REVIEW ARTICLE



Neospora caninum infection in Iran (2004–2020): A review

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Abstract Knowledge on neosporosis and associated risk factors in different species of animals are so important for designing the control programs and reduce the economic losses globally. This literature review targeted for evaluating the infection rate of Neospora caninum in animals in Iran. Until April 2020, all of published documents in the main English and Persian-language databases were searched. A total number of 110 documents (English = 85 and Persian = 25) were extracted. Most of reports were seroepidemiological studies using ELISA in Iranian cattle population. The range of Neospora infection was 3.8-76.2% in cattle, 0-54.6% in dogs, 0.9-9.9% in sheep, 6.2% in goats, 19.2-55.9% in buffaloes, 20-42.2% in horses, 52% in donkeys, 3.2-27% in camels, 14% and 19% in cats, and 0-20.4% in rodents. This rate in birds was 17.3% in chicken, 9.8% and 30.4% in pigeons, 2.8% and 3.7% in sparrows, and 9.9% in hooded crows. This is a comprehensive literature review on Neospora infection in Iran at the first time. The infection of N. caninum is

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widespread in Iran especially in dogs and cattle population. This review can provide baseline information for future research. Study on other hosts especially on wild and exotic animals is recommended for exact estimate of neosporosis in Iran. Investigations into molecular diagnosis and genotyping of *N. caninum* strains are also needed; this will be helpful for developing vaccines and finding the connection among wild and domestic cycles of disease. Education on the risk factors associated with *N. caninum* infection for is suggested farmers and rural public.

Keywords *Neospora caninum* · Neosporosis · Animal · Bird · Risk factor · Review article · Iran

Introduction

Neospora caninum (Protozoa: Apicomplexa) is causative agent of neosporosis, a polysystemic parasitic disease with worldwide distribution (Dubey 2003). The parasite was isolated from five Boxer puppies with congenital encephalitis and myositis in Norway in 1984 for the first time (Bjerkas et al. 1984). Since then, *N. caninum* has been recognized as causative agent of abortion, stillbirth, and birth of weak neonates in several mammalian species, particularly cattle (Dubey et al. 2007).

Different species of canines such as domestic dogs (*Canis familiaris*), coyotes (*C. latrans*), dingoes (*C. domesticus*), and gray wolves (*C. lupus*) are definitive hosts for *N. caninum* (Dubey et al. 2005). Based on data from parasite isolation as well as DNA and antibody detection, a wide range of domestic and wild ruminants, equids, camels, rodents, and avian species can serve as intermediate host of this parasite (Dubey and Schares 2011).

Interestingly, dogs can be the definitive and intermediate host at the same time (Dubey et al. 2007).

Intermediate hosts become infected either via ingestion of oocysts with food and water (horizontal transmission), or via transplacental route (vertical transmission) (Dubey et al. 2005, 2007). In the intermediate host, oocysts transform in to motile and rapidly dividing tachyzoites, which subsequently change to bradyzoite which can localize in several tissues such as brain and muscles and form cysts. Upon consuming *Neospora*-tissue cysts by the definitive host, formation of oocysts in its intestine and its excretion in feces completes the life cycle (Dubey 2003).

In Iran, *N. caninum* infection was reported in dairy cattle from Mashhad, Northeast region at the first time (Sadrebazzaz et al. 2004). Since then, many epizootological studies have been conducted on various animals and avian species. However, there are no comprehensive reviews on the prevalence of *N. caninum* in Iran. Therefore, we aimed at collecting all the available information from 2004 to April 2020 to have a better picture of the situation in the country.

Materials and methods

Introduction to the study area: Iran

The country is located in western Asia between latitudes $24^{\circ}-40^{\circ}$ N, and longitudes $44^{\circ}-64^{\circ}$ E. It has an area of 1,648,195 km² with different climatic regions: (1) Mediterranean in north on the coast of the Caspian Sea which is covered by the lush lowland and the vast forests, (2) continental and arid in the central plateau, (3) cold in mountains of west and northwest, and (4) desert and hot in the southeast (Fig. 1).

Search criteria

All peer reviewed documents published from 2004 to April 2020 in English-language databases (Google Scholar, Science Direct, Scopus PubMed and Proquest) and Persianlanguage databases (SID, Magiran, Iran Medex, and Iran Doc) were searched. The search terms which were used alone and/or combined included "*Neospora*," "*Neospora caninum*," "neosporosis," "epidemiology," "prevalence," "anti-*Neospora* antibodies," "animals," "cattle," "abortion," "dog," "sheep," "goat," "horse," "donkey," "cat," "bird," "chicken," "pigeon," "sparrow," "rodent," "camel," and "buffalo," "Iran." All of the enlisted terms were searched in Persian language too. Exclusion criteria were articles that had no data or insufficient data about prevalence and/or infection rate of *Neospora*.

Results

A total of 110 documents including 85 English and 25 Persian articles and theses were reviewed. Most of the papers was on cattle (n = 53) and dogs (n = 25), respectively. The most of reports were sero-epidemiological studies using ELISA in Iranian cattle population. The range of *Neospora* infection was 3.8–76.2% in cattle, 0–54.6% in dogs, 0.9–9.9% in sheep, 6.2% in goats, 19.2–55.9% in buffaloes, 20–42.2% in horses, 52% in donkeys, 3.2–27% in camels, 14% and 19% in cats, and 0–20.4% in rodents. This rate in birds was 17.3% in chicken, 9.8% and 30.4% in pigeons, 2.8% and 3.7% in sparrows, and 9.9% in hooded crows. All of articles are summarized in Tables 1, 2, 3, 4, and 5 according to study regions, number and type of sample, method of examination, and gender.

Discussion

Neospora caninum in cattle

According to the latest report of Iranian Ministry of Agriculture (2019), overall cattle population in different regions of Iran is 1,202,000 in dairy farms, 4,616,000 in both beef and dairy farms and 2,271,000 of native breed in rural regions. The main raised breed is Holstein with a milk average production of 20 L per head/day. Also, the annually milk and meat production are estimated approximately 1,059,000 and 830 (×1000 Tons), respectively. The global median losses caused by abortions due to N. caninum are estimated of US\$ 1.3 billion annually (Reichel et al. 2013). Reichel et al. (2013) reported that 46.4 million cows were at annual risk of abortion. The economic losses were calculated to be US \$1,600.00 and US \$150.00 in the per dairy and beef farm, respectively (Reichel et al. 2013). Though many animals are involved in the life cycle of N. caninum, it is common between dog and cattle (Dubey et al. 2007). Most of Iranian researchers have concentrated on neosporosis in cattle; because of that's great impact on economic losses for farmers and also dairy industry. Vertical transmission is the most important infection route in cattle with frequency of 93.7% (Nicolino et al. 2017). Neosporosis in bovine has been related with endemic, epidemic and sporadic abortions, neonatal mortality, stillbirth, genitally infections, and decreasing of milk production (Guido et al. 2016). Most of the economic losses are associated with increase in the abortion rate, culling and decrease in the volume of milk production (Santos et al. 2016).

