). The results of this study revealed that temperature significantly affected the density of *Ph. papatasi*. The densities of *Ph. papatasi* were significantly positively associated with factors such as max temperature of warmest month (°C), temperature annual range (BIO5-BIO6) (°C), temperature seasonality (standard deviation $\times 100$), mean diurnal range (mean of max/min temp) (°C), mean temperature of driest quarter (°C), and mean temperature of warmest quarter (°C). However, there was no significant relationship between the densities of *Ph. papatasi* with annual mean temperature (°C), isothermality (BIO2/BIO7) ($\times 100$), mean temperature of wettest quarter (°C), mean temperature of coldest quarter (°C), and min temperature of coldest month (°C). It can be concluded that max and mean temperatures of driest and warmest quarters and months were significantly associated with the density of *Ph. papatasi*. However, mean temperature of wettest and coldest quarters and months had no significant role in the density of *Ph. papatasi*, which may be due to the effect of degree-day on the growth, development, and density of different species of insects. Degree-day is a measure of the amount of heat that accumulates above a specified temperature during a 24h period (27). Ac-

ording to results of studies on the growth of this species, a specific temperature limit is required for growth, and temperatures higher or lower than this limit will lead to a stop of growth. The maximum temperature threshold for the growth of this species is 35 °C, and the optimal minimum temperature for the growth according to different growth stages of this species is 20, 25, and 11.6 °C. As such, pre-oviposition activities are carried out at a temperature above 20 °C (8, 28-29). Therefore, temperature in the cold seasons, in which the air temperature does not reach 20 °C, cannot have a significant relationship with the density of this species. Colacicco-Mayhugh MG et al. (23) found that max temperature of warmest month was one of the most critical factors associated with the density of *Ph. papatasi* in Eastern Mediterranean, which is supported by the findings of our study. They also showed that mean temperature of coldest quarters and min temperature of coldest month were the most influential factors in the density of this species (23); while, in our study, there was no significant relationship between the values of these variables and the densities of *Ph. papatasi*. Further, the results of the present research showed no significant relationship between the densities of *Ph. papatasi* with isothermality (BIO2/BIO7) ($\times 100$) and mean temperature of wettest quarter (°C). These variables had low effects on the density of *Ph. papatasi* in another study conducted in Golestan Province (21). However, in a study conducted in Iran (22), contrary to the results of this study, isothermality (BIO2/BIO7) ($\times 100$) and mean temperature of wettest quarter (°C) showed high impacts on the density of this species.

In addition, we found no significant relationship between annual mean temperature and densities of *Ph. papatasi*. This result is in accordance with the results of a study in Iran (22), showing a relatively low correlation between this factor and the density of *Ph. papatasi*.

While in a study conducted in Golestan Province (21), this variable showed a significant effect on the density of this species. Differences between the results of these studies and those of our study may be due to differences in the nature of these studies. All of these studies applied the MaxEnt Model as an ecological Niche model, in which modeling and prediction are based on the existence of a species in a region (30) and quantity is not measured. However, the present study analyzed the effect of each variable quantity on the densities of *Ph. papatasi* and quantity had a direct effect on the study outcomes.

Precipitation was another main factor associated with the density of *Ph. papatasi*. The findings of the current study suggested that the densities of *Ph. papatasi* were positively associated with precipitation seasonality (coefficient of variation). In other words, the increased precipitation increased the density of sand flies, which is in accordance with the results of studies conducted in Iran and other countries (22, 31-32). Rainfall influences the dynamics, reproduction, and breeding of vectors such as sandflies (33-34), as sandflies need a certain amount of moisture for their development and survival. However, heavy rainfalls can kill adults and immature stages of sand flies (35-38).

