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Climate change and potential distribution of zoonotic cutaneous leishmaniasis in Central Iran: Horizon 2030 and 2050

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ABSTRACT

Objective: To investigate and predict the effects of climate change on the potential distribution of the main vector and reservoir hosts of the disease in Yazd province in the future.

Methods: Distribution data for vector and reservoir hosts of zoonotic cutaneous leishmaniasis in Yazd province were obtained from earlier studies conducted in the area. MaxEnt ecological niche modeling was used to predict environmental suitability. BCC-CSM1-1(m) model and two climate change scenarios, RCP 4.5 and RCP 8.5 were used for horizons 2030 and 2050 climate projections. Future projections were based on data of a regional climate change model. Results: With both scenarios in 2030 and 2050, the results of jackknife test indicated that the mean temperature of wettest quarter and temperature annual range had the greatest effect on the model for the vector and the reservoir hosts, respectively.

Conclusions: The climate conditions are the major determinants of zoonotic cutaneous leishmaniasis incidence rate in Yazd Province. These climate conditions provide favorable habitats for ease transmission of zoonotic cutaneous leishmaniasis in this endemic area. Habitats suitability for the vector and reservoir will be expanding in the coming years compared with the current conditions, such that, in horizon 2030 & 2050, the probability of the presence of the vector and reservoir within 38 580 and 37 949 km², respectively, from Yazd province is above 60%. Moreover, an increase is predicted in the presence of the vector in the western parts and the reservoir in the northern and central parts of the province in the future. Understanding the role of environmental and bioclimatic factors in zoonotic cutaneous leishmaniasis occurrence can provide a guide for policy-makers in the creation and implementation of more effective policies for prevention and control.

1. Introduction

Leishmaniasis is a widespread vector-borne disease in the tropical

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and subtropical regions of zoonotic cutaneous leishmaniasis of the world with approximately two million new human cases reported each year[1,2]. It is estimated that more than 12 million people are infected worldwide[3]. Cutaneous leishmaniasis (CL) continues to

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be a major public health problem in Iran, with many endemic foci of zoonotic cutaneous leishmaniasis (ZCL) been reported in different parts of the country[4,5]. The sand fly Phlebotomus (P.) papatasi (Diptera: Psychodidae) is the principal vector of Leishmania major (Kinetoplastida: Trypanosomatidae), the causative agent of ZCL in the old world[6]. A number of gerbils (Rodentia: Gerbillidae) are known as reservoir hosts of ZCL in the old world, including Rhombomys (R.) opimus, Meriones libycus, Tatera (T.) indica and Meriones hurrianae which are more endemic in Iran[7]. The prevalence of ZCL has been reported in different provinces of Iran ranging from 1.8% to 37.9%[8,9]. Yazd province, located in the central part of Iran, is one of the major endemic foci of ZCL in the country. From 2007 to 2016, a total of 4229 cases have been formally reported in Yazd province[10]. Leishmania major is responsible for most of the ZCL cases reported in Yazd province, and has been isolated from various rodent reservoirs (R. opimus, Meriones libycus and T. indica) and sand fly vector (P. papatasi)[11].

Vector-borne diseases including ZCL are generally sensitive to climatic conditions[12]. It seems like that the incidence of human cases strongly correlates with the presence of sufficiently high densities of vector and reservoir species[13]. It has been suggested that climate change can cause a shift in the geographical spread of insect populations in the future through the modification of climatic conditions which affect the reproduction and length of annual activity of these insect species[14]. Different climatic conditions can cause species' range shifts, therefore, analyses of the distributional response of vector and reservoir species to climate change scenarios can provide insight into how the spatial epidemiology of ZCL may be affected by climate change[15].

Representative concentration pathway (RCPs) scenarios are a set of greenhouse gas concentration and emission pathways designed to support research on the impacts of climate and potential policymakers' response to climate change[16,17]. RCP scenarios provide interpolated climate layers based on the 2003 3rd assessment report of the intergovernmental panel on climate change and are publicly available[18]. These scenarios assume different climate futures based on greenhouse gases emission pathways and their associated radiative forcing to the year 2100[18]. In total, a set of four RCPs have been developed and are named according to the change in radiative forcing by the end of the century: 8.5, 6, 4.5 and 2.6 W/m².

Ecological niche models (ENMs) based on point occurrence data, digitized environmental layers, and machine learning algorithms, which are usually all overlaid on a geographic information system (GIS) platform, provide a better understanding of the distribution of vector-borne diseases[19,20]. ENMs attempt to predict the fundamental niche of a species, defined as the set of environmental conditions in which it can maintain populations without the need for immigration[21]. The fundamental niche provides the potential distributional pattern of a species when projected to a geographical space[22]. Effects of geographical constraints on dispersal and negative

ecological interactions such as competition may prevent a species from occupying its entire fundamental niche[23]. Thus, taking such factors into consideration defines the actual geographical distribution of a species. Practically, ENMs combine both the ecological requirements and spatial locations of species and predict the occurrence of species in an area between the potential and actual distributions[24]. Modification of these distributions using the constraining factors mentioned above generates the actual distributions. ENMs can thus provide useful framework for testing hypotheses concerning the role of different environmental variables in determining species' distributional patterns[25]. Estimating the potential future distributions of vector and reservoir species can help to identify potential risk areas for ZCL. The aim of this study was to investigate the effect of climate change on the potential distribution of ZCL vector and reservoir species in Central Iran in horizon 2030 and 2050.

2. Material and methods

2.1. Vector and reservoir data

Existing records from all previous studies conducted on *P. papatasi*, the main vector of ZCL, in Yazd province were collected and entered into a database. Previous studies in the area have reported *R. opimus* as the main reservoir host of ZCL in the province. Coordinates of collection sites for this gerbil were also recorded in the database.