Wide range of infection is reported according to climate and geographic regions, type of samples (blood, milk,



Fig. 1 The map of Iran with different geographic and climate regions

semen, and abortion material), type of animals (race, healthy, pregnant, and antecedents of abortion), and laboratory diagnostic techniques (Table 1). The global prevalence of *N. caninum* in cattle was calculated 20% (95% CI 18–21%) by Ribeiro and colleagues (2019). They reported that seropositive cows were 1.6 times more likely to abort (OR = 2.6) than seronegative ones. In a meta-analyses study from Iranian dairy cattle, overall seroprevalence is estimated 23.6% in limit scale on eligibility articles until September 2018 (Ansari-Lari 2020). In addition, sero-prevalence rate showed significantly positive association with antecedents of abortion (OR = 2.5) and studied regions, opposite to age and breed.

Dairy cattle were considered more susceptible to bovine neosporosis (OR = 1.6) than beef cattle (Ribeiro et al. 2019). In Iran, the most reports of *N. caninum* infection were in dairy cattle with range of 3.8-76.2% (Binaei 2016; Noori et al. 2019). The lowest infection rate (3.8%) in Sistan is due to hot and dry climate in the region (Noori et al. 2019). Also, a high infection (57.3%) in North of Iran is related to moderate and rainy climate, which is favorite for sporulation of oocysts (Youssefi et al. 2010). This rate was 10-95.2% in milk samples (Shakerian et al. 2015; Binaei 2016) and 7.4-10.5% in semen samples (Sharifzadeh et al. 2012; Doosti et al. 2015). Bulk milk examination is using routinely for detection of antibodies to *N. caninum* in dairy farms due to ease of sampling, and the tests are cheap and rapid (Hurkova et al. 2005). 14.2% and

20% of native cattle were seropositive in Hamedan district, West of Iran (Gharekhani et al. 2012; Gharekhani and Heidari 2014a). In pregnant cows, seroprevalence of 24.8%, 30.4%, and 38.8% was reported in Hamedan, Shiraz, and Tehran, respectively (Salehi et al. 2010; Gharekhani and Yakhchali 2019). DNA of *N. caninum* is isolated in 11–66.7% of aborted fetus materials using molecular techniques (Habibi et al. 2005; Rafati and Jafarian 2014). The seropositivity in pregnant cows is a risk factor for increase the abortion. According to De Meerschman et al. (2002) report, risk of abortion had a direct correlation with level of *N. caninum*-specific antibodies in the animals. The abortion risk in seropositive animals is estimated 1.6–26-fold higher than seronegative animals (Ribeiro et al. 2019).

Impact of age on seropositivity was variable, according to studied regions and also the reasons were not clear because of the various opinions. Razmi et al. (2006) and Gharekhani et al. (2013a) reported a significant correlation between seropositivity and age of animals (P < 0.05). Youssefi et al. (2010) found a significant increase in seropositivity in 4–5-year-old animals. Sadrebazzaz et al. (2004) reported equal levels of seroprevalence in all age groups for the most of the herds. Gharekhani and Heidari (2014a, b) opined that the increase of seroprevalence with age is related to sample size. On other hand, no significant correlation was seen between seropositivity and age by Nourollahifard et al. (2008), Gharekhani and Heidari

Table 1 Infection rate of Neospora caninum in cattle population from different regions of Iran

Study area	Sample type	Methods	No. of sam	ple (positiv	ve %)	References	
			Total (%)	Male (%)	Female (%)		
Mashhad	Blood of dairy cattle	IFAT	810 (15.2)			Sadrebazzaz et al. (2004)	
Mashhad	Tissue of aborted fetuses	Semi-nested PCR	6 (66.7)	6 (66.7) NA		Habibi et al. (2005)	
Mashhad	Blood of dairy cattle	ELISA ^a	337 (46.3)			Razmi et al. (2006)	
Mashhad	Brain of aborted fetuses	PCR	100 (13)			Razmi et al. (2007)	
Mashhad	Sera and fluids of aborted fetuses	IFAT	12 (50)	NA	NA	Sadrebazzaz et al. (2007)	
	Brain of aborted fetuses	Semi-nested PCR	6 (33)				
Mashhad	Brain of aborted fetuses	PCR	151 (11.9)			Razmi et al. (2010)	
	fluids of aborted fetuses	ELISA ^a	151 (9.9)				
Mashhad	Blood of dairy cattle	ELISA ^a	200 (19)			Razmi et al. (2013)	
	Brain of aborted fetuses	PCR	200 (11.5)				
Mashhad	Blood of dairy cattle	ELISA ^b	116 (24.3)			Mikhchi et al. (2013)	
Mashhad	Bulk milk	ELISA ^c	123 (35)			Razmi and Barati (2017)	
Neishabour, Khorasan Razavi province	Blood of dairy cattle	ELISA ^c	250 (18)			Atashgahi (2015)	
Neishabour, Khorasan Razavi province	Blood	ELISA ^b	100 (26)	9 (22.22)	91 (26.3)	Nourollahifard et al. (2017)	
Torbatheydariyeh, Khorasan Razavi province	Blood of dairy cattle	ELISA ^c	280 (16.1)			Shahidi (2016)	
Kerman province	Blood	ELISA ^b	285 (12.6)	98 (10.2)	187 (13.9)	Nourollahifard et al. (2008)	
Shahrbabak, Kerman province	Blood of cattle	ELISA ^a	93 (12.9)	3 (0)	90 (13.3)	Asadi (2016)	
Sirjan, Kerman province	Blood of dairy cattle	ELISA ^c	150 (9.3)			Motamedipour (2016)	
Ahvaz	Blood	ELISA ^a	557 (21)	NA	NA	Hajikolaei et al. (2008)	
Golestan province	Blood of dairy cattle	ELISA ^a	800 (13.4)			Sattari et al. (2011)	
Babol, Mazandaran province	Blood of rural cattle	ELISA ^a	82 (43.9)	NA	NA	Youssefi et al. (2009)	
	Blood of industrial cattle		155 (25.8)				
	Total		237 (32)				
Ardabil	Blood of cattle with abortion	ELISA ^a	46 (7)			Youssefi et al. (2010)	
Garmsar, Semnan province	history		84 (45.2)				
Babol, Mazandaran province			96 (57.3)				
Moghan, Ardabil province	Different tissue of aborted fetuses	PCR	82 (41.5)			Hoseini et al. (2018)	
Garmsar, Semnan province	Blood of dairy cattle	ELISA ^a	104 (38.5)			Ranjbarbahadori et al. (2010)	
Semnan	Blood of dairy cattle	ELISA ^b	237 (76.2)			Binaei (2016)	
	Milk of dairy cattle		104 (95.2)				
Tehran	Blood of pregnant dairy cattle	ELISA ^a	768 (38.8)			Salehi et al. (2010)	
Tabriz	Blood of dairy cattle	ELISA ^a	266 (10.5)			Nematollahi et al. (2011a)	
Tabriz	Blood of cattle with abortion history	ELISA ^a	32 (20)			Nematollahi et al. (2011b)	
Tabriz	Blood of aborted cattle	ELISA ^a	76 (18.4)			Nematollahi et al. (2013)	