We found no significant relationship between the density of *Ph. papatasi* with precipitation of wettest quarter and precipitation of wettest month. However, Hanafi-bojd et al. (22) and Rodgers et al. (39) revealed that precipitation of wettest quarter was a major factor associated with the density of *Ph. papatasi*. Colacicco-Mayhugh MG et al. (23) revealed that precipitation of wettest quarter and precipitation of wettest month were not significantly associated with the abundance of *Ph. papatasi*, which is consistent with our study. Perhaps the reason for this is that the wettest months and quarters of the year occur in winter and sand flies have no activity in this season; with the end of the wet months of the

year, the population of sand flies increases, as it was pointed out in another article (40).

Moreover, NDVI is another factor associated with the density of *Ph. papatasi*. The findings of our study revealed a significant reverse relationship between NDVI and the densities of *Ph. papatasi*, as this species is more abundant in areas with lower NDVI, which is in compliance with other studies conducted in this province and in Morocco County (15, 21, 41). Moreover, Abdel-dayme et al. (20) showed vegetation type as a major factor associated with the density of *Ph. papatasi*. Colacicco-Mayhugh MG (23) also introduced land cover as a significant factor associated with the density of *Ph. papatasi*. Moreover, Mollalo et al. (42) revealed a significant negative relationship between NDVI and the incidence of ZCL, as ZCL is more prevalent in areas with lower NDVI. Therefore, it can be concluded that NDVI is an essential factor in the incidence of ZCL.

Another factor associated with the density of *Ph. papatasi* is elevation. Accordingly, in a previous study conducted in Golestan Province, *Ph. papatasi* was collected at altitudes ranging from -32m from sea level to 598m above sea level, and densities of this species were more in plain areas with a lower altitude than in areas with a higher altitude (15). In a study conducted in Libya (20), it was observed that areas with an altitude lower than 600m were mostly suitable for the distribution of *Ph. papatasi*. Further, in a study conducted in Pakistan (43), altitude had the highest effect on the density of CL among various examined variables and *Ph. papatasi* was the dominant species in areas with a higher altitude. However, in this study, there was no significant relationship between altitude and the abundance of *Ph. papatasi*, which is inconsistent with previous studies.

Regarding slope as another factor associated with the density of *Ph. papatasi*, in different studies conducted in Golestan Province (21, 44) and Iran (22), slope was introduced as a major factor in the density of *Ph. pa-*

papatasi. However, the findings of the present study showed no significant relationship between slope and the densities of *Ph. papatasi*.

The density of sandflies might also depend on other environmental factors, such as soil type, land-use, or wind, which can impair their flight activity (38). In the present study, we could not assess the relationship between these factors and the density of *Ph. papatasi*, which could be one of the limitations of the study.

Conclusions

Air temperature and precipitation were shown as the most important factors in the density of *Ph. papatasi* in Golestan Province as the endemic focus of zoonotic cutaneous leishmaniasis in Iran.

Acknowledgements

This paper is part of a project approved by Infectious Diseases Research Center, Golestan University of Medical Sciences (grand No: 930708134). The authors would like to express their sincere gratitude to all staff and supervisors of this center who provided financial assistance and contributed to this thesis. The authors declare that they have no competing interests.

