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A review of bovine fasciolosis and other trematode infections in Nigeria

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Short title: Bovine trematodiasis in Nigeria
Abstract

Trematode infections cause serious economic losses to livestock worldwide. Global production losses due to fasciolosis alone exceed US$3 billion annually. Many trematode infections are also zoonotic and thus a public health concern. The World Health Organisation has estimated that about 56 million people worldwide are infected by at least one zoonotic trematode species and up to 750 million people at risk of infection. Fasciolosis caused by the fluke, *Fasciola gigantica* is endemic in Nigeria and is one of the most common causes of liver condemnation in abattoirs. Cattle total losses from *Fasciola* infection in Nigeria have been estimated to cost £32.5 million. Other trematode infections of cattle including paramphistomosis, dicrocoeliasis and schistosomiasis have all been reported in various parts of Nigeria with varying prevalence.

Most publications on trematode infections are limited to Nigerian local and national journals with very few international. This paper therefore summarised the current data on distribution, control and zoonotic trematode infections in Nigeria and other African countries. We also identified research gaps and made recommendations for future research and areas for funding for policy/planning.
Introduction

Nigeria is one of the four leading livestock producers in Sub - Saharan Africa (FAO, 2009) contributing up to 12.7% of the total Nigerian agricultural gross domestic products (CBN, 1999). In the tropics, cattle are generally reared under the transhumance husbandry systems with little supplementary feeding resulting in low productivity and high pre-weaning mortality (Ulayi et al., 2007). Also, acute shortage of feeds during the dry season remains a common occurrence, compelling these animals to graze around water bodies that are often heavily infested with potential intermediate hosts of trematode infections (Ulayi et al., 2007).

Trematode infections of cattle including fasciolosis (Babalola & Schillhorn van Veen, 1976, Dipeolu et al., 2000), paramphistomosis (Bogatko, 1975), dicrocoeliasis (Schillhorn van Veen et al., 1980, Ulayi et al., 2007) and schistosomiasis (Pugh et al., 1980, Ndifon et al., 1988) have all been reported in various parts of Nigeria. They are referred to as 'digenetic' because they require at least two different kinds of hosts for their full development. The first are the intermediate hosts of mollusc species (slug, snail or shell fish) while the final hosts are vertebrates (Taylor, 1964). Mixed infection with more than one trematode species is a common occurrence in animals due to similarities in life cycles. Fasciola infection has been reported to occur concurrently with Schistosoma andparamphistomid species(Schillhorn van Veen et al., 1980, Yabe et al., 2008). It is also reported to occur concurrently with Dicrocoelium species (Schillhorn van Veen et al., 1980, Ulayi et al., 2007). Infection with trematode parasites can lead to severe losses to farmers hence affecting sustainability of food production (Alvarez Rojas et al., 2014).

Fasciolosis, which is also referred to as distomatosis or liver fluke disease, is a parasitic disease caused by the trematode of the genus Fasciola. The most important species are F. hepatica and F. gigantica, found in the temperate and tropical regions of the world respectively. The geographical distribution of Fasciola species is determined by the distribution of the snail intermediate hosts (Boray, 1985). The temperate disease occurs in Europe, America and Oceania while the tropical fasciolosis is common in Africa, Indian sub-continent, Central
and Southeast Asia and other subtropical and tropical areas of the world (FAO, 1993). Both species overlap in many areas of Africa and Asia (Walker et al., 2008). Adults of each species can occur concurrently in the same animal host, either because of local overlap or because of livestock movement (Mas-Coma et al., 2009). The two Fasciola species may hence interbreed resulting in hybrids species (Agatsuma et al., 2000, Ashrafi et al., 2006). They are hermaphrodites and are found in the bile ducts of a large number of herbivorous ruminants, equine, pigs, rabbits (FAO, 1993) and wild animals (Hammond, 1972). Man is also a suitable host (Boray, 1969). The intermediate hosts of F. gigantica are true water snails belonging to the phylum Mollusca, class Gastropoda and subclass Pulmonata (Wright, 1971). They belong to the family Lymnaeidae and super species Lymnaea (Radix) auricularia sensu lato.

Paramphistomes Fischoeder 1901 also known as rumen flukes are gastro-intestinal trematodes belonging to the family of Paramphistomatidae (Soulsby, 1982). These are conically shaped flukes measuring 5-12 mm × 2-4 mm. The adults predilection sites are the rumen and reticulum of ruminants while the immature parasites are found in the small intestines and abomasums (Rojo-Vázquez et al., 2012). Snails of the families Planorbidae, Bulinidae and Lymnaeidae act as intermediate hosts (Soulsby, 1982, Castro-Trejo et al., 1990). They are largely non-pathogenic but clinical outbreaks have been reported to occur. The most important species in Africa is Paramphistomum microbothrium (Dinnik, 1964). Others are: P. cervi (European Environment), P. ichikawar (Australasia) (FAO, 1993) and P. daubneyi first described in Kenya and common in Europe (Abrous et al., 1996). Recently there are reports of increasing number of cases of rumen flukes identified mainly as Calicophoron daubneyi in cattle and sheep in the republic of Ireland (Zintl et al., 2014, Toolan et al., 2015). The adult paramphistomes are regarded as commensals in the rumen as heavy infections are tolerated without causing any damage to the rumen (Dinnik, 1964) although immature parasites in the small intestine cause clinical disease (Aiello, 1998).