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Table 1 continued

Study area	Sample type	Methods	No. of samp	ole (positive	: %)	References	
			Total (%)	Male (%)	Female (%)		
	Brain of aborted fetuses	PCR	14 (42.8)				
East Azerbaijan province	Blood	ELISA ^b	236 (17.8)	74 (10.8)	162 (20.9)	Gharedaghi (2012a)	
Hamedan	Blood of native cow	ELISA ^a	400 (20)			Gharekhani et al. (2012)	
Hamedan province	Blood of dairy cattle	ELISA ^a	492 (12.8)	514	892 (16)	Gharekhani et al. (2014)	
	Blood of beef cattle Total		514 (19.8) 1406 (17.4)	(19.8)		Gharekhani and Tavoosidana (2013)	
						Gharekhani et al. (2013a)	
Hamedan province	Blood of aborted cattle	ELISA ^a	85 (61.2)			Gharekhani (2014)	
Hamedan province	Blood of pregnant dairy cattle	ELISA ^c	476 (24.8)			Gharekhani and Yakhchali (2019)	
Hamedan and Kurdistan	Blood of native cattle	ELISA ^a	768 (14.2)	NA	NA	Gharekhani and Heidari (2014a)	
Kurdistan province	Blood of slaughtered cattle	ELISA ^a	368 (7.8)	NA	NA	Heidari et al. (2014)	
Sanandaj	Blood of dairy cattle	ELISA ^a	336 (17.6)			Adhami et al. (2014)	
Kermanshah	Blood of cattle	ELISA ^c	92 (35.9)	NA	NA	Ghahvei (2014)	
Shahrekord	Blood of dairy cattle	ELISA ^c	184 (28.3)			Fooroghi-Pardenjani (2013)	
Shahrekord	Brain of aborted fetuses	Nested PCR	100 (11)			Rafati and Jaafarian (2014)	
Shahrekord	Milk of rural cattle	PCR	100 (10)			Shakerian et al. (2015)	
Shahrekord	Milk from rural native cattle	Nested PCR	100 (24)			Alipour et al. (2018)	
Lorestan province	Blood of dairy cattle	ELISA ^a	347 (9.8)			Nayebzadeh et al. (2015)	
Qom	Blood of dairy cattle	ELISA ^c	200 (9)			Javanshir (2015)	
Isfahan province	Blood of dairy cattle	ELISA ^a	611 (32.1)			Morovati and Noaman (2016)	
Isfahan province	Blood of dairy cattle	ELISA	1500 (26.3)			Hosseininejad et al. (2017)	
Kashan, Isfahan province	Milk from dairy cattle	ELISA ^c	187 (18.2)			Hadadi et al. (2018)	
Isfahan province	Blood of dairy cattle	ELISA ^a	216 (19)			Noaman and Nabinejad (2020)	
Shiraz	Blood of pregnant dairy cattle	ELISA ^a	253 (30.4)			Ansari-Lari et al. (2017)	
Fars province	Blood of dairy cattle	ELISA	180 (32.1)			Tavanaee and Namavari (2017)	
		NAT	180 (29.3)				
Shahriyar, Alborz province	Milk from dairy cattle	ELISA ^c	150 (22)			Taheri Lak et al. (2017)	
Arak	Brain of aborted fetuses	Nested PCR	38 (26.3)			Khani et al. (2018)	
Sistan region	Blood of dairy cattle	ELISA ^c	184 (3.8)			Noori et al. (2019)	
Different regions	Fresh semen of Iranian bulls	PCR	90 (9.7)			Sharifzadeh et al. (2012)	
	Frozen semen of Iranian bulls		85 (7.4)				
Different regions	Brain of aborted fetuses	PCR	395 (45)	NA	NA	Kamali et al. (2014)	
Different regions	frozen semen	PCR	57 (10.5)			Doosti et al. (2015)	

 $a = IDEXX^{(i)}$ Company; $b = Svanova Biotech^{(i)}$ Company; $c = ID-Vet^{(i)}$ Company

NA not available, PCR polymerase chain reaction, NAT Neospora agglutination test, IFAT indirect fluorescent antibody test, ELISA enzymelinked immunosorbent assay

(2014a) and Heidari et al. (2014). Some researchers reported low seroprevalence in cattle of > 2 yr, which they found were due to decrease of antibodies in vertically transmitted infection (Dubey et al. 2007).

The high rate of seropositivity was determined in rural cattle (20%) followed by beef (19.8%) and dairy cattle (12.8%) (P = 0.004) (Gharekhani et al. 2013a). Gharekhani and Tavoosidana (2013) stated that 21.5% and 18.5% of animals with Hybrid and Holstein breed were seropositive,