2.2. General circulation models

The results of the research conducted by National Climate Research Institute[26] were the basis for selecting General Circulation Models in this study. BCC-CSM1-1(m) model at a spatial resolution of 30 s (1 km²) was used in our analysis. The Beijing Climate Center Climate System Model version 1.1 (BCC_CSM1.1) was used to develop the BCC_CSM1.1 (m) model with a moderate resolution[27]. In the present study, two scenarios were used for modeling: RCP4.5 and RCP8.5. In the RCP4.5 emission scenario, CO2 concentration is estimated to be 650 PPM by 2100, with a radiative forcing level of 4.5 watts per square meter[28,29]. Global Change Assessment Model team at the Joint Global Change Research Institute (a branch of the Pacific Northwest National Laboratory) in the United States developed the RCP 4.5 scenario[30]. The RCP 4.5 scenario is considered as a stabilization scenario, without an overshoot, in which total radiative forcing is stabilized shortly after 2100[31]. RCP 8.5 scenario corresponds to the highest greenhouse gas emission trajectory compared to the scenario literature[32], and hence also to the upper bound of the RCPs. The greenhouse gas concentrations and emission trajectories in RCP 8.5 is estimated to increase considerably over time, leading to a radiative forcing level of 8.5 W/m² at the end of the century with a temperature range between 3.5-4.5 $^{\circ}$ C[33]. The bioclimatic data based on RCP 4.5 and RCP 8.5 scenarios for the years 2030 and 2050 were downloaded from www.ccafs.cgiar.org (http://ccafs-climate.org/data) and www. worldclim.org (http://www.worldclim.org/cmip5_30s), respectively, at resolution 30 s.

2.3. Modeling potential occurrence of sand flies and gerbils

MaxEnt software Ver. 3.3.3 was used to predict the most appropriate ecological niches for the target species[34]. The contribution of the environmental variables was calculated by Jackknife analysis. Variables with no contribution were omitted from the final analysis. Eighty percent of the vector/reservoir collection points were used for model training and the remaining 20% were used as Test-set. MaxEnt was used for random point's selection. ESRI ArcGIS 10.3 software was used for the preparation, management and editing of spatial and climatic data, modeling, and presentation of the model results. The management of climatic data and preparation of the expressions for modeling were assisted by Microsoft Excel 2010 software.

3. Results

3.1. Modeling for ZCL vector and reservoir hosts in Horizon 2030: RCP 4.5 scenario, BCC-CSM1-1(m) model

The estimates of relative contributions of the environmental variables to the MaxEnt model for the vector and reservoir hosts of ZCL are shown in Table 1. The variables with the highest contributions to the models for *P. papatasi* and *R. opimus* were bio8 (73.5%) and bio7 (50.4%), respectively. Area under curve (AUC)

for *P. papatasi* was calculated as 0.917. The results of jackknife test of variable importance for *P. papatasi* show that the environmental variable with highest gain when used in isolation is bio8, which therefore appears to have the most useful information by itself. Bio8 is the environmental variable that decreases the gain the most when it is omitted, which therefore, appears to have the most information that is not present in the other variables (Figure 1). The AUC for *R. opimus* was calculated as 0.851. Jackknife test of variable importance for *R. opimus* revealed that the environmental variable with highest gain when used in isolation is bio7, which therefore appears to have the most useful information by itself. The environmental variable that decreases the gain the most when it is omitted is bio7, which therefore appears to have the most information that isn't present in the other variables (Figure 2).

3.2. Modeling for ZCL vector and reservoir hosts, Horizon 2030: RCP 8.5 scenario, BCC-CSM1-1(m) model

As shown in Table 1, the variables with the highest contributions to the models for *P. papatasi* and *R. opimus* were bio8 (38.4%) and bio7 (93.8%), respectively. The AUC for *P. papatasi* was calculated as 0.889 (Figure 3). The results of jackknife test of variable importance for *P. papatasi* show that the environmental variable with highest gain when used in isolation is bio8, which therefore appears to have the most useful information by itself. The environmental variable that decreases the gain the most when it is omitted is bio8, which therefore appears to have the most information that isn't present in the other variables (Figure 3). The AUC for *R. opimus* was calculated as 0.794 (Figure 4). Jackknife test of variable importance for *R. opimus* revealed that the environmental variable with highest gain when used in isolation is bio7, which therefore appears to have the most useful information by itself. The environmental variable

Table 1. Contribution of variables used for the prediction of the potential distribution of *Phlebotomus papatasi* and *Rhombomys opimus* as vector and reservoir hosts of zoonotic cutaneous leishmaniasis in Yazd Province, Central Iran, horizon 2030: RCP 4.5 and 8.5 scenarios, BCC-CSM1-1(m) model (%).

