

Helminth parasites of laboratory mice and rats

Agnieszka Perec-Matysiak, Anna Okulewicz, Joanna Hildebrand and Grzegorz Zaleśny

Department of Parasitology, Institute of Genetics and Microbiology, Wrocław University, Przybyszewskiego 63, 51-148 Wrocław

Corresponding author: Agnieszka Perec-Matysiak, Department of Parasitology, Institute of Genetics and Microbiology, Wrocław University, Przybyszewskiego 63, 51-148 Wrocław; E-mail: agap@microb.uni.wroc.pl

ABSTRACT. Rodents, as mice and rats are the most common laboratory animals used in research and testing. They are seldom investigated for autochthonous ecto- and endoparasites prior their utilization in the experiments. Helminth parasites can alter the interpretation of final results. Pinworms commonly infecting laboratory rodents include mainly the mice pinworms *Syphacia obvelata* and *Aspicularis tetraptera*, and in rats *Syphacia muris*. The fact that many laboratory rodent colonies were found to be parasite contaminated suggests a need for eradication and improvement of the quality of laboratory rodents. This review reports the data on the presence of helminth parasites in laboratory rodent colonies, and suggests to pay special attention on controlling the sanitary conditions of animal houses.

Key words: helminths, laboratory rodents, pinworms.

Development of many biological assays still depends on the use of living laboratory animal models. Rodents, as mice and rats are the most common laboratory animals used in research and testing. Experimental results from research performed with living laboratory animals may be affected by environmental conditions provided to breeding and maintaining of these animals and by infectious agents [1-3]. In some papers it is showed that infected animals are not suitable for biomedical researches as well as for helminth biology experiments [4, 5].

The parasite infections can affect investigations by inducing physiological and immunological alterations in the hosts, increasing or diminishing host susceptibility to experimental stress, inducing tissue damages, stimulating abnormal tissue growth, competing with the host for nutrients, decreasing the volume of host's blood and body fluids and by mechanical interference [6].

This review reports the data on the presence of helminth parasites, mainly with regard to pinworm species, and its interference with research in laboratory rodent colonies. It is suggested to pay special

attention on controlling the sanitary conditions and barriers of animal houses.

Still little is known about the effects of environmental changes on the biological variation in experimental results [7] as well as laboratory rodents are seldom investigated for autochthonous ecto- and endoparasites prior their utilization in the experiments [8, 9]. The control or eradication of worm burdens in laboratory animals ensures the proper procedures in scientific research.

Among helminth parasites infecting laboratory animals the most common are pinworms belonging to the family Oxyuridae. Rodent pinworms display some host-specificity, although some can cross species barriers. In general *Syphacia obvelata* and *Aspicularis tetraptera* are considered as mouse pinworms, *Syphacia muris* the rat pinworm, *Syphacia mesocricetus* the hamster pinworm and *Denstomella translucida* the gerbil pinworm [10, 11]. *S. obvelata* has also been reported to infect humans [12]. It is common high even in well-managed animal colonies [8]. While pinworm infections are usually subclinical, rectal prolapse, intussusception, faecal impaction, rough hair coat and poor weight gain

have been reported in heavily infected rodents [10, 13]. Very heavy parasite loads may lead to catarrhal enteritis, liver granulomas and perianal irritation [10]. There are a few reports documenting how pinworms infection influenced experimental results. A significant reduction of activity of mice infected with *S. obvelata* was observed in behavioral studies [14]. Wagner et al. [15] reported definite growth differences between pinworm-free and pinworm-infected rats. *S. muris* experimentally infected animals grew slower than uninfected animals. It was reported that in laboratory rats infected with *S. muris*, despite of absence of apparent histopathological changes, intestinal transport of water and electrolytes is significantly decreased due to pinworm infection [16]. Additionally infection with *S. muris* retarded the growth of young mice and accelerated the development of their hepatic monooxygenase system [17].