 Table 2 Infection rate of Neospora caninum in dogs in different regions of Iran

Study area	Sample type	Methods	No. of samp	le (positiv	References	
			Total (%)	Male (%)	Female (%)	
Tehran	Blood of farm dogs	IFAT	50 (46)	NA	NA	Malmasi et al. (2007)
	Blood of household dogs		50 (20)	NA	NA	
Tehran	Blood of farm dogs	IFAT	50 (28)	NA	NA	Haddadzadeh et al. (2007)
	Blood of urban dogs		53 (11.3)	NA	NA	
Tehran	Blood of stray dogs	ELISA ^b	42 (2.2)	NA	NA	Pouramini et al. (2017)
	Brain and other tissues	Nested PCR	42 (35.7)	22 (40.9)	20 (3)	
Mashhad	Stool of household dogs	PCR	85 (0)	NA	NA	Razmi (2009)
	Stool of farm dogs		89 (2.2)			
Chaharmahal-va Bakhtiari	Blood of stray dogs	IFAT	50 (46)	NA	NA	Hosseininejad et al. (2010a)
	Blood of household dogs		50 (20)	NA	NA	
	Total		100 (32)	NA	NA	
Isfahan	Blood of dogs referred to clinic	IFAT	233 (10.3)	NA	NA	Hosseininejad et al. (2010b)
Isfahan	Blood of farm dogs	IFAT	113 (17.7)	NA	NA	Hosseininejad et al. (2017)
Isfahan	Stool of shelter dogs	Nested PCR	90 (24.4)	NA	NA	Motamedi et al. (2020)
Chaharmahal-va Bakhtiari, Isfahan and Khuzestan	Blood of stray or shepherd dogs	ELISA	317 (43.3)	NA	NA	Hosseininejad and Hosseini (2011)
	Blood of household dogs		231 (8.7)	NA	NA	
	Total		548 (29.01)	300 (29.3)	248 (28.6)	
Chaharmahal-va Bakhtiari province	Blood of dogs	IFAT	200 (27.5)	NA	NA	Raeisi (2010)
Meshkin-Shahr (Ardabil)	Blood or rural dogs	ELISA	171 (30.4)	146 (30.8)	25 (28)	Sharifdini et al. (2011)
Urmia	Blood of stray dogs	IFAT	135 (26.6)	81 (29.6)	54 (22.2)	Yakhchali et al. (2010)
Sarab (East Azerbaijan)	Blood or rural dogs	IFAT	384 (10.7)	306 (10.1)	78 (12.8)	Khanmohammadi and Fallah (2011)
Tabriz	Blood of stray dogs	IFAT	100 (31)	63 (33.3)	37 (27)	Gharedaghi (2012b)
Tabriz	Stool of dogs	PCR	100 (21)	NA	NA	Fakheri (2019)
Shiraz	Blood of farm and household dogs	MAT	108 (44.4)	64 (37.5)	44 (52.3)	Khordadmehr et al. (2012)
		Dot- ELISA	108 (54.6)	64 (50)	44 (61.4)	
Hamedan	Blood of stray dogs	IFAT	200 (52.8)	NA	NA	Gharekhani et al. (2013a)
	Blood of shepherd dogs		70 (18)	NA	NA	
	Total		270 (27)	158 (23.4)	112 (32.1)	
	Blood of dairy farm dogs	ELISA ^b	185 (8.6)	56 (17.9)	129 (4.7)	Gharekhani and Yakhchali (2019)
	Blood of stray dogs	ELISA ^b	180 (5)	64 (7.8)	116 (3.5)	Gharekhani and Yakhchali (2019)
	Blood of pet dogs		184 (4.9)	79 (1.3)	105 (7.6)	Gharekhani et al. (2020)
Lorestan province	Stool of farm dogs	PCR	428 (2.1)	NA	NA	Dalimi et al. (2014)
Qom	Blood of farm dogs	ELISA ^b	50 (4)	NA	NA	Javanshir (2015)
Ahvaz	Blood of dogs referred to clinic	ELISA	100 (18)	NA	NA	Hosseininejad et al. (2018)

Table 2 continued

Study area	Sample type	Methods	No. of sample (positive %)			References	
			Total (%)	Male (%)	Female (%)		
Ahvaz	Blood of rural and urban dogs	NAT	150 (20)	85 (17)	65 (20)	Mosallanejad et al. (2018a)	
Different regions	Blood of stray dogs	IFAT	137 (12.4)	90 (16.7)	47 (4.3)	Yakhchali et al. (2017)	

 $b = ID-Vet^{\mathbb{R}}$ Company

NA not available, PCR polymerase chain reaction, NAT Neospora agglutination test, MAT modified agglutination test, IFAT indirect fluorescent antibody test, ELISA enzyme-linked immunosorbent assay

Animal	Study area	Sample type	Methods	No. of sam	ple (positiv	References	
type				Total (%)	Male (%)	Female (%)	
Sheep	Aleshtar, Lorestan province	Blood of aborted ewes	ELISA ^b	170 (1.1)			Ezatpour et al. (2013)
		Blood of healthy ewes		410 (1.7)			
		Total		586 (1.5)			
	East Azerbaijan province	Blood of aborted ewes	ELISA ^a	70 (5.7)			Asadpour et al. (2013)
		Fetal tissue	PCR	70 (8.5)			
	Mianeh, East Azerbaijan	Blood	ELISA ^c	317 (4.1)	NA	NA	Vajdi et al. (2014)
province Tehran	province		IFAT	317 (2.5)	(2.5)		
	Tehran	Healthy slaughtered sheep	Nested PCR		NA	NA	Arbabi et al. (2016)
		Heart		180 (6.7)			
		Brain		150 (0.7)			
		Total		330 (3.9)			
	Hamedan	Blood of aborted ewes	ELISA ^a	358 (2.2)			Gharekhani et al. (2013b)
	Golpayegan, Isfahan province	Blood of ewes	ELISA ^b	250 (6.8)			Shahi (2016)
	Mashhad	Fetal brain	PCR	71 (9.9)	NA	NA	Razmi and Naseri (2017)
	Khuzestan province	Blood	ELISA ^b	550 (6.8)	163 (6.1)	392 (7.1)	Gharekhani et al. (2018)
	Different provinces	Fetal brain	Semi- nested	109 (0.9)			Sasani et al. (2013)
			PCR				
Goats	Hamedan	Blood	ELISA ^b	450 (6.2)	54 (11.1)	396 (5.6)	Gharekhani et al. (2016)
	Khuzestan province	Blood	ELISA ^b	185 (10.8)	30 (16.7)	155 (9.7)	Gharekhani et al. (2018)

Table 3 In	fection rate	of Neospora	caninum	in sheep	o and	goats	from	different	regions	of	Iran
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 $a = IDEXX^{\text{(B)}}$ Company; $b = ID-Vet^{\text{(B)}}$ Company; $c = VMRD^{\text{(B)}}$ Company