Variables	Description	Phlebotomus papatasi		Rhombomys opimus	
	Description	RCP4.5	RCP8.5	RCP4.5	RCP8.5
Bio1	Annual mean temperature ($^{\circ}$ C)	0.0	0.0	0.0	0.0
Bio2	Mean diurnal range [Mean of monthly (max temp-min temp)] (°C)	4.6	6.1	41.0	5.1
Bio3	Isothermality (Bio2/Bio7) (×100)	2.3	8.3	0.0	0.0
Bio4	Temperature seasonality (standard deviation $\times 100$)	0.0	1.7	0.0	0.0
Bio5	Max temperature of warmest month (°C)	0.0	0.0	0.0	0.0
Bio6	Min temperature of coldest month ($^{\circ}$ C)	0.8	0.0	0.0	0.0
Bio7	Temperature annual range (Bio5-Bio6) (℃)	0.0	0.0	50.4	93.8
Bio8	Mean temperature of wettest quarter ($^{\circ}$ C)	73.5	38.4	0.0	0.0
Bio9	Mean temperature of driest quarter ($^{\circ}$ C)	1.7	0.0	0.0	0.0
Bio10	Mean temperature of warmest quarter (°C)	0.0	0.0	0.0	0.0
Bio11	Mean temperature of coldest quarter (°C)	0.2	10.8	0.0	1.0
Bio12	Annual precipitation (mm)	6.5	4.9	0.0	0.0
Bio13	Precipitation of wettest month (mm)	0.0	4.7	0.0	0.0
Bio14	Precipitation of driest month (mm)	0.5	0.0	0.0	0.0
Bio15	Precipitation seasonality (Coefficient of variation)	4.7	2.8	8.5	0.1
Bio16	Precipitation of wettest quarter (mm)	0.0	0.0	0.0	0.0
Bio17	Precipitation of driest quarter (mm)	2.8	0.0	0.0	0.0
Bio18	Precipitation of warmest quarter (mm)	0.8	16.3	0.0	0.0
Bio19	Precipitation of coldest quarter (mm)	1.5	5.9	0.0	0.0

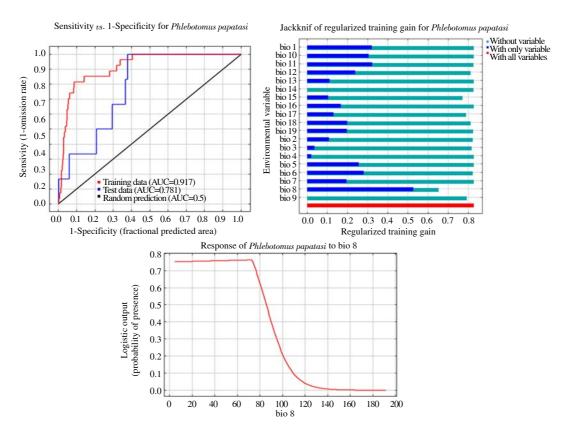


Figure 1. Receiver operating characteristic curve and output of MaxEnt model and jackknife analysis for *Phlebotomus papatasi* in Yazd province, Central Iran, Horizon 2030: RCP 4.5 scenario, BCC-CSM1-1(m) model.

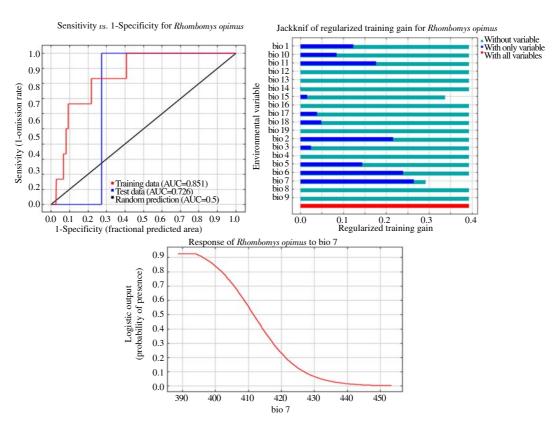


Figure 2. The receiver operating characteristic curve and output of MaxEnt model and jackknife analysis for *Rhombomys opimus* in Yazd province, Central Iran, Horizon 2030: RCP 4.5 scenario, BCC-CSM1-1(m) model.

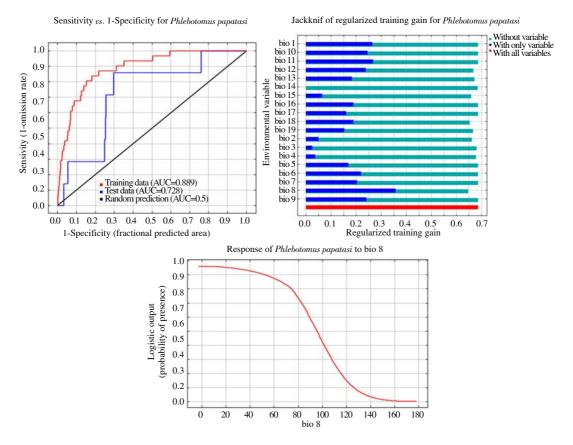


Figure 3. The receiver operating characteristic curve and output of MaxEnt model and jackknife analysis for *Phlebotomus papatasi* in Yazd province, Central Iran, Horizon 2030: RCP 8.5 scenario, BCC-CSM1-1(m) model.

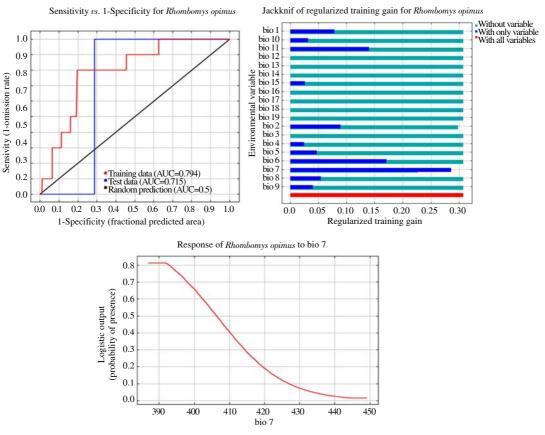


Figure 4. The receiver operating characteristic curve and output of MaxEnt model and jackknife analysis for *Rhombomys opimus* in Yazd province, Central Iran, Horizon 2030: RCP 8.5 scenario, BCC-CSM1-1(m) model.