Dicrocoelium species, commonly referred to as the lancet fluke, are found in the bile duct of domestic and wild ruminants (Otranto & Traversa, 2002, Taylor et al., 2007). Other animals such as rabbits, pigs, dogs, horses and human can also be
infected (Rojo-Vázquez et al., 2012). There are several species; with *D. dendriticum* (Otranto et al., 2007) having the widest distribution worldwide while *D. hospes* is common in Africa (Aiello, 1998). Other species are *D. chinensis* and *D. suppereri* (Rojo-Vázquez et al., 2012). Two intermediate hosts are required. The first intermediate host is a terrestrial snail of the genus *Limicolaria* while the second intermediate hosts which are brown ants of the genus *Formica* (Soulsby, 1982).

*Schistosoma* species are elongate, unisexual trematodes commonly referred to as blood flukes and widely distributed throughout Africa, Middle East, Asia and some Mediterranean countries (Soulsby, 1982). They are found in the blood vessels such as portal, mesenteric and intestinal veins of domestic animals (Aiello, 1998). The species, *Schistosoma matteei* and *S. bovis* are common in Africa, Middle East and southern Europe (Vercruysse & Gabriel, 2005, Taylor et al., 2007). *S. bovis* are the most pathogenic in animals in Africa (Vercruysse & Gabriel, 2005). They occur in the portal and mesenteric vessels of cattle, sheep and goats. They are similar to the human parasite *S. haematobium* (Soulsby, 1982). *S. matteei* has also been reported in sporadic human infection. Other African species in ruminants includes *S. curassoni* (common in Western Africa), *S. margreboweiei* and *S. leiperi*. In Asia, *S. spindale*, *S. nasale* (nasal worm), *S. indicum*, *S. incognitum* and *S. japonicum* (which are zoonotic in Far East) (De Bont & Vercruysse, 1997). Aquatic snails such as *Bulinus, Physopsis, Oncomelania, Lymnaea* and *Indoplanorbis* act as the intermediate hosts (Taylor, 2007).

This review aims at describing the current status of bovine fasciolosis and other trematode infections in Nigeria and surrounding African countries, in relation to economic losses, distribution, molecular studies, control strategies and human infections. Data were collected from publications obtained from online search as well as local national journals. Bovine fasciolosis in reported in literatures across different states of Nigeria were mapped by using QGIS® software v2.12 (Lyon, France).
Economic losses from trematode infections

Tropical fasciolosis caused by *Fasciola gigantica* is regarded as one of the most important single helminth infection of ruminants in Asia and Africa (Boray, 1985, Fabiyi, 1987). Economic losses from fasciolosis are often difficult to estimate and may result directly from increased liver condemnation or indirectly from decreased livestock productivity (Taylor, 1964). Although direct losses are easier to measure, indirect losses are considered to be far more economically important (Kaplan, 2001). With the total cattle population in Africa estimated at 201 million animals, an annual loss of about US$840 million from fasciolosis infection is predicted (Spithill *et al.*, 1999), but this cost is likely to have increased significantly in the last eighteen years. Production losses in liver fluke infected cattle often take the form of reduced milk production in dairy cattle and poor feed conversion in beef cattle (Armstrong, 1982). A linear relationship between burden of adult *F. gigantica* and weight gain of cattle has been described, with infected animals gaining only about half the annual weight compared to control (Sewell, 1966). The amount of weight loss may also, however, be dependent on age, level of nutrition and intensity of infection (Spithill *et al.*, 1999). Other clinical manifestations are anaemia, reduced fertility and reduced work capacity (Hillyer, 2005), and where there is massive infection with immature parasites, sudden death could occur and lead to serious economic losses (Torgerson & Claxton, 1999, Rojo-Vázquez *et al.*, 2012).

Severe infection with immature paramphistomes can cause great economic losses due from reduced weight gain and decrease in milk production (Horak, 1971). Paramphistomosis has been reported to cause between 30-40% mortality in cattle and sheep. These deaths may result from anaemia, hypo-proteinaemia, profuse diarrhoea and marked emaciation (Soulsby, 1982), and a decrease in milk yield in dairy cattle (Spence *et al.*, 1996).

Economic losses from dicrocoeliasis is less apparent compared to those from other trematode infections and is difficult to quantify due to concurrent infection with other gastrointestinal parasites. This may also be largely due to the asymptomatic nature of the disease (Otranto & Traversa, 2003). Clinical signs are mostly absent
but losses occur mainly from liver condemnation due to fibrosis of bile ducts and cirrhosis (Taylor, 2007).

An abattoir survey to determine the prevalence and species of cattle schistosomiasis in Zambia revealed an overall prevalence of 51%, with 93% of the infected animals having up to 100 worm pairs in the mesenteric veins; *Schistosoma mattheei* was the predominant species (75%); while *S. leiperi* (12%) and *S. margrebowiei* (2%) were also reported (De Bont et al., 1994).