NA not available, PCR polymerase chain reaction, IFAT indirect fluorescent antibody test, ELISA enzyme-linked immunosorbent assay

respectively (P > 0.05). The high infection in dairy cattle as against beef cattle is related to different production systems for dairy and beef cattle. As few studies have been conducted on the animals breed, planning and conducting extensive researches on the association between various breed and infection is recommended. Abortion is a predominant clinical sign of neosporosis in cattle (Dubey 2003). There is a direct association between the level of antibodies to *N. caninum* in cows and occurrence of abortion (De Meerschman et al. 2002). The risk of abortion due to neosporosis is estimated 14.3% in different regions (Reichel et al. 2013). In addition, the risk of

Animal type	Study area	Sample type	Methods	No. of sam	ple (Positiv	e %)	References	
				Total (%)	Male (%)	Female (%)		
Cat	Ahvaz	Blood (feral cat)	NAT	100 (19)	57 (19.3)	43 (18.6)	Hamidinejat et al. (2011a)	
	Ahvaz	Blood (domestic Persian cat)	NAT	100 (14)	52 (3.6)	48 (3.4)	Hamidinejat et al. (2011b)	
Rodent	Arak	Brain tissue of Mus musculus	PCR	68 (0)			Khani (2016)	
	Ahvaz	Blood of Rattus rattus	NAT	150 (6)	83 (6)	67 (5.9)	Mosallanejad et al. (2018b)	
	Meshgin-Shahr	Brain tissue	PCR	70 (0)	40 (0)	30 (0)	Nazari et al. (2019)	
	Meshgin-Shahr	Blood	IFAT	157 (20.4)	97 (33)	60 (38.3)	Nazari et al. (2020)	
Camel (Camelus	Mashhad	Blood (slaughterhouse)	IFAT	120 (5.8)	NA	NA	Sadrebazzaz et al. (2006)	
dromedarius)	Isfahan	Blood (slaughterhouse)	IFAT	310 (3.2)	NA	NA	Hosseininejad et al. (2009)	
	Yazd	Blood	NAT	254 (3.9)	208 (3.85)	46 (4.35)	Hamidinejat et al. (2013)	
	Bushehr	Blood	NAT	92 (27)			Namavari et al. (2017)	
Buffalo (Bubalus bubalis)	Ahvaz	Blood (slaughterhouse)	ELISA ^a	181 (37)	40 (15)	141 (43.3)	Hajikolaei et al. (2007)	
	Ahvaz	Blood (slaughterhouse)	ELISA ^a ELISA ^b NAT	188 (55.9)	121 (53.7)	67 (59.7)	Pourmahdi-Borujeni et al. (2015)	
	Urmia	Blood (slaughterhouse)	ELISA Nested	83 (19.2)	32 (3.6)	51 (15.6)	Rezvan et al. (2019)	
			PCR					
Horse (Equus caballus)	Tabriz	Blood	MAT	100 (28)	NA	NA	Gharedaghi (2011)	
	Shiraz	Blood	MAT	200 (40)	80 (45)	120 (55)	Moraveji et al. (2011)	
	Mashhad	Blood	MAT	150 (30)	NA (55)	NA (45)	Hosseini et al. (2011)	
	Hamedan	Blood of riding club animals	MAT	45 (42.2)	26 (57.9)	19 (42.1)	Gharekhani et al. (2013c)	
		Blood of rural animals	MAT	75 (40)	17 (52.9)	58 (36.2)		
		Total	MAT	120 (40.8)	43 (46.5)	77 (37.7)		
	Khuzestan province	Blood of Arab breed	MAT	235 (20)	122 (16.4)	113 (23.9)	Tavalla et al. (2015)	
Donkey (Equus africanus asinus)	Hamedan	Blood	MAT	100 (52)	22 (36.1)	78 (56.4)	Gharekhani et al. (2013c)	

Table 4	Infection rate	of Neospora	caninum (a	and/or i	Neospora sp.)) in cats,	camels,	buffaloes,	and e	equine	population	from	different	regions	of
Iran															

 $a = IDEXX^{\text{(B)}} Company; b = ID-Vet^{\text{(B)}} Company$

NA not available, PCR polymerase chain reaction, NAT Neospora agglutination test, MAT modified agglutination test, IFAT indirect fluorescent antibody test; ELISA Enzyme-linked immunosorbent assay

abortion in seropositive cows estimated 1.6–26 time higher than seronegative ones (Schares et al. 2004; Razmi et al. 2006; Ribeiro et al. 2019; Lefkaditis et al. 2020). Benavides et al. (2012) reported the risk of abortion in beef cattle is low compared to dairy cattle. Abortion rate associated with neosporosis in Iranian cows is reported between 7.8 and 66.7% (Sadrebazzaz et al. 2004; Razmi et al. 2006; Salehi et al. 2010; Nematollahi et al. 2011a; Gharekhani et al. 2014; Ansari-Lari et al. 2017). By attention to Ansari-Lari et al. (2017) idea, up to 30% of abortions can be eliminated by control of *N. caninum* infection in the herd level.

In a study by Gharekhani and Heidari (2014a), 26% of animals with history of abortion were seropositive (P < 0.0001 and odds ratio = 2.9). Youssefi et al. (2010) reported that 7%, 45.2% and 57.3% of aborted cattle were seropositive to *N. caninum* in Ardebil (Northwest of Iran, cold climate), Garmsar (Central of Iran, warm and dry climate) and Babol (North of Iran, rainy and mild climate), respectively. In a study from Northeast of Iran, the

 Table 5 Infection rate of Neospora caninum in birds from different regions of Iran

Birds type	Study area	Sample type	Methods	No. of sample (positive%)	References
Chicken (Gallus domesticus)	Shiraz	Blood	MAT	150 (17.3)	Sayari et al. (2016)
Pigeon (Columba livia)	Ahvaz	Blood	NAT	102 (30.4)	Bahrami et al. (2016)
		Brain tissue	PCR	102 (9.8)	
Sparrow (Passer domesticus)	Ahvaz	Brain tissue	PCR	210 (2.8)	Bahrami et al. (2015)
	Tehran	Brain tissue	Nested PCR	217 (3.7)	Abdoli et al. (2015)
hooded crows (Corvus cornix)	Tehran	Brain tissue	Nested PCR	55 (9.9)	Abdoli et al. (2018)

NA not available, PCR polymerase chain reaction, NAT Neospora agglutination test, MAT modified agglutination test

prevalence of abortion in seropositive animals was reported higher than in seronegative (P < 0.05, OR = 1.78) ones (Razmi et al. 2006). Also, in the report of Hosseininejad et al. (2017), infection rate was detected significantly high in animals with history of abortion. Recently, a prospective cohort investigation in the country revealed that approximately 30% of abortions may be managed by control of N. caninum (Ansari-Lari et al. 2017). This is so important that the time is right to initiate vigorous inclusive efforts for preventing economic losses due to pathogen in Iran. The complication of N. caninum co-infections with other pathogens related with abortions can increase the rate of abortion. Noaman and Nabinejad (2020) found that, N. caninum plus bovine herpesvirus type 1 (BHV-1) co-infection was 12% versus 6.9% of N. caninum infection without BHV-1. Also, N. caninum seroprevalence was 10.7% and 8.5% in plus and without bovine viral diarrhea virus (BVDV) infection, respectively.