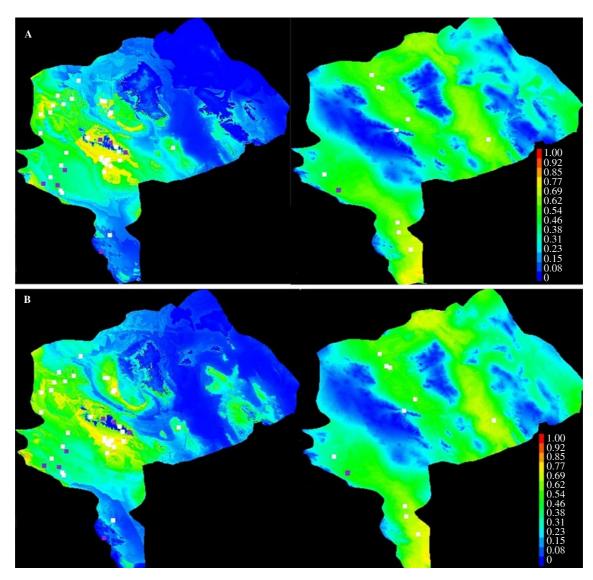


Figure 5. A: Predicted environmental suitability for vector (left) and reservoir (right) of ZCL in Yazd province of Iran using RCP 4.5 scenario, BCC-CSM1-1(m) model: Horizon 2030. B: Predicted environmental suitability for vector (left) and reservoir (right) of ZCL in Yazd province of Iran using RCP 8.5 scenario, BCC-CSM1-1(m) model: Horizon 2030.

that decreases the gain the most when it is omitted is bio7, which therefore appears to have the most information that isn't present in the other variables (Figure 4). Environmentally suitable areas for the vector and reservoir hosts in horizon 2030 using RCP 4.5 and RCP 8.5 scenarios have been presented in Figure 5.

3.3. Modeling for ZCL vector and reservoir hosts, Horizon 2050: RCP 4.5 scenario, BCC-CSM1-1(m) model, P. papatasi

Table 2 shows estimates of relative contributions of the environmental variables to the MaxEnt model for the vector and reservoir hosts of ZCL. As shown below, the variables with the highest contributions to the models for *P. papatasi* and *R. opimus* were bio8 (61.1%) and bio7 (50.4%), respectively. The AUC for *P. papatasi* was calculated as 0.903 (Figure 6). Analysis of Jackknife test of variable importance for *P. papatasi* shows that the environmental variable with highest gain when used in isolation is

bio8, which therefore appears to have the most useful information by itself. The environmental variable that decreases the gain the most when it is omitted is bio8, which therefore appears to have the most information that isn't present in the other variables (Figure 6).

The AUC for *R. opimus* was calculated as 0.849 (Figure 7). Jackknife test of variable importance for *R. opimus* revealed that the environmental variable with highest gain when used in isolation is bio7, which therefore appears to have the most useful information by itself. The environmental variable that decreases the gain the most when it is omitted is bio2, which therefore appears to have the most information that isn't present in the other variables (Figure 7).

3.4. Modeling for ZCL vector and reservoir hosts, Horizon 2050: RCP 8.5 scenario, BCC-CSM1-1(m) model, P. papatasi

Estimates of relative contributions of the environmental variables to the MaxEnt model for the vector and reservoir hosts of ZCL have

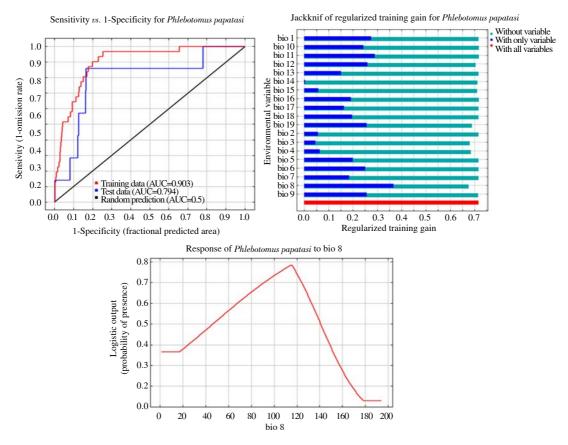


Figure 6. The receiver operating characteristic curve and output of MaxEnt model and jackknife analysis for *Phlebotomus papatasi* in Yazd province, Central Iran, Horizon 2050: RCP 4.5 scenario, BCC-CSM1-1(m) model.

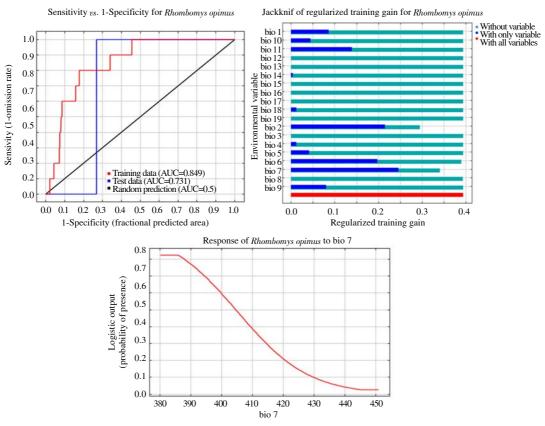


Figure 7. The receiver operating characteristic curve and output of MaxEnt model and jackknife analysis for *Rhombomys opimus* in Yazd province, Central Iran, Horizon 2050: RCP 4.5 scenario, BCC-CSM1-1(m) model.

been presented in Table 2. As shown in the Table 2, the variables with the most contributions to the models for *P. papatasi* and *R. opimus* were bio8 (53.0%) and bio7 (60.6%), respectively. The AUC curve calculated for *P. papatasi* was 0.891 (Figure 8). Analysis of jackknife test of variable importance for *P. papatasi* showed that the environmental variable with highest gain when used in isolation is bio8, which therefore appears to have the most useful information by itself. The environmental variable that decreases the gain the most when it is omitted is bio8, which therefore appears to have the most information that isn't present in the other variables (Figure 8).