Schistosomiasis due to *Schistosoma bovis* has also been reported with a prevalence rate of 4.8% in Tanzania (Nzalawahe et al., 2015). These parasites cause economic losses due to their long-term effect on growth and productivity and also due to increased susceptibility to other parasitic and bacterial diseases (De Bont & Vercruysse, 1998). The clinical signs associated with the intestinal and hepatic form of disease in ruminants are haemorrhagic enteritis, haematuria, anaemia and loss of weight with death within few months of the disease. Nasal schistosomiasis, on the other hand, is a chronic disease that causes coryza and dyspnea in infected animals (Aiello, 1998). Production losses reported due to schistosomiasis in cattle are attributed mainly to losses occurring in animals aged between 6-30 months and are due to reduced weight gain, liver condemnation, poor future reproductive performance and death (Hunt et al., 1984). Also approximately 165 million cattle are likely to be infected with *Schistosoma* species worldwide (De Bont et al., 1994).

Many trematode infections are also zoonotic and thus a public health concern (Chen & Mott, 1990, Mas-Coma et al., 2005, Wolfe, 1966). The World Health Organisation has estimated that about 56 million people worldwide are infected by at least one zoonotic trematode species (Furst et al., 2012) and up to 750 million people at risk of infection (Keiser & Utzinger, 2009). It has been suggested that recent environmental changes associated with global warming which favours the development of snail intermediate hosts contribute to an increase in disease outbreaks (Mitchell, 2002).
Epidemiology of bovine fasciolosis in Nigeria

Tropical fasciolosis due to *F. gigantica* Cobbold 1855 has been reported in several parts of Africa (Schillhorn van Veen, 1980, Phiri *et al.*, 2005a, Abebe *et al.*, 2010, Nzalawahe *et al.*, 2014) and is likely to be a problem throughout the continent. The two species of *Fasciola: F. gigantica* and *F. hepatica* and their respective snail intermediate hosts have both been reported in Africa. Fasciolosis is endemic in Nigeria and is of great economic importance (Ogunrinade & Ogunrinade, 1980). The disease has been reported with varying prevalence across the country and is one of the most common causes of liver condemnation in abattoirs. The prevalence distribution of bovine fasciolosis is mapped (fig 1) according to available data published between 1980 and 2016 (Schillhorn van Veen, 1980, Nwosu & Srivastava, 1993, Nkwu *et al.*, 2004, Opara, 2005, Ekwunife, 2006, Adedokun *et al.*, 2008, Umar *et al.*, 2009, Ibironke & Fasina, 2010, Sugun *et al.*, 2010, Omoleye, 2012, Gboeloh, 2012, Odigie & Odigie, 2013, Abraham, 2014, Ardo, 2014, Magaji, 2014, Ngele & Ibe, 2014, Yahaya & Tyav, 2014, Onyeabor, 2014, Ejeh *et al.*, 2015, Elelu *et al.*, 2016). The high prevalence rate from liver condemnation in a 3-year period reported in Lagos State (Ibironke & Fasina, 2010) can be explained by the fact that Lagos State is the commercial nerve centre of Nigeria with a very large human population, large number of cattle from all over the country are often slaughtered from the abattoir.

Cattle total losses from *Fasciola* infection alone in Nigeria have been estimated to cost £32.5 million (Fabiyi & Adeleye, 1982). A survey of abattoirs in Nigeria showed that about 70% of organ condemnation, mainly of livers was due to fasciolosis (Alonge & Fasanmi, 1979). A recent study reported up to 88.1% liver condemnation due to liver flukes alone in Lagos abattoir (Ibironke & Fasina, 2010). These studies are in agreement with a 3-year abattoir study in Southwestern Nigeria in which losses from liver condemnation were estimated at US$134,000 (Ibironke & Fasina, 2010). These data translate to huge economic losses from cumulative condemnation across the country.

Acute fasciolosis due to the migration of juvenile flukes is rare in cattle; however, a few cases have been reported in Nigeria. For example, on a dairy farm in North-central Nigeria, a pregnant cow was reported to have died during parturition,
where post mortem examination revealed inflammatory lesion and fibrosis due to fasciolosis (Okaiyeto et al., 2012). The chronic form of the disease is the most common in cattle in Nigeria (Ogunrinade & Adegoke, 1982) and it occurs when small numbers of flukes finally enter the bile duct and infection becomes patent. This results in a chronic wasting disease from slow acquisition of liver flukes for months or even years. The normal tissues of the liver are replaced by fibrous tissues, with hyperplastic cholangitis and macroscopically visible calcification of the bile ducts giving rise to so called 'pipe-stem liver' (Taylor et al, 2007; Soulsby, 1982). Clinically, the condition is characterised by weight loss, anaemia, and oedema at the jaw (bottle jaw), thoracic and lower abdominal regions (Radostits et al., 2000, Mitchell, 2002, Taylor et al., 2007, Rojo-Vázquez et al., 2012).