References

1. Yaghoobi-Ershadi MR (2012) Phlebotomine sand flies (Diptera: Psychodidae) in Iran and their role on *Leishmania* transmission. *J Arthropod Borne Dis*. 6: 1–17.
2. Parvizi P, Akhoundi M, Mirzaei H (2012) Distribution, fauna and seasonal variation of sandflies, simultaneous detection of nuclear internal transcribed spacer ribosomal DNA gene of *Leishmania major* in *Rhombomys opimus* and *Phlebotomus papatasi*, in Natanz District in central part of Iran. *Iran Biomed J*. 16(2): 113–120.
3. Karmaoui A (2020) Seasonal distribution of *Phlebotomus papatasi*, vector of zoonotic cutaneous leishmaniasis. *Acta Parasit*. 65: 585–598
4. Cherabin M, Sofizadeh A, Palideh AR, Gharavi AH, Gharavi M (2012) Epidemiological characteristics of cutaneous leishmaniasis in Maraveh tapeh District, Golestan Province during 2006–2010. *J Zabol Univ Med Sci*. 4(1): 19–27 (Persian).
5. Sofizadeh A, Faraji Far AA, Cherabin M, Badiei F, Cherabin M, Sarli J, Yapang Gharavi M, Mehravaran A (2013) Cutaneous leishmaniasis in Gonbad Kavoods, north of Iran (2009–2011): an epidemiological study. *J Gorgan Univ Med Sci*. 14(4): 100–106 (Persian).
6. Sofizadeh A, Ghorbani M, Gorganli Davaji A, Gharemeshk Gharavi A (2015) Epidemiological status of cutaneous leishmaniasis and ecological characteristics of sandflies in Maraveh-Tapeh County, Golestan Province, 2011–2012, Iran. *Qom Univ Med Sci J*. 9(6): 53–65 (Persian).
7. Sofizadeh A, Vatandoost H, Rassi Y, Hanafi-Bojd AA, Rafizadeh S (2016) Spatial analyses of relation between rodent's active burrows and incidence of zoonotic cutaneous leishmaniasis in Golestan Province, northeastern of Iran. *J Arthropod Borne Dis*. 10(4): 569–576.
8. Agh-Ataby MD, Sofizadeh A, Ozbaki GM, Malaki-Ravasan N, Ghanbari MR, Mozafari O (2016) Ecoepidemiological characteristics of a hypoendemic focus of zoonotic cutaneous leishmaniasis in north Iran (southeast of Caspian Sea). *J Vector Borne Dis*. 53: 248–256.
9. Rassi Y, Sofizadeh A, Abai MR, Oshaghi MA, Rafizadeh S, Mohebail M, Mohtaram F, Salahi R (2008) Molecular detection of *Leishmania major* in the vectors and reservoir hosts of cutaneous leishmaniasis in Kalaleh District, Golestan Prov-