The AUC curve calculated for *R. opimus* was 0.839 (Figure 9). Jackknife test of variable importance for *R. opimus* showed that the environmental variable with highest gain when used in isolation is bio7, which therefore appears to have the most useful information by itself. The environmental variable that decreases the gain the most when it is omitted is bio7, which therefore appears to have the most information that isn't present in the other variables (Figure 9). The environmentally suitable area for both vector and reservoir in horizon 2030 using RCP 4.5 and RCP 8.5 scenarios is presented in Figure 10.

Table 2. Contribution of variables used to predict the potential distribution of *Phlebotomus papatasi* and *Rhombomys opimus* as vector and main reservoir of ZCL in Yazd Province, Central Iran, horizon 2050: RCP4.5 and RCP8.5 scenarios, BCC-CSM1-1 (m) model.

Variables	Phlebotomus papatasi		Rhombomys opimus		
variables	RCP4.5	RCP8.5	RCP4.5	RCP8.5	
Bio1	2.1	1.7	0.0	0.0	
Bio2	0.2	0.5	48.7	35.6	
Bio3	6.5	3.5	0.0	0.0	
Bio4	10.4	6.7	0.0	0.0	
Bio5	0.0	0.0	0.0	0.0	
Bio6	0.0	0.2	0.7	3.8	
Bio7	0.0	0.0	50.4	60.6	
Bio8	61.1	53.0	0.2	0.0	
Bio9	0.6	1.0	0.0	0.0	
Bio10	0.0	0.0	0.0	0.0	
Bio11	0.0	2.8	0.0	0.0	
Bio12	5.4	0.8	0.0	0.0	
Bio13	0.3	1.5	0.0	0.0	
Bio14	1.1	1.3	0.1	0.0	
Bio15	1.1	18.1	0.0	0.0	
Bio16	1.3	1.7	0.0	0.0	
Bio17	0.2	1.9	0.0	0.0	
Bio18	2.3	1.5	0.0	0.0	
Bio19	7.4	3.8	0.0	0.0	

3.5. Climatic suitability for vector and reservoir hosts of ZCL in Yazd province

The results of the projected (2030 & 2050) climatic suitability for the two species (*P. papatasi* and *R. opimus*) are shown in Table 3. Western and central part of Yazd province will provide more suitable climatic conditions for *P. papatasi* and *R. opimus* to the end of the horizon 2030 and 2050.

Table 3. Projected climatic suitability (%) and the area (km²) covered by different classes for *Phlebotomus papatasi* and *Rhombomys opimus* in Yazd province, center of Iran.

Study period	Suitability	Phlebotomus papatasi		Rhombon	Rhombomys opimus		
Study period		RCP 4.5	RCP 8.5	RCP 4.5	RCP 8.5		
2030s	0-20	55 314	52 904	24 965	22 571		
	21-40	25 108	23 637	30 432	34 626		
	41-60	13 519	15 600	34 848	32 359		
	61-80	6 974	8 680	10 712	11 403		
	81-100	43	137	0	0		
2050s	0-20	55 155	60 563	31 144	31 083		
	21-40	18 221	20 093	30 178	33 613		
	41-60	19 783	13 016	28 368	24 599		
	61-80	7 948	7 218	11 370	11 817		
	81-100	5	222	0	0		

4. Discussion

To investigate the potential changes in the status of ZCL vector and reservoir species in the climate zone of Yazd province in the future, the probability of the presence of *P. papatasi* and *R. opimus* in 2030 and 2050 were examined. Probability of the presence of the two studied species was grouped into five classes using equal intervals in ArcGIS. Areas with more than 60% presence probability of species were considered as areas with high potential of CL transmission. In Horizon 2030, by using RCP 4.5 scenario, we found that the presence probability of the vector within 7 017 km² from Yazd province is above 60%. In this scenario, the variable with the highest contribution to the models for *P. papatasi* was bio8. Variation in this variable from about 70 to 140 has had a negative trend in the presence of *P. papatasi*, and then it does not change. The value of AUC was 0.917, which indicates that the output of the model is valid.

RCP 4.5 scenario revealed a potential expansion in the habitat of *R. opimus* to 10 712 km² from Yazd province. Jackknife test of variable importance for *R. opimus* showed that bio7 has the maximum effect on the dispersion of this species. Variation in this variable from about 395 to 445 has had a negative trend in the presence of *R. opimus*, and then it does not change. The value of AUC was 0.851, which indicates that the output of the model is valid.

In Horizon 2030, RCP 8.5 scenario revealed a potential dispersion of *P. papatasi* within 8 817 km² from Yazd province at various probabilities. The results of jackknife test show that bio8 has the highest effect on the dispersion of this species. Variation in this variable from about 0 to 160 has had a negative trend in the presence of *P. papatasi*, and then it does not change. The calculated AUC was 0.889, which indicates that the output of the model is valid.

In this model, we observed a potential presence of *R. opimus* within 11 403 km² from Yazd province. Jackknife analysis for *R. opimus* showed that bio7 has the highest gain among the environmental variables. Variation in this variable from about 390 to 445 has had a negative trend in the presence of *R. opimus*, and then it does not

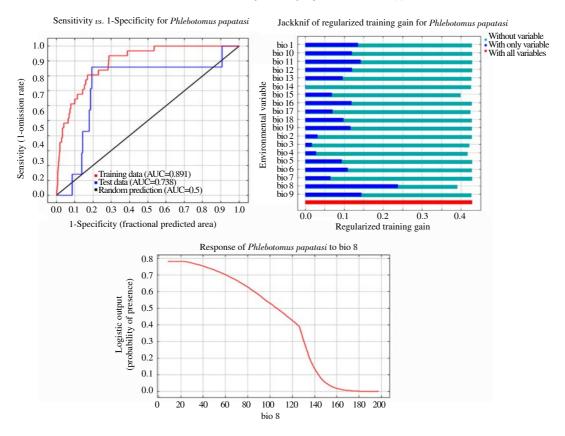


Figure 8. The receiver operating characteristic curve and output of MaxEnt model and jackknife analysis for *Phlebotomus papatasi* in Yazd province, Central Iran, Horizon 2050: RCP 8.5 scenario, BCC-CSM1-1(m) model.