**Factors influencing outbreak of bovine fasciolosis**

*Season of the year*

There are varying reports on the seasonal infection rates of *F. gigantica* across Nigeria. Majority of studies reported high infection rates in the beginning of the dry season (Schillhorn van Veen et al., 1980, Umar et al., 2009) which is in agreement with studies in other parts of Africa like Zambia, where bovine fasciolosis was also reported with higher fluke abundance reported in the post-rainy season (Phiri et al., 2005b). Similar observations were made in Zimbabwe, where *Fasciola* faecal egg counts were reported to follow a seasonal variation with increase during end of dry season with highest liver condemnation reported during the rainy season (reviewed by Pfukenyi et al., 2005). However, another study carried out in the Southern part of Nigeria reported an all year occurrence of infection (Gboeloh, 2012). This all year occurrence in the South was attributed to the favourable climatic condition (warm and humid) which favours development of parasites, and the increase in the density of snail intermediate hosts (Gboeloh, 2012). The climatic condition in Southern Nigeria could be likened to those of highland areas of Iringa, Tanzania (Nzalawahe et al., 2015) and Zimbabwe (Pfukenyi et al., 2006a) where wet/swampy grazing areas reportedly favour availability and distribution of snail intermediate host. In addition a recent study in Tanzania have associated trematode infections including *Fasciola* with irrigation practices during the dry season that favours growth of intermediate snails hosts and development of trematode larval stages (Nzalawahe et al., 2014).
**Type of management system**

The type of management system has been shown to significantly influence the prevalence of *Fasciola* infection (Keyyu et al., 2005). A low prevalence rate was also reported to occur in cattle reared by sedentary husbandry system in Lake Chad area (Jean-Richard et al., 2014). While a high prevalence rate of 54.3% has been reported in cattle managed extensively in South western Nigeria (Adediran et al., 2014), this is not surprising because, cattle come into contact with snail-infected habitat during extensive communal grazing. In contrast, under intensive management systems, metacercaria-free water and herbage can be supplied to the cattle thereby minimizing the likelihood of outbreaks of fasciolosis.

**Age of cattle**

In a recent study from Nigeria, *F. gigantica* infection was reported to be higher in adult cattle than weaners (Elelu et al., 2016). This is in agreement with past studies in Africa with similar findings (Pfukenyi et al., 2006a, Nzalawahe et al., 2014). The difference in prevalence within age group was attributed to longer length of exposure to the infection in adults compared to weaners (Pfukenyi et al., 2005, Nzalawahe et al., 2014). It was also concluded that adult cattle act as a constant source of *F. gigantica* infection for the more susceptible young animals (Pfukenyi et al., 2005).

**Distribution of snail intermediate host**

The intermediate snail host *Lymnaea (Radix) natalensis* that is widespread throughout Africa is also widely distributed in Nigeria, although its occurrence is restricted to permanent water bodies (Ndifon & Ukoli, 1989). *Lymnaea* is fairly common in regions with rainfall over 1000mm (Schillhorn van Veen, 1980) and it tolerates relatively high temperatures (Njoku-Tony, 2011) consistent with tropical climates. In Uganda, *L. (Radix) natalensis*, the intermediate snail host of *F. gigantica*, is the vector commonly found abundant at lower altitudes below 1800m, while *Galba truncatula* (intermediate snail host of *F. hepatica*) was found only at altitudes above 3000m (Howell et al., 2012). The presence of the two
vectors indicates the possibility of the two *Fasciola* parasites in Uganda. Other countries of Africa have however reported the presence of the two species of *Fasciola* (Yilma & Mesfin, 2000, Walker et al, 2008; Sisay & Nibret, 2013). To date the authors of this review are not aware of any report of *Galba truncatula* in Nigeria, although specific studies to identify these species in areas of high altitude might be worthwhile.

In Nigeria, during the wet season, many of the developing *Lymnaea* snails are washed away in torrential streams after heavy rain and this may be important in the spread of *Fasciola* (Schillhorn van Veen, 1980). Snails were also reported to be more abundant during the beginning of dry season, reach a climax in the middle of the dry season, but decrease towards the end of the dry season when most streams and pools dry up (Schillhorn van Veen, 1980). This is supported by a field study in south-western Nigeria which shows that dry season conditions favour snails and this was said to be due to low turbidity, reduced currents and substantial growths of algae and macrophytes (Ndifon & Ukoli, 1989). In the sub-tropical and tropical countries with distinct wet and dry seasons, optimal development of fluke eggs to miracidia occurs at the start of the wet season and development within the snail is complete by the end of rains (Taylor, 2007). The dry season therefore coincides with the snail shedding of cercaria and more animals grazing closer to streams and ponds thereby predisposing them to infection. Herdsmen migrate in search of water and grazing during the dry season and thousands of cattle often converge on the few ponds, which fail to dry up (Ikeme & Obioha, 1973).

**Diagnosis of bovine fasciolosis**

The most common method of diagnosis is by faecal egg counts and pathological lesions in the liver during abattoir examination. Serological diagnostic method to detect antibodies, such as indirect ELISA (Damwesh & Ardo, 2013, Aliyu *et al.*, 2014) and direct ELISA (Fagbemi *et al.*, 1997) has been carried out in Nigeria. Another method of diagnosis is testing for precipitating antibodies using the Agar Gel Precipitation Test (AGPT) (Adedokun *et al.*, 2008). Comparatively, this method has been shown to detect more positive cases than faecal and bile egg counts (Adedokun *et al.*, 2008). In Kwara State, North-central Nigeria, survey of
cattle for fasciolosis revealed a higher prevalence rates from faecal analysis compared to a previous abattoir study that utilized liver examination. This further showed the lack of sensitivity of the abattoir method of diagnosis as positive samples are likely to be lost (Elelu et al., 2016).