Gharekhani et al. (2014) reported that seroprevalence rate was 17.2% and 20.9% in farms with the presence of dog and also stray canids such as fox and jackal, respectively. In dairy farms, the seroprevalence rate had significant connection with history of abortion in individual level (P = 0.009), stillbirth (P = 0.02), and metritis in animals (P = 0.004), while no significant correlation was seen between stage of gestation (P = 0.977) and number of gestations (P = 0.986) in pregnant cows and also herd size (P = 0.178) with seropositivity of animals. Additionally, 33.7% of seropositive animals had history of artificial insemination (P < 0.0001) (Gharekhani and Yakhchali 2019). The risk of infection may increase with gestation, suggesting that horizontal transmission of N. caninum was particular importance in some herds with the presence of dog in farm (Dubey et al. 2007). Gharekhani et al. (2014) reported that the infection in dairy herds with > 100 population was 2.7-fold higher than < 100 (P = 0.0005). The number of dogs in each farm can be in direct relation with the herd size (Dubey and Schares 2011). Implementation of modern management with preventive approach inside the farms, and also farmers training is significantly effective for reducing the national economic losses due to bovine neosporosis.

Neospora caninum in buffaloes

Buffaloes, while being more susceptible to subclinical infection, are also more tolerant to neosporosis, and the inflammatory responses to infection are mild. Seroprevalence appears to be much higher in buffalo as compared to both dairy and beef cattle. Even when cattle and buffalo cohabit, thus sharing the same environmental risk factors, buffalo appears to show much higher infection rates. Abortions in buffalo seem infrequent in comparison with cattle (Reichel et al. 2015).

In Iran, buffalo breeding is traditional and limited in Southwest (Ahvaz) and Northwest (Urmia) regions. In the traditional condition, the chance of infection increase due to close contact with final hosts. Little is known about N. caninum infection in buffaloes in Iran and other countries. Reported seroprevalences are between 1.5% in Vietnam and 100% in Australia (Dubey et al. 2007), but the most differences are probably related to the different laboratory diagnostic tools. The overall seroprevalence is estimated 48% worldwide (Reichel et al. 2015). In a study (Hajikolaei et al. 2007), the seroprevalence of N. caninum was reported 37% in Ahvaz. There were significant differences between male (15%) and female (43.3%) animals. In females, 7.7% of heifers and 56.8% of cows were seropositive (P < 0.05). In another survey from Ahvaz, seroprevalence rate was 55.9%, 66.5%, and 56.9% using different ELISA and also NAT techniques. No significant statistical differences were detected between seroprevalence rate, age, and sex of animals (P > 0.05) (Pourmahdi-Borujeni et al. 2015). Rezvan et al. (2019) found that in Urmia district, where all of seropositive samples (19.2%) were confirmed using nested PCR for detection of Neospora NC5 gene, a 100% concurrency was detected among the high levels of anti-Neospora antibodies and parasitemia. The results indicated that the buffalo has no significant role in the cycle of disease in Iran because of low population in limit scale.

Neospora caninum in dogs

Dogs play a significant role for maintaining the infection and also occurrence of 'storms of abortions' due to horizontal transmission in dairy herds (Dubey et al. 2005, 2007). Adult infected dogs may represent various neuromuscular neurologic disorders. Pups are usually born without clinical signs. They typically show progression toward ascending paralysis of the rear limbs after 21 days of birth. In addition, paralysis of jaw, atrophy, dysphagia, and heart failure are gradually observed (Barber and Trees 1996). The presence of infected dog in dairy farms is a risk factor for herd population (Gharekhani and Yakhchali 2019; Ribeiro et al. 2019). Also, dogs commuting between farms and rural areas have an important impact in spreading of the infection. These factors are the possibility to consumption of cattle fetuses or placentas infected with tachyzoites or cysts of N. caninum, materials of aborted fetuses, or uterine discharge by dogs living in farm suburb (Haddadzadeh et al. 2007; Malmasi et al. 2007).

Infection to *N. caninum* was reported in range zero to 54.6% in different type of animals, samples, and diagnostic methods from Iran (Razmi 2009; Khordadmehr et al. 2012) (Table 2). This rate was varied from 1 to 100% throughout the world (Dubey and Schares 2011). Results of metaanalysis, the pooled seroprevalence was estimated 17.1% in global scale (Anvari et al. 2020) and also the infection rate was calculated to be higher in males (15.8%), as compared to females (15.2%).

The high risk of infection was seen in farm dogs with a close contact with seropositive animals and secreted materials. In the most of reports, a high infection was detected in stray dogs above 2 years old. This is due to their more probable of exposure to *N. caninum* infection and infected postnatally (Hossininejad and Hosseini 2011). Some of researchers stated that the highest infection is in male dogs (Malmasi et al. 2007; Yakhchali et al. 2010; Sharifdini et al. 2011). Most of the farmers tend to have male dogs in their farms. Therefore, the male dogs might have been more infectious than females (Malmasi et al. 2007; Khanmohammadi and Fallah 2011).

Study by Gharekhani et al. (2013a) found that 52.8% of stray dogs and 18% of shepherd dogs were seropositive (P < 0.001). Haddadzadeh et al. (2007) reported 28% and 11.3% of seropositivity in farm and urban dogs, respectively (P = 0.032). Additionally, antibodies to *N. caninum* in farm dogs (46%) were higher than household dogs (20%) (P = 0.005) (Malmasi et al. 2007). It was found that only one of the evaluated cases with antibodies titers of more than 1:800 in IFA technique had severe clinical signs

of dermatitis and locomotion difficulties (Malmasi et al. 2007). Hosseininejad et al. (2010a) reported no statistical evaluation was possible to compare the infection rate and different breeds. Because of limited information on breed impact on *N. caninum* infection in dogs, focus on this subject in future investigations is essential.