- ince, Iran. Iran J Arthropod Borne Dis. 2(2): 21–27.
10. Akhoundi M, Mohebbali M, Asadi M, Mahmoodi MR, Amraei K, Mirzaei A (2013) Molecular characterization of *Leishmania* spp. in reservoir hosts in endemic foci of zoonotic cutaneous leishmaniasis in Iran. Folia Parasitol. 60(3): 218–224.
 11. Bordbar A, Parvizi P (2014) High density of *Leishmania major* and rarity of other mammals' *Leishmania* in zoonotic cutaneous leishmaniasis foci, Iran. Trop Med Int Health. 19(3): 355–363.
 12. Roshanghalb M, Parvizi P (2012) Isolation and determination of *Leishmania major* and *Leishmania turanika* in *Phlebotomus papatasi* main vector of zoonotic cutaneous leishmaniasis in Turkmen Sahra, Golestan Province. J Mazandaran Univ Med Sci. 21(Suppl 1): 74–83 (Persian).
 13. Mozafari O, Sofizadeh A, Shoraka HR, Namroodi J, Kalteh EA (2020) Eco-epidemiology of cutaneous leishmaniasis in Golestan Province, northeastern Iran: a systematic review. Jorjani Biomed J. 8(1): 60–78.
 14. Jorjani O, Mirkarimi K, Charkazi A, Dadban Shahamat Y, Mehrbakhsh Z, Bagheri A (2019) The epidemiology of cutaneous leishmaniasis in Golestan Province, Iran: A cross-sectional study of 8-years. Parasite Epidemiol Control. 5: e00099.
 15. Sofizadeh A, Rassi Y, Hanafi-Bojd AA, Shoraka HR, Mesgarian F, Rafizadeh S (2018) Distribution and ecological aspects of sand flies (Diptera: Psychodidae) species northeastern Iran. Asian Pac J Trop Med. 11(9): 526–533.
 16. Correa Antonialli S, Torres TG, Paranhos Filho AC, Tolezano JE (2007) spatial analysis of american analysis of american visceral leishmaniasis in Mato Grosso do Sul State, Central Brazil. J Infect. 54: 509–514.
 17. Azimi F, Shirian S, Jangjoo S, Ai A, Abbasi T (2017) impact of climate variability on the occurrence of cutaneous leishmaniasis in Khuzestan Province, Southwestern Iran. Geospat Health. 12(1): 15–22.
 18. Rodriguez-Morales AJ (2005) Ecoepidemiology and satellite epidemiology: new tools in the management of public health problems. Rev per Med Exper Salud Publ. 22: 54–63.
 19. Rassi Y, Hanafi-Bojd AA (2006) Sandflies, Vectors of Leishmaniasis. 1nd. Noavaran-e- Elm Publ, Tehran (Persian).
 20. Abdel-Dayem MS, Annajar BB, Hanafi-Bojd HA, Obenauer PJ (2012) The potential distribution of *Phlebotomus papatasi* (Diptera: Psychodidae) in Libya based on ecological niche model. J Med Entomol. 49(3): 739–745.
 21. Sofizadeh A, Rassi Y, Vatandoost H, Hanafi-Bojd AA, Mollalo A, Rafizadeh S, Akhavan AA (2017) Predicting the distribution of *Phlebotomus papatasi* (Diptera: Psychodidae), the primary vector of zoonotic cutaneous leishmaniasis, in Golestan Province of Iran using ecological niche modeling: comparison of MaxEnt and GARP Models. J Med Entomol. 54(2): 312–320.
 22. Hanafi-Bojd AA, Yaghoobi-Ershadi MR, Haghdoost AA, Akhavan AA, Rassi Y, Karimi A, Charrahy Z (2015) Modeling the distribution of cutaneous leishmaniasis vectors (Psychodidae: Phlebotominae) in Iran: a potential transmission in disease prone areas. J Med Entomol. 52(4): 557–565.
 23. Colacicco-Mayhugh MG, Masuoka PM, Grieco J (2010) Ecological niche model of *Phlebotomus alexandri* and *P. papatasi* (Diptera: Psychodidae) in the Middle East. Int J Health Geog. 9(2): 1–9.
 24. Ahmadpour M, Rezaei HR, Oshaghi MA, Hosseinzadeh Colagar A (2018) Modeling of the geographical distribution effects of great gerbil (*Rhombomys opimus*) on distribution of sandfly *Phlebotomus*