**Epidemiology bovine paramphistomosis in Nigeria**

Rumen flukes are paramphistome parasites commonly seen in the abattoir in Nigeria (Bunza et al., 2008). The snail vector of the disease (*Lymnaea, Planorbis* and *Bulinus*) has also been reported in Nigeria (Ndifon & Ukoli, 1989, Brown & Kristensen, 1993). The species reported to occur in Nigerian domestic livestock include *Paramphistomum microbothrium, Carmyerius gregarious, Carmyerius spatiosus Cotylophorum cotylophorum* (Schillhorn van Veen et al., 1975) and *Paramphistomum cervi* (Bogatko, 1975). Others species recovered from Nigerian cattle are *Ceylonocotyle dicranocoelium, Bothriophoron bothriophoron, Calicophoron calicophorum* and *Calicophoron microbothrioides* (Dube et al., 2013).

Prevalence rates of 2.2% (Edosomwan & Shoyemi, 2012); 18.8% (Nwigwe et al., 2013); 16.1% (Elelu et al., 2016) has been reported for paramphistomes in past studies in Nigeria. A higher prevalence rate of 41.67% for *P. cervi* was reported in another abattoir study in Northern Nigeria (Nnabuife et al., 2013). Similarly, also in Northern Nigeria a high paramphistome prevalence rate of up to 56.0% in slaughtered cattle has been reported in a study area around a marshy river valley. The grazing land associated with this river valley provides suitable breeding sites for snail intermediate hosts of the parasites (Bunza et al., 2013). The convergence of cattle in common graze land during dry season as well as irrigation practices has been implicated in high prevalence of paramphistomosis (Nzalawhe et al., 2014). In a cross-sectional study in Ethiopia, 51.8% of cattle slaughtered were positive for paramphistomosis with peak prevalence rates observed during October to November (Ayalew et al., 2016). Although adult cattle were more likely to be infected with paramphistomes because they are more likely to be infected when taken out for grazing, however several studies have shown that there is no statistically significant difference in paramphistome infection between age groups of cattle (Titi et al., 2010, Tagesse et al., 2014, Khedri et al., 2015, Ayalew et al., 2016).
Epidemiology of bovine dicrocoeliasis in Nigeria

There are records of *Dicrocoelium hospes* infection in cattle in Nigeria (Nwosu & Srivastava, 1993, Ulayi et al., 2007), and snail species of *Limicolaria flammea* have been experimentally shown to be a suitable intermediate host of the disease (Fashuyi & Adeoye, 1986). The majority of prevalence data are based on abattoir surveys (FAO, 1992). Studies carried out in Zaria abattoir, Northern Nigeria recorded high prevalence rates of 56.0% (Schillhorn van Veen et al., 1980) and 35.4% (Ulayi et al., 2007) from cattle for *D. hospes*. A low prevalence rate of 18.3% in Borno State (Nwosu & Srivastava, 1993), and 22.33% was also reported in Plateau State in bile duct of cattle (Omowaye et al., 2012). Seasonal infection rates have also been reported for of *D. hospes* in Nigeria, with highest rate occurring during and directly after the rainy season (Schillhorn van Veen et al., 1980).

Epidemiology of bovine schistosomiasis in Nigeria

*Schistosoma curassoni* and *S. bovis* have been reported in cattle in Nigeria (Ndifon et al., 1988, Elelu et al., 2016). The intermediate snail host (*Bulinus globosus*) of *Schistosoma* species is present in Nigeria (Ndifon and Ukolı, 1989) and has been successfully experimentally infected with *S. bovis* miracidia originating from a Nigerian cow (Ndifon et al., 1988). The transmission of *S. mattheei* has been reported with an all year round occurrence with high prevalence during the wet months (Pfukenyi et al., 2005). The prevalence rates reported in Nigeria were 7.8% for *S. bovis* and 2.2% for *S. curassoni* based on examination of rectal scrapings, however a higher prevalence of 31.1% (including both species of *Schistosoma*) was observed in 502 slaughtered cattle by examination of mesenteric and rectal veins (Ndifon et al., 1988). In another study that utilised a more sensitive ELISA technique, a higher prevalence rate (16.2%) was recorded in cattle of the Kuri breed compared with other breeds (Hambali et al., 2016). This is not surprising as these breed of cattle with their characteristic bulbous horns are adapted to swimming, they are therefore more likely to come in contact with *Schistosoma* infected water. Schistosomiasis is transmitted via active skin penetration during contact with water. Other studies carried out in parts of North-Western Ethiopia located close to swampy areas reported higher prevalence rates.
up to 24% in cattle (Lulie & Guadu, 2014). In addition, significantly higher schistosomiasis prevalence was observed in the wet season compared to the dry season in cattle in Zimbabwe (Pfukenyi et al., 2006b). Higher prevalence rate (25.2%) has also been reported in cattle managed extensively compared to those under a semi-intensive management system (15.38%) (Lulie & Guadu, 2014).

Molecular identification of trematode species

Several PCR-based techniques have been used to identify digenetic trematodes (Lotfy et al., 2008, Lotfy et al., 2010). Molecular identification has an important contribution to make in many areas where there are problems associated with definitive speciation. These areas include instances of species overlap as seen in Fasciola (Kendall, 1965), the existence of hybrids between different genotypes (for example, the Japanese triploid forms (Itagaki & Tsutsumi, 1998), the identification of sexually immature paramphistome species (Horak, 1971), and difficulty in speciation based on morphology of parasite eggs (Itagaki et al., 2003). Other reasons for molecular identification are the close relatedness between taxa as seen between S. haematobium and S. bovis (human and animal parasites) as well as difficulty in cercarial identification (Webster et al., 2010).