Animals with latent disease and/or co-infection with other pathogens are high susceptible to N. caninum infection (Anvari et al. 2020). The simultaneous presence of N. caninum and T. gondii infections was detected in 2.8%, 4%, and 8.9% of dogs from Hamedan, Ahvaz, and other regions of Iran, respectively (Hosseininejad and Hosseini 2011; Hosseininejad et al. 2018; Gharekhani and Yakhchali 2019). Also, N. caninum and Leishmania infantum co-infection was 9.4% in Northwest of Iran (Sharifdini et al. 2011). Dogs are a matter of concern also to public health as a source of N. caninum infection. Accordingly, the preparation of a comprehensive information on N. caninum infection in dogs is necessary. It is suggested that researchers carry out further projects for blocking the neosporosis in dogs especially working on efficient vaccine.

Neospora caninum in sheep and goats

In Iran, most of the sheep and goat breeding farms are traditional, and animals have a direct contact especially with stray dogs. The clinical and economic reality of neosporosis in sheep and goats is unclear. The disease has been sporadically reported as a reason for reproductive failures such as abortion and neonatal mortality in these animals, similar to toxoplasmosis (Dubey and Schares 2011; Razmi and Naseri 2017). Two researches on neosporosis in goats were conducted in Iran by Gharekhani et al. (2016, Gharekhani et al. 2018a, b). In Hamedan, 6.2% and 13.3% of examined animals were seropositive in individual and herd level, respectively. The infection was 5.8% in flocks with 30-99 population and 23.7% in flocks with 100–300 population (P = 0.013, OR = 5). The highest infection was significantly found in animals above 2 years of age (54.4%, P = 0.024, OR = 2.6). Also, 11.1% of male and 5.6% of females were seropositive (P = 0.112). This rate in animals with history of abortion (60%) was 53-fold higher than those without abortion (P < 0.0001) (Gharekhani et al. 2016). In Khuzestan province, 10.8% of goats and 32.4% of sheep flocks were positive for N. caninum and also the rate of mixed infection with N. caninum and T. gondii was 5.4% (Gharekhani et al. 2018a, b). No significant correlation observed between sheep and goats (P = 0.082). Seropositivity in animals with history of abortion (25%) was 3.6-time higher than others (8.3%, P = 0.022). There were significantly high infection in < 1year-old animals (25.9%) versus 1-2 (15.8%) and > 2 (5.8%) age groups (P = 0.005). According to herd size, higher infection was in animals that sampled in herds with more than 500 populations (P = 0.016). No connection with gender was found (Table 3, P = 0.330).

The lowest and highest seroprevalence rate of N. caninum in sheep was reported 1.5% in Aleshtar (Lorestan province) and 6.8% in Golpayegan (Isfahan province), respectively (Table 3) (Ezatpour et al. 2013; Shahi 2016). Gharekhani et al. (2013b) reported 2.2% of seropositivity in aborted ewes in Hamedan. In addition, the significant difference was neither observed in different age groups (P = 0.999) nor between different fetal age groups (P = 0.671). In Ezatpour et al. (2013) work, no significant correlation was seen between seropositivity in aborted and healthy sheep from Aleshtar district (P = 0.516). Gharekhani et al. (2018a, b) reported the seropositivity in 6.8% of sheep and 37.8% of flocks in Khuzestan province. N. caninum and T. gondii co-infection was 3.2% in individual and 21.6% in herd levels. Population size in sheep flocks had a significant influence on seropositivity (12.2% in <250, 1.7% in 250–500 and 8.9% in > 500 population; P < 0.0001) unlike to history of abortion (P = 0.094), age (P = 0.597), and sexes (P = 0.807) (Gharekhani et al. 2018a, b).

Histopathological examination in animals with neosporosis demonstrated extensive suppurative and nonsuppurative meningoencephalitis (Sasani et al. 2013). Along with various lesions incidence of cellular and vascular, the glial reactions were also assessed in aborted fetal brain tissues containing focal or diffuse gliosis by the authors. Some researchers (Asadpour et al. 2013; Sasani et al. 2013; Razmi and Naseri 2017) believe that neosporosis may act as a causative agent of abortion in sheep, as N. caninum DNA was isolated in 9.9% and 8.5% of brain samples derivate from aborted fetuses' in Mashhad and Tabriz regions, respectively. In investigations by Arbabi et al. (2016), DNA of N. caninum was detected in 6.7% and 0.7% of heart and brain samples, respectively, in animals without clinical signs.

Neospora in equine

Another species of *Neospora*, *N. hughesi* is considered as equine parasite. Abortion, protozoal myeloencephalitis, and neuromuscular disorder are the main findings of neosporosis in horses (Dubey and Schares 2011). Seroprevalence of *Neospora* infection was reported 20–40.8% and 52% in horses and donkeys, respectively (Gharedaghi 2011; Hosseini et al. 2011; Moraveji et al. 2011; Gharekhani et al. 2013c; Tavalla et al. 2015). In a study by Gharekhani et al. (2013c), 42.2% of riding club horses and 40% of rural horses from Hamedan were seropositive (P = 0.811); also, no significant connection was recognized among seropositivity, age and genders equivalent to other reports (Moraveji et al. 2011; Tavalla et al. 2015). The detail of published articles is demonstrated in Table 4.

Neospora caninum in camels

Camel husbandry constitutes an important economic activity in the desert populations of the East, South, and Central regions of Iran (Fig. 1). According to the Iranian Veterinary Organization, average populations of 150,000 camels are reared in Iran. The camel is used as a load animal, playing a crucial role in transportation from rural areas. It is also a source of meat, skin, and leathers (Sadrebazzaz et al. 2006; Hosseininejad et al. 2009). The seroprevalence rate was detected as 3.2% and 3.9% in Isfahan and Yazd central regions of Iran (Hosseininejad et al. 2009; Hamidinejat et al. 2013), 5.8% in Mashhad (Northwestern Iran) (Sadrebazzaz et al. 2006), and 27% in Bushehr (South of Iran) (Namavari et al. 2017) (Table 4). Hamidinejat et al. (2013) reported that 3.85% of male animals and 4.35% of females were seropositive; in addition, this rate was 3.66% and 4.44% in animals living that hot-dry and cold-dry conditions. Data analyses revealed no statistical significant differences were seen between the seroprevalence of N. caninum in sex groups and different climates (P > 0.05). Many of the camel population is kept in communities lacking equipment and trained personnel for carrying out parasitological examinations; therefore, the development of rapid techniques for rapid detection of the main camel parasites such as N. caninum is essential.