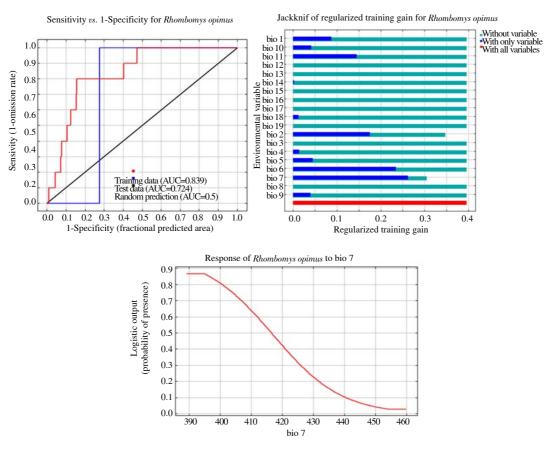


Figure 9. The receiver operating characteristic curve for *Rhombomys opimus* and output of MaxEnt model and jackknife analysis for *Rhombomys opimus* in Yazd province, Central Iran, Horizon 2050: RCP 8.5 scenario, BCC-CSM1-1(m) model.

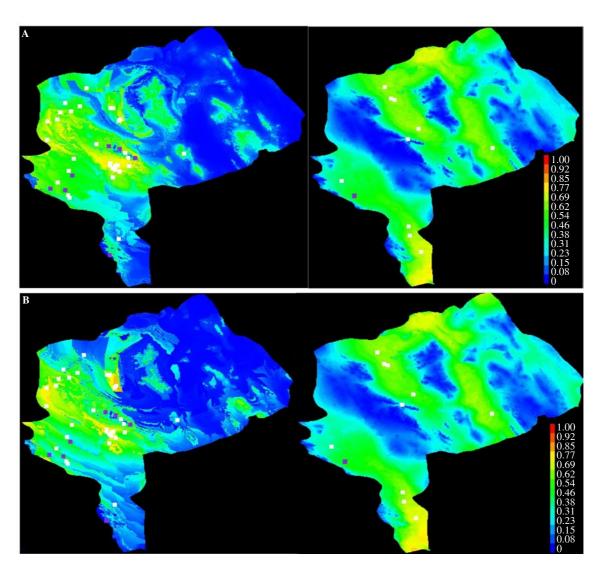


Figure 10. A: Predicted environmental suitability for vector and reservoir of zoonotic cutaneous leishmaniasis in Yazd province of Iran using RCP 4.5 scenario, BCC-CSM1-1(m) model, Horizon 2050: vector (left) and reservoir (right). B: Predicted environmental suitability for vector and reservoir of zoonotic cutaneous leishmaniasis in Yazd province of Iran using RCP 8.5 scenario, BCC-CSM1-1(m) model, Horizon 2050: Vector (left) and Reservoir (right).

change. The value of AUC was 0.794 indicating that the output of the model is valid. In Horizon 2030, in general, we observed a higher expansion of habitat suitability of ZCL vector and reservoir hosts in scenario 4.5 compared with scenario 8.5.

In Horizon 2050, RCP 4.5 scenario revealed a potential dispersion of the vector within 7 953 km² from Yazd province at various probabilities. The results of jackknife test of variable importance to the models for *P. papatasi* show that the environmental variable with highest gain when used in isolation is bio8. Variation in this variable from 20 to 115 has had a positive trend in the presence of *P. papatasi*, but in higher values up to about 175 has had a negative effect, and after that it does not change. The calculated value of AUC was 0.903, which indicates that the output of the model is valid. In this scenario, there is a potential presence of *R. opimus* within 11 370 km² from Yazd province. The results of jackknife test of variable importance for *R. opimus* show

that the environmental variable with highest gain when used in isolation is bio7. Variation in this variable from about 385 to 450 has had a negative trend in the presence of *R. opimus*, and then it does not change. The value of AUC was 0.849 indicating that the output of the model is valid.

In Horizon 2050, by using RCP 8.5 scenario, we observed that there is a potential presence of *P. papatasi* within 7 440 km² of Yazd province. The results of jackknife test of variable importance for *P. papatasi* indicate that the environmental variable with the highest gain when used in isolation is bio8. Variation in this variable from about 20 to 180 has had a negative trend in the presence of *P. papatasi*, and then it does not change. The value of AUC was 0.891, which indicates that the output of the model is valid.

In this Horizon, there is a potential dispersion of *R. opimus* within 11 817 km² of Yazd province. Jackknife test of variable importance

for *R. opimus* revealed that the environmental variable with the highest gain when used in isolation is bio7. Variation in this variable from about 395 to 445 has had a negative trend in the presence of *R. opimus*, and then it does not change. The calculated value of AUC was 0.839, indicating that the output of the model is valid.

According to the prediction of the model in the study area, in both 2030 and 2050 there is a decrease in the environmental suitability for $P.\ papatasi$, while the condition will increase the suitable areas for $R.\ opimus$, although the changes are not so big. Previous studies conducted in other countries revealed a significant relationship between CL and climatic factors. Toumi $et\ al$ in Sidi Bouzid reported that only average temperature was not a statistically significant predictor of ZCL incidence[35]. Contrary to our findings, in another study, Temperature Annual Range & Mean Temperature of Wettest Quarter (°C) had the greatest effect on the presence of ZCL vector and reservoir, respectively. The results of our study are consistent with the findings of other researchers around the world including Roger $et\ al$ in South America and Sing in Rajasthan, India. Both studies showed that the incidence of the disease increased with rise of temperature and decreased with decline of relative humidity[36,37].