Molecular studies by analysis of the first (ITS1) and second (ITS2) internally transcribed spacer of the ribosomal rDNA and the mitochondrial cytochrome c oxidase I (COI) gene has been carried out for Fasciola (Mas-Coma et al., 2005, Ali et al., 2008, Amor et al., 2011, Amer et al., 2011), paramphistome (Lotfy et al., 2010) and Dicrocoelium (Otranto et al., 2007). The ITS-1 and ITS-2 are highly conserved and useful for differentiating closely related taxa that have diverged relatively recently (<50 million years ago) (Mas-Coma & Bargues, 2009). The mitochondrial NADH dehydrogenase subunit 1 and COI were recently used to characterize F. gigantica from Nigeria (Ichikawa-Seki et al., 2017). However further molecular studies from different parts of Nigeria is recommended in order to understand these important pathogens.

Other genetic methods such as the use of restriction fragment length polymorphism (PCR-RFLP) of ribosomal or mitochondrial genes using common
restriction enzymes (such as AvaII and DraI) have been used to analyse the whole of mitochondrial DNA to distinguish Fasciola species (Marcilla et al., 2002), to identify species of paramphistomes (Itagaki et al., 2003), to study genetic variability of Dicrocoelium species (Sandoval et al., 1999), and also to distinguish between S. haematobium and S. bovis (Barber et al., 2000, Webster et al., 2010).

A polymerase chain reaction technique (PCR) has also been used to detect the F. gigantica infection status of snail intermediate hosts (Velusamy et al., 2004, Kaset et al., 2010). PCR technique used to amplify specific fragments of mitochondrial DNA in faecal samples in sheep has shown promise in early detection of F. hepatica infection (Martínez-Pérez et al., 2012). Molecular identification of paramphistome species from Asia and elsewhere in Africa has been carried out (Lotfy et al., 2010), however at time of writing this review, there are no records of such studies having been carried out in Nigeria. However, PCR to amplify species-specific genes has been used in differentiating human species of Schistosoma in Nigeria (Akinwale et al., 2014) and also to identify S. bovis in Kenya (Kamanja et al., 2011).

Control strategies for trematode infections

The principles behind the control of trematode infections are similar for all species (FAO, 1993). Conventional methods for control include strategic anthelmintic treatment to reduce environmental contamination, (FAO, 1993), snail eradication by use of molluscicides and improved drainage systems to adversely influence the snail habitats(Armour, 1975, De Bont & Vercruysse, 1997). The current control of F. hepatica in cattle in temperate regions where the disease is prevalent is often based on strategically timed flukicide treatment, which is determined by studying the seasonal transmission dynamics in numerous location throughout the world (Kaplan, 2001). The tropical trematode infection caused by F. gigantica may be less amenable to this approach because the intermediate hosts L. (Radix) natalensis are true water snails (Sewell, 1966) and cattle can come into contact with infected snails while grazing around water bodies all year round. Control of Dicrocoelium infection is more difficult due to the complex life cycle involving terrestrial intermediate hosts. Control is therefore based on
chemotherapy and husbandry practices (avoiding grazing during early morning and late evening to avoid infective ants) (Otranto & Traversa, 2002).

**Strategic chemotherapy**

Strategic anthelmintic treatment based on epidemiological and meteorological data is important for the control of flukes (FAO, 1993). There are several drugs for the treatment of fasciolosis. These include: halogenated phenol (niclofolan, bithionol, hexachlorophene and nitroxynil); salicylanides (rafoxanide, oxyclozanide and closantel), benzimidazoles (triclabendazole and albendazole); sulphonamides (clorsulon), phenoxyalkanes (diamphenethides) (Fairweather & Boray, 1999a). Triclabendazole has been the preferred drug for treating liver fluke since 1983 due to its high efficacy against early immature, immature and adult flukes (Boray et al., 1983). These drugs differ in their effectiveness against adults and immature flukes. Strategic antihelminthic use at 12-13 week intervals is effective against both mature and immature flukes and reduces intensity of infection over time. Seasonal occurrence of fasciolosis and suggestions for strategic antihelminthic treatments has been suggested of which in the tropics with fasciolosis outbreak all year round, treatment up to 4 times per year is recommended (Torgerson & Claxton, 1999). In Nigeria, some parts have reported seasonal trends in fasciolosis while some Southern parts of the country have reported an all year round occurrence (Gboeloh, 2012). Some authorities recommend that cattle should be dewormed regularly (Aliyu et al., 2014), while others recommend treatment upon onset of clinical fasciolosis. Damwesh and Ardo (2015) proposed 2-3 annual treatments: at the start of the rainy season, mid rainy season and at the start of the dry season.

The anthelmintic drugs currently in use in Nigeria include: albendazole, nitroxynil, closulon and levamisole. However, a search in available literatures revealed that the preferred drug of choice against Fasciola (triclabendazole) is currently not available for use in Nigeria.