Neospora caninum in cats

Cats are susceptible to N. caninum and can be infected both transplacentally and postnatally. The clinical aspects of Neospora infection are unclear in naturally infected animals (Dubey et al. 2007). Experimental infection has been reported in immunocompetent cats (Hamidinejat et al. 2011a). The presence of cats in the farms is a risk factor; as they have important role in the remaining of infection with eating the infected rodents (Dubey and Schares 2011). In Hamidinejat et al. (2011a, b) study from Ahvaz, 19% of feral cats and 14% of domestic Persian cats were seropositive to N. caninum. There was no significant correlation between sex and seroprevalence rate (Table 4). Also, logistic regression showed that the prevalence rate increased significantly with age (P < 0.05). Available information confirmed that cats are the suitable intermediate hosts for N. caninum and may have a significant impact in epidemiology of the parasite (Dubey and Schares 2011). In the farm, cats become infected via consumption of cysts in rodents and also oocysts in environment (Donahoe et al. 2015).

Neospora caninum in rodents

Rodents are one of the intermediate hosts for N. caninum and plays a significant role in continuity of parasite cycle (Dubey et al. 2007). The presence of rodents in dairy farms had a positive correlation with seroprevalence rate in cows and also is a risk factor for neosporosis in herd (Gharekhani and Yakhchali 2019). Yet, there are only four published papers on Neospora infection in Iranian rodents, only (Table 4). Mosallanejad et al. (2018b), working from Ahvaz, found antibodies to N. caninum in 6% of black rats (Rattus rattus). The seroprevalence rates were not significant statistically (P < 0.05) according to sexes. Brain tissues of 70 different rodent species (Persian jird (Meriones persicus): 50, gray dwarf hamster (Cricetulus migratorius): 9, and house mouse (Mus musculus): 11) were evaluated for N. caninum infection using molecular methods in Meshkin-Shahr district, Northwest Iran. All of the examined rodents were negative (Nazari et al. 2019) similar to Khani (2016), who investigated on N. caninum infection in rodents that were captured from dairy farms in Arak, Central Iran. In another research from Meshkin-Shahr by Nazari et al. (2020), antibodies to N. caninum were detected 12.9% in Persian jird, 20% in gray dwarf hamster, and 39% in house mouse using the indirect fluorescence antibody test (IFAT); a significant correlation (P = 0.002) was seen among the seroprevalence rate and different species of rodents. Interestingly, although low infection rates have been reported in rodents from Iran, there are some reports on notable infection in other countries such as Taiwan (16.4%) and USA (10%) (Huang et al. 2004; Hughes et al. 2006). In addition, N. caninum infection rate was recognized in 77% in house mouse, 71% in rock squirrel (Spermophilus variegatus), and 50% in brown rat (Rattus norvegicus) from Mexico (Medina-Esparza et al. 2013). We recommended regular control of rodents to eliminate a potential reservoir of N. caninum infection in the farms.

Neospora caninum in birds

Neospora caninum antibodies and DNA were detected in different species of birds, but viable parasites have neither been demonstrated nor isolated by bioassays. Additionally, antibody responses to *N. caninum* were not persistent in experimentally infected birds (Dubey et al. 2007; Barros et al. 2018). None of the birds inoculated with tachyzoites developed clinical signs. It seems that chickens can be successfully infected with tachyzoites, but the host eliminates the infection. The role of birds in the life cycle of *N. caninum* is uncertain, but they may contribute to parasite transmission in sylvatic cycles either as mechanical vectors or as intermediate hosts (Barros et al. 2018).

There is little information on presence of N. caninum in birds in Iran and also in other countries (Abdoli et al. 2015; Barros et al. 2018). In Iran, N. caninum infection has been reported 2.8% and 3.7% in sparrows (Passer domesticus) (Abdoli et al. 2015; Bahrami et al. 2015), 17.3% in chickens (Gallus domesticus) (Savari et al. 2016), 9.8% and 30.4% in Pigeons (Columba livia) (Bahrami et al. 2016), and 9.9% in hooded crows (Corvus cornix) (Table 5). The high infection in pigeons is probably related to the pigeon's diet. Pigeons commonly feed directly from the ground and are probably exposed to N. caninum after ingestion of the parasite oocysts from the soil (Bahrami et al. 2016). Bartels et al. (1999) reported a significant association between the presence and number of poultry with the increase of seroprevalence and abortion storms related to N. caninum in the farms. The occurrence of N. caninum infection in ground-foraging birds can be used to indicate environmental contamination with N. caninum oocysts (Abdoli et al. 2015). However, more research is needed to reveal the role of birds in the epidemiology of neosporosis.

Neospora caninum in humans

There is some information on neosporosis in humans worldwide. Various levels of N. caninum antibodies in immunodeficient humans are documented (Oshiro et al. 2015). In Lobato et al. (2006) study, antibodies to N. caninum were recognized 38% and 18% in HIV-infected patients and humans with neuro-psychological disorders, respectively, while this rate was 5% in newborns and 6% in healthy and normal man. In other hand, no antibodies were found in women with history of abortion (Petersen et al. 1999). Duarte et al. (2020) reported 24.3% of seropositivity for IgG anti-N. caninum in humans using IFAT; all of samples were negative for IgM. Also, 1% of umbilical cord blood samples were positive in molecular evaluation using Nc5 gene. No seen parasite-tissue cysts and/or inflammatory infiltration in histopathology. There was a significant connection (P < 0.05) among seroprevalence rate and the presence of dogs.

Neospora caninum has been successfully cultured in human cell lines, but that zoonotic aspect has not been defined yet (McCann et al. 2008). However, till date, no reports are available on this topic from Iran.

Conclusion

This research compiled the comprehensive information on *Neospora* infection in animals and birds in Iran for the first time. It is clear that *N. caninum* infection is distributed in wide scale from Iranian animals especially in cattle

population. Both horizontal and vertical ways have an active role in transmission of the infection in this region. There are significant economic losses in Iranian dairy hers due to abortion, culling, and reproductive failures. This review can provide baseline information for future research. Study on other hosts, especially on wild and exotic animals, is recommended for exact estimate of neosporosis in Iran. Focus on molecular and genotyping diagnosis of *N. caninum* strains is needed; which will be helpful for developing vaccine and establish the link between wild and domestic cycles of disease. Public education on risk factors associated with *N. caninum* infection for farmers and rural people is suggested.

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Compliance with ethical standards

Conflict of interest The authors declare that they have no conflict of interest.

Ethical approval The ethical guidelines have been consistence during the designing the project, date collection, and preparation of results. All of authors gave their informed consent prior to their inclusion in the research. We also hereby declare all ethical standards respected in the submitted article.

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