- papatasi* in Golestan Province. J Animal Environment. 9(4): 73–80 (Persian).
25. Theodor O, Mesghali A (1964) On the Phlebotominae of Iran. J Med Entomol. 1(3): 285–300.
 26. Seyedi-Rashti MA, Nadim A (1992) The genus *Phlebotomus* (Diptera: Psychodidae: Phlebotominae) of the countries of the Eastern Mediterranean Region. Iran J Publ Health. 21(1–4): 11–50.
 27. Zalom FG, Goodell PB, Wilson LT, Barnett WW, Bentley WJ (1983) Degree-days: The calculation and use of heat units in pest management. Division of Agriculture and Natural Resources, University of California, UC DANR Leaflet 21373, California.
 28. Kasap OE, Alten B (2005) Laboratory estimation of degree-day developmental requirements of *Phlebotomus papatasi* (Diptera: Psychodidae). J Vector Ecol. 30(2): 328–333.
 29. Oshaghi MA, Maleki Ravasan N, Javadian E, Rassi Y, Sadraei J, Enayati AA, Vatandoost H, Zare Z, Emami SN (2009) Application of predictive degree day model for field development of sandfly vectors of visceral leishmaniasis in north-west of Iran. J Vector Borne Dis. 46(4): 247–55.
 30. Elith J, Graham CH, Anderson RP, Dudik M, Ferrier S, Guisan A, Hijmans RJ, Huettmann F, Leathwick JR, Lehmann A (2006) novel methods improve prediction of species' distributions from occurrence data. Ecography. 29(2): 129–151.
 31. González C, Paz A, Ferro C (2014) Predicted altitudinal shifts and reduced spatial distribution of *Leishmania infantum* vector species under climate change scenarios in Colombia. Acta Trop. 129: 83–90.
 32. Queiroz MFM, Varjão JR, Moraes SC, Salcedo GE (2012) Analysis of sandflies (Diptera: Psychodidae) in Barra do Garças State of Mato Grosso, Brazil, and the influence of environmental variables on the vector density of *Lutzomyia longipalpis* (Lutz and Neiva, 1912). Rev Soc Bras Med Trop. 45(3): 313–317.
 33. Peterson AT, Shaw J (2003) *Lutzomyia* vectors for cutaneous leishmaniasis in southern Brazil: ecological niche models, predicted geographic distributions, and climate change effects. Int J Parasitol. 33(9): 919–931.
 34. Salomón OS, María Gabriela Quintana MG, Isolina Flores I, Ana María Andina AM, Silvia Molina S, Lucía Montivero L, Isabel Rosales I (2006) Phlebotominae sand flies associated with a tegumentary leishmaniasis outbreak, Tucumán Province, Argentina. Rev Soc Bras Med Trop. 39(4): 341–346.
 35. Kasap OE, Alten B (2006) Comparative demography of the sand fly *Phlebotomus papatasi* (Diptera: Psychodidae) at constant temperatures. J Vector Ecol. 31: 378–385
 36. Kasap OE, Alten B (2005) Laboratory estimation of degree-day developmental requirements of *Phlebotomus papatasi* (Diptera: Psychodidae). J Vector Ecol. 30: 328–333.
 37. Simsek FM, Alten B, Caglar SS, Ozbel Y, Aytekin AM, Kaynas S, Belen A, Kasap OE, Yaman M, Rastgeldi S (2007) Distribution and altitudinal structuring of phlebotomine sand flies (Diptera: Psychodidae) in southern Anatolia, Turkey: their relation to human cutaneous leishmaniasis. J Vector Ecol. 32: 285–291.
 38. Koch LK, Kochmann J, Klimpel S, Cunze S (2017) Modeling the climatic suitability of leishmaniasis vector species in Europe. Scientific Reports. 7(13325): 1–10.
 39. Rodgers MSM, Bavia ME, Fonseca EOL, Cova BO, Silva MMN, Trabuco Carneiro DDM, Cardim LL, Malone JB (2019) Ecological niche models for sand fly species and predicted distribution of *Lutzomyia longipalpis* (Diptera: Psychodidae) and visceral leishmaniasis in Ba-

- hia state, Brazil. Environ Monit Assess. 191(Suppl 2): 331.
40. Sherlock IA (1996) Ecological interactions of visceral leishmaniasis in the state of Bahia, Brazil. Mem Inst Oswaldo Cruz. 91: 671–683.
 41. Boussaa S, Kahime K, Samy A, Boumezough A (2016) Species composition of sand flies and bionomics of *Phlebotomus papatasi* and *P. sergenti* (Diptera: Psychodidae) in cutaneous leishmaniasis endemic foci, Morocco. Parasite Vectors. 9: 60.
 42. Mollalo A, Alimohammadi A, Shahrisvand M, Shirzadi MR, Malek MR (2014) Spatial and statistical analyses of the relations between vegetation cover and incidence of cutaneous leishmaniasis in an endemic province, northeast of Iran. Asian Pac J Trop Dis. 4(3): 176–180.
 43. Zaidi F, Fatima SH, Jan T, Fatima M, Ali A, Khisroon M, Adnan M, Rasheed SB (2017) Environmental risk modelling and potential sand fly vectors of cutaneous leishmaniasis in Chitral District: a leishmanial focal point of mount Tirich Mir, Pakistan. Trop Med Int Health. 22(9): 1130–1140.
 44. Mollalo A, Sadeghian A, Israel GD, Rashidi P, Sofizadeh A, Glass GE (2018) Machine learning approaches in GIS-based ecological modeling of the sand fly *Phlebotomus papatasi*, a vector of zoonotic cutaneous leishmaniasis in Golestan Province, Iran. Acta Trop. 188: 187–194.