Neither triclabendazole nor niclofolan have been found to be effective against Dicrocoelium and paramphistome species (Güralp & Tina, 1984). The drugs of choice for paramphistome infection are resorantel, oxyclozanide and the combination of bithional and levamisole (Aiello, 1998). A combination of
oxyclozanide and levamisole is most effective against paramphistomosis in cattle given in two treatments 3 days apart (Rolfe & Boray, 1987). Strategic treatment during the dry season may reduce contamination of snail habitat in the following raining season (Rolfe et al., 1991).

Benzimidazoles (except triclabendazole) and pro-benzimidazoles (thiophanate, netobimin) at higher doses are effective against Dicrocoelium species (Otranto & Traversa, 2002). Praziquantel is highly effective against all bovine visceral schistosomiasis (De Bont & Vercruysse, 1997) but is advised only in severe outbreaks due to the risk of portal occlusion from heavy worm burdens (McCully & Kruger, 1969). A combination of praziquantel and artemether is also effective against mature and immature schistosomes (Pfunkenyi et al., 2005).

**Anthelmintic drug resistance**

Resistance of parasites to anthelmintic drugs is a growing global problem (Wanyangu et al., 1996). Anthelmintic drug resistance is a real problem in the UK (Gordon et al., 2012) and Australia (Brockwell et al., 2014) and has also been reported in human in the Netherlands (Winkelhagen et al., 2012). Anthelmintic resistance has been demonstrated using faecal egg count reduction (Coles et al., 1992), copro-antigen reduction tests (Brockwell et al., 2014) and egg hatch assay (Fairweather et al., 2012). Demonstration of resistance to triclabendazole TCBZ (also the drug of choice in human fasciolosis) against immature and adult *Fasciola* indicates serious potential problems in controlling fasciolosis in future (Fairweather & Boray, 1999b). Burdens of 20–34 drug-resistant flukes were observed in cattle following treatment with triclabendazole in Australia (Brockwell et al., 2014). However, studies have shown that, the anthelmintic drugs, nitroxynil and oxyclozanide were able to kill 100% and 99.6% of adult triclabendazole-resistant flukes respectively. Also albendazole caused up to 95% reduction in triclabendazole-resistant fluke egg counts while clorsulon showed 73.2% reduction (Coles & Stafford, 2001). This finding is consistent with those carried out recently that showed that the anthelmintic drugs, clorsulon and oxyclozanide were effective in removing adult triclabendazole-resistant flukes (Elliott et al., 2015).

There are also reports of some isolates of *F. hepatica* being resistant to
albendazole in sheep (Sanabria et al., 2013, Novobilsky et al., 2012). A reduced efficacy of albendazole and oxyclozanide against F. gigantica in naturally infected cattle has also been reported in Tanzania (Keyyu et al., 2008). This could probably be the case with F. gigantica in Nigeria, hence further validation tests may be needed. There are currently few if any data on anthelminthic drug resistance in Nigerian cattle. However, a recent study in small ruminants suggests low resistance to ivermectin and levamisole with susceptibility to albendazole (Adediran and Uwalaka, 2015)

The incomplete elimination of infection leads to subclinical fasciolosis with continuous contamination of pasture especially in sheep (Ollerenshaw, 1971). An alternative strategy is to use a combination of drugs-synergistic drug usage (Fairweather & Boray, 1999b) and also treatment based on degree of infection (Malone & Craig, 1990) may be practical treatment strategies.

**Human trematodiasis in Nigeria**

Human fasciolosis has been reported from 51 countries over the last 25 years in the continents of Africa, America, Asia, Europe and Oceania (Mas-Coma et al., 1999). Fasciolosis is among the most neglected tropical diseases (Mas-Coma et al., 2009) with an estimated 180 million humans at risk of Fasciola species infection worldwide (WHO, 1995), and with as many as 2.4 to 17 million humans which may be infected (Hopkins, 1992, Toledo et al., 2012). Human infection with fasciolosis have been reported to occur from eating uncooked watercress derived from endemic areas where infected cattle range freely, and also probably from contaminated water (Stemmermann, 1953, Toledo et al., 2012). Man is not a natural host of F.gigantica and most flukes derived from human beings are not fertile. As such coprological examination is not likely to be of value and only symptomatic cases would be discovered (Stemmermann, 1953). Human fasciolosis is determined by the presence of the intermediate snail host, domestic herbivorous animals, climatic conditions and the dietary habits of man (Chen and Mott, 1990). Mas-Coma et al., (1999) reviewed the prevalence of Fasciola species reported in various parts of the world (Mas-Coma et al., 1999).

The main methods of diagnosing human trematodiasis are direct parasitological
detection of fluke eggs in stool (Mas-Coma et al., 2006) and other biofluids-
duodenal and biliary aspirates) and immune-diagnosis (indirect diagnosis). Other
non-invasive diagnostic techniques such as radiology, ultrasound, computer
tomography and magnetic resonance can also be used (Esteban et al., 1998, Mas-
Coma et al., 2005, Keiser & Utzinger, 2009). In a recent review, stool and blood
techniques have been reported to be improved for diagnosis of human fasciolosis
and during surveys. However the author also identified difficulties of diagnosing
fascioliasis in humans due to different infection phases and parasite migration
capacities, clinical heterogeneity, immunological complexity, different
epidemiological situations and transmission patterns (Mas-coma et al., 2014).