In a study conducted by Yazdanpanah and Rostamianpur^[38] in Ilam Province in the West of Iran, they found a positive correlation between average temperature and CL incidence, whereas this correlation was negative in another study conducted by Mozafari and Bakhshizadeh^[39] in Yazd-Ardakan plain in central of Iran. In contrast to our findings, both studies conducted in Ilam and Yazd reported that the association between CL epidemic and relative humidity was not significant, whereas in our study, relative humidity appeared to be the most significant factor.

Climate factors are the major determinants of ZCL incidence rate in Yazd Province. These climate conditions provide the favorable habitats for propagation and transmission of ZCL in this endemic area. Habitat suitability for the vector and reservoir hosts in the coming years will be expanding such that in horizon 2030 and 2050 the probability presence of the vector and reservoir hosts in 38 580 and 37 949 km², respectively, from Yazd province is above 60%. Also, we predict an increase in the presence of the vector in the western parts and the reservoir in the northern and central parts of the province in the future. In the present study, we observed a complex relationship between environmental factors, bioclimatic factors and ZCL occurrence in Yazd province. Understanding the role of environmental and bioclimatic factors in ZCL occurrence can provide a guide for policy-makers in the creation and implementation of more effective policies for the prevention and control.

Conflict of interest statement

Authors declare no conflict of interest.

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References

- [1] Postigo JR. World Health Organization. Control in selected countries of the WHO Eastern Mediterranean and African regions. Casablanca Morocco; 2014 [Online]. Available from: https://www.who.int/ leishmaniasis/resources/9789241508773/en/. [Cited on 15th December 2018].
- [2] Carvalho BM, Rangel EF, Ready PD, Vale MM. Ecological niche modelling predicts southward expansion of *Lutzomyia* (Nyssomyia) flaviscutellata (Diptera: Psychodidae: Phlebotominae), vector of *Leishmania* (Leishmania) amazonensis in South America, under climate change. PLoS One 2015; 10(11): e0143282.
- [3] Loeuillet C, Bañuls AL, Hide M. Study of Leishmania pathogenesis in mice: Experimental considerations. *Parasite Vector* 2016; 9(1): 144.
- [4] Akhavan P, Karimi M, Pahlavani P. Risk mapping of cutaneous leishmaniasis via a fuzzy C means-based neuro-fuzzy inference system. Int Arch Photogramm Remote Sens Spatial Inf Sci 2014; XL-2/W3: 19-23. doi: 10.5194/isprsarchives-XL-2-W3-19-2014.
- [5] Shirzadi MR, Mollalo A, Yaghoobi-Ershadi MR. Dynamic relations between incidence of zoonotic cutaneous leishmaniasis and climatic factors in Golestan Province, Iran. J Arthropod Borne Dis 2015; 9(2): 148-160.
- [6] Salloum T, Khalifeh I, Tokajian S. Detection, molecular typing and phylogenetic analysis of *Leishmania* isolated from cases of leishmaniasis among Syrian refugees in Lebanon. *Parasite Epidemiol Control* 2016; 1(2): 159-168
- [7] Gholamrezaei M, Mohebali M, Hanafi-Bojd AA, Sedaghat MM, Shirzadi MR. Ecological niche modeling of main reservoir hosts of zoonotic cutaneous leishmaniasis in Iran. Acta Trop 2016; 160: 44-52.
- [8] Holakouie-Naieni K, Mostafavi E, Boloorani AD, Mohebali M, Pakzad R. Spatial modeling of cutaneous leishmaniasis in Iran from 1983 to 2013. Acta Trop 2017; 166: 67-73.
- [9] Nikonahad A, Khorshidi A, Ghaffari HR, Aval HE, Miri M, Amarloei A, et al. A time series analysis of environmental and metrological factors impact on cutaneous leishmaniasis incidence in an endemic area of Dehloran, Iran. Environ Sci Pollut Res Int 2017; 24(16): 14117-14123.