Human fasciolosis have been reported in Africa involving both species of
Fasciola (Mas-Coma et al., 1999). There are hardly any studies targeted at
detecting the zoonotic effect of fasciolosis in Nigeria. In a study there was a report
of the positive detection of Fasciola species in HIV-infected human subjects in
Nigeria though at low prevalence rate of 1% (Abaver et al., 2012). This suggests
that human infection may be significant although further research on risk factors
as well as possible routes of transmission needs to be carried out.

Human cases of dicrocoeliasis are rare and often occur by accidental ingestion of
infected ants on unwashed vegetables or by drinking contaminated water (Haridy
et al., 2003). Human cases of dicrocoeliasis have been reported in Nigeria (Roche,
1948, Samaila et al., 2009), Czechoslovakia (Ondriska et al., 1989), Egypt
(Massoud et al., 2003), Turkey (Cengiz et al., 2010) and Kyrgyzstan (Jeandron et
al., 2011). Spurious infection with D. hospes has also been reported in Ghana due
to accidental ingestion of infected animal liver (Wolfe, 1966). The subclinical
form of the disease, which is most common, is characterized by cholangitis and
adenomatous proliferation of the bile duct (Cabeza-Barrera et al., 2011). Other
signs are constipation, diarrhoea, vomiting and abdominal pain (Ondriska et al.,
1989). Human dicrocoeliasis has been reported in a 7-year old Nigerian child of a
nomadic pastoralist showing clinical signs of fever, jaundice and anterior
subcutaneous abdominal mass (Samaila et al., 2009).
The amphistome species reported in man is *Gastrodiscoides hominis* and is widely distributed in Asia. It is located in the caecum and colon, with pigs being the main animal reservoir (Dutt & Srivastava, 1972) and has been reported in Nigeria in a malnourished child (Dada-Adegbola *et al*., 2004).

Although *S. bovis* is primarily infects ruminant, it has been isolated in man in various parts of Africa (Raper, 1951, Chunge *et al*., 1986, Santoro, 1988, Kinoti & Mumo, 1988). Moreover, the mesenteries of cattle, which may be infected with *S. bovis*, are usually sold at the local markets in Africa as edible tripe thus making it public health risk (Kamanja *et al*., 2011). Hybridization between *S. bovis* and *S. haematobium* (cattle and human infection) in Senegal has been reported with an impending risk of the emergence of a new disease (Huyse *et al*., 2009).

Several drugs for the effective treatment of human fasciolosis are available. These include dihydroemetine-emetine derivative, bithionol, praziquantel and triclabendazole (effective against both acute and chronic fasciolosis). These are currently the drugs of choice for human fasciolosis caused by both *F. hepatica* and *F. gigantica* (Esteban *et al*., 1998, Savioli *et al*., 1999). Praziquantel has also been used in treatment of human dicrocoeliasis (Massoud *et al*., 2003). Mebendazole was effective in treatment of the human amphistomosis (Dada-Adegbola *et al*., 2004).

The prevention of human trematode infections may be achieved by strict control of water cress and other metacercariae carrying aquatic plants for human consumption especially in endemic zone (Mas-Coma *et al*., 2005). Drinking water must be boiled or purified. Integrated control approaches and inter-sectoral collaboration between public health and veterinary medicine has also been suggested for control (Keiser & Utzinger, 2009).

**Conclusions and recommendations.**

Trematode infections are a significant limiting factor in livestock production; therefore development of sustainable strategies for their control is a priority. In order to develop sustainable control, gap in knowledge must be identified to guide research projects. The outcome of such research would enhance proper allocation
of funds for disease control and policy formulation. In addition, it is also important to carry out detailed study on the overall economic losses due from trematode infections in order to design and implement appropriate systematic disease prevention and control methods.

The prevalence rates of trematodes reported across the various abattoirs in Nigeria are an indication that trematode infection poses widespread risk to livestock and possibly humans. However, it is worthy of note that neither oxyclozanide nor triclabendazole, which are the 2 drugs of choice for mixed and human *Fasciola* infection respectively is currently not available for use in Nigeria. Availability of these two drugs could be helpful in reducing diseases occurrence. These drugs could be made available by formulating favourable policies that ensure the drug retailers have basic training to identify and stock the appropriate drugs of choice and informing livestock farmers on rationale for use (example is information on effectiveness of triclabendazole against both mature and immature flukes). The gap in technical knowledge needed by drug retailers for dispensing drugs has been identified in a previous study (Bett *et al.*, 2004) and veterinarians have a huge responsibility in advising/training these retailers on the appropriate drugs to stock based on the diseases prevalent to different localities.

There are hardly any studies targeted at detecting the zoonotic effect of fasciolosis infections in Nigeria. Although a few studies exist on other trematodes in Nigeria, these are often case reports. Further research on burden, risk factors as well as possible routes of transmission of trematode infection to human needs to be carried out.

Molecular studies on trematodes in different species of livestock including those targeted at understanding anthelmintic drug resistance should also be carried out across Nigeria. These studies would be useful in providing more data on the diversity of these important trematodes.

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Statement of interest

None

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Figure 1: Geographical distribution of bovine fasciolosis across different states of Nigeria (1980–2016) from published prevalence (%) data based on abattoir records, liver and coprological examination. States with 0% have no available data as at time of review.