- [10]Shiravand B, Tafti AAD, Hanafi-Bojd AA, Almodarresi A, Mirzaei M, Abai MR. Modeling spatial risk of zoonotic cutaneous leishmaniasis in Central Iran. Acta Trop 2018; 185: 327-335.
- [11] Yaghoobi-Ershadi M, Jafari R, Hanafi-Bojd A. A new epidemic focuses of zoonotic cutaneous leishmaniasis in central Iran. *Ann Saudi Med* 2004; 24(2): 98-101.
- [12]Medlock JM, Leach SA. Effect of climate change on vector-borne disease risk in the UK. Lancet Infect Dis 2015; 15(6): 721-730.
- [13] Chalghaf B, Chlif S, Mayala B, Ghawar W, Bettaieb J, Harrabi M, et al. Ecological niche modeling for the prediction of the geographic distribution of cutaneous leishmaniasis in Tunisia. Am J Trop Med Hyg 2016; 94(4): 844-851
- [14]Boussaa S, Kahime K, Samy AM, Salem AB, Boumezzough A. Species composition of sand flies and bionomics of *Phlebotomus papatasi* and *P. sergenti* (Diptera: Psychodidae) in cutaneous leishmaniasis endemic foci, Morocco. *Parasite Vector* 2016; 9(1): 60.
- [15]Tlamçani Z, Er-Rami M. The current status of cutaneous leishmaniasis in Morocco. Türkiye Parazitol Dergisi 2014; 38(1): 5.
- [16] Van Vuuren DP, Kriegler E, O'Neill BC, Ebi KL, Riahi K, Carter TR, et al. A new scenario framework for climate change research: Scenario matrix architecture. Clim Change 2014; 122(3): 373-386.
- [17]O'Neill BC, Kriegler E, Riahi K, Ebi KL, Hallegatte S, Carter TR, et al. A new scenario framework for climate change research: The concept of shared socioeconomic pathways. Clim Change 2014; 122(3): 387-400.
- [18]Shrestha S, Bach TV, Pandey VP. Climate change impacts on groundwater resources in Mekong Delta under representative concentration pathways (RCPs) scenarios. *Environ Sci Policy* 2016; 61: 1-13.
- [19]Abdullah AYM, Dewan A, Shogib MRI, Rahman MM, Hossain MF. Environmental factors associated with the distribution of visceral leishmaniasis in endemic areas of Bangladesh: Modeling the ecological niche. Trop Med Health 2017; 45(1): 13.
- [20]Carvalho B, Rangel E, Vale M. Evaluation of the impacts of climate change on disease vectors through ecological niche modelling. *Bull Entomol Res* 2017; 107(4): 419-430.
- [21] Alvarado-Serrano DF, Knowles LL. Ecological niche models in phylogeographic studies: Applications, advances and precautions. *Molecular Ecol Resour* 2014; 14(2): 233-248.
- [22] Soberón J. Grinnellian and Eltonian niches and geographic distributions of species. *Ecol Lett* 2007; 10(12): 1115-1123.
- [23]Sánchez-Cordero V, Stockwell D, Sarkar S, Liu H, Stephens CR, Giménez J. Competitive interactions between felid species may limit the southern distribution of bobcats Lynx rufus. *Ecogr* 2008; 31(6): 757-764.
- [24]González C, Wang O, Strutz SE, González-Salazar C, Sánchez-Cordero V, Sarkar S. Climate change and risk of leishmaniasis in North America: Predictions from ecological niche models of vector and reservoir species. PLoS Negl Trop Dis 2010; 4(1): e585.
- [25]Sarkar S, Pressey RL, Faith DP, Margules CR, Fuller T, Stoms DM, et al. Biodiversity conservation planning tools: Present status and challenges for the future. Annu Rev Environ Resour 2006; 31: 123-159.

- [26]Ghahreman N, Tabatabaei M, Babaeian I. Investigation of uncertainty in the IPCC AR5 precipitation and temperature projection over Iran under RCP Scenarios. Poster on COP21-CMP11, Paris 2015. doi: 10.13140/ RG.2.1.1808.3683.
- [27]Kan M, Huang A, Zhao Y, Zhou Y, Yang B, Wu H. Evaluation of the summer precipitation over China simulated by BCC_CSM model with different horizontal resolutions during the recent half century. *J Geophys* Res Atmos 2015; 120(10): 4657-4670.
- [28]He C, Zhou T. Responses of the western North Pacific subtropical high to global warming under RCP4. 5 and RCP8. 5 scenarios projected by 33 CMIP5 models: The dominance of tropical Indian Ocean-tropical western Pacific SST gradient. J Climate 2015; 28(1): 365-380.
- [29]Knutson TR, Sirutis JJ, Zhao M, Tuleya RE, Bender M, Vecchi GA, et al. Global projections of intense tropical cyclone activity for the late twenty-first century from dynamical downscaling of CMIP5/RCP4. 5 scenarios. J Clim 2015; 28(18): 7203-7224.
- [30] Anderson TR, Hawkins E, Jones PD. CO₂, the greenhouse effect and global warming: From the pioneering work of Arrhenius and Callendar to today's earth system models. *Endeavour* 2016; 40(3): 178-187.
- [31] Lee JW, Hong SY, Chang EC, Suh MS, Kang HS. Assessment of future climate change over East Asia due to the RCP scenarios downscaled by GRIMs-RMP. Clim Dyn 2014; 42(3-4): 733-747.
- [32]O'Neill BC, Kriegler E, Riahi K, Ebi KL, Hallegatte S, Carter TR, et al. A new scenario framework for climate change research: the concept of shared socioeconomic pathways. Clim Change 2014; 122(3): 387-400.
- [33]Fann N, Nolte CG, Dolwick P, Spero TL, Brown AC, Phillips S, et al. The geographic distribution and economic value of climate changerelated ozone health impacts in the United States in 2030. J Air Waste Manag Assoc 2015; 65(5): 570-580
- [34]Radosavljevic A, Anderson RP. Making better Maxent models of species distributions: Complexity, overfitting and evaluation. *J Biogeogr* 2014; 41(4): 629-643.
- [35]Toumi A, Chlif S, Bettaieb J, Alaya NB, Boukthir A, Ahmadi ZE, et al. Temporal dynamics and impact of climate factors on the incidence of zoonotic cutaneous leishmaniasis in central Tunisia. *PloS Negl Trop Dis* 2012; 6(5): e1633.
- [36]Roger A, Nacher M, Hanf M, Drogoul AS, Adenis A, Basurko C, et al. Climate and leishmaniasis in French Guiana. Am J Trop Med Hyg 2013; 89(3): 564-569.
- [37]Singh KV. Studies on the role of climatological factors in the distribution of phlebotomine sandflies (Diptera: Psychodidae) in semi-arid areas of Rajasthan, India. J Arid Environ 1999; 42(1): 43-48.
- [38] Yazdanpanah HA, Rostamianpur M. Analysis of spatial distribution of leishmaniasis and its relationship with climatic parameters (case study: Ilam Province). Bull Environ, Pharmacol Life Sci 2013; 2(12): 80-86.
- [39]Mozafari G, Bakhshizadeh KF. Analysis of bioclima factors on the leishmaniasis diseases in Yazd-Ardakan plain. Geogr Develop 2011; 9(23): 185-202.