Immunity to infection is mostly humoral as for many other helminthoses. In this regard, studies carried out by Sato et al. [18] reported that antibodies against *S. obvelata* somatic antigen were detected in experimental infection of pinworms in mice. Additionally pinworm infection altered higher antibody production to nonparasitic antigenic stimuli [18]. Moreover study of Beattie et al. [19] showed that infection with *S. obvelata* induces a proliferation of T- and B-lymphocytes in spleen and lymph nodes and occasional germinal center formation. In addition athymic mice infected with pinworms develop a lymphoproliferative disorder which eventually leads to lymphoma [4]. These studies indicate that infection with this parasite can modulate the immune system of the host and affect experimental final results [8, 18, 20].

Data on the prevalence and mean intensity of the helminth fauna recovered from outbred and inbred mice conventionally maintained in Brazilian animal houses were presented by Bazzano et al. [20]. As it was found out the frequency of *S. obvelata* in laboratory animals ranged from 9 to 74% (intensity 13–67 specimens/host) and *A. tetraptera* from 17 to 83% (intensity 6–17 specimens/host). Infections due to a single species were observed in 62% of the animals, compared to 16% related to associations. In the latests study carried out by us [21] the prevalence of infection was as high as 66% however the intensity ranged from 1 to 250 specimens. According to Taffs [22] the prevalence of *S. obvelata* infection in laboratory animals may be a function of age (the intensity diminishes with increasing age

of the host), sex (higher prevalence in male hamsters than females), strain and the host status.

In Poland *S. obvelata* was noted several times in laboratory mice colonies, e.g. by Szymańska et al. [23] and Obruśnik and Stankiewicz [24] — in 52% of mice in open conventional colonies, and in 5.5–15% in closed colonies respectively. On the contrary, the reports on *A. tetraptera* infections are not so common. Klauska and Złotorzycka [25] demonstrated 20% prevalence of laboratory mice infected with this nematode. Our previous examinations of 176 laboratory mice (BALB/c strain) [26] demonstrated that the prevalence of *S. obvelata* in males was 70% (with the intensity of infection of 1–102) and in females was 52% (with the intensity of 1–34 individuals). In our other mice colony of BALB/c strain which was obtained from different laboratory, the invasion of *A. tetraptera* developed rapidly. Examinations of 40 mice showed the prevalence as high as 75% (with the intensity of 1–193 individuals).

A parasitological survey of laboratory mice of 18 different strains from three animal facilities in Wrocław, Poland was presented by Pacoń and Piekarska [27]. Coproscopic examinations of mice revealed the presence of pinworm eggs in every colony. The prevalence of *S. obvelata* ranged from 5.5% to 15% while *A. tetraptera* infection was higher ranging from 33% to 75%. Additionally the necropsy of three mice from one colony demonstrated the presence of *Taenia taeniaeformis* cysticerci — the tapeworm, which in adult stage parasitises cats.

A study carried out by Gilioli et al. [28] showed that parasite infections were present in most of the 18 Brazilian animal houses investigated. A high proportion of animals infected with four or more parasite species was observed in a single group of six animals, indicating that the colonies were heavily infected. As far as only helminth infections in rat colonies are concerned the prevalence of *Syphacia muris* was as high as 80%, *Trichosomoides crassicauda* 55.5% and *Hymenolepis nana* 40% respectively. Prevalence of helminths in mice colonies was as follows: *Syphacia obvelata* 86.6%, *Aspicularis tertaptera* 60% and *H. nana* 53.3% respectively.

A parasitological monitoring of mice and rats facilities at the Central Institute of Experimental Animals of Japan was carried out in years 1996–1998. The inspection of mice facilities showed the detection of parasites in 125 of 444 facilities. The number of positive facilities with respect to helminths as *Syphacia obvelata* and

Aspiculuris tetraptera was in 14% and 7% respectively [29].

It is obvious that the scientists should no longer minimize the influence of parasites on experimental results. Because the measures taken to combat parasitic contamination in laboratory mice and rats have conspicuously lagged behind those taken against other pathogens e.g. bacteria, viruses and protozoa, parasites may currently be found even in animal facilities that appear to be well maintained without any problems. Ultimately pinworms often serve as possible indicators of biologically contaminated animal facilities and it is advised that pinworm existence and species name should be clearly indicated in the health monitoring reports [29]. The fact that many laboratory rodent colonies were found to be parasite contaminated [30], suggests a need for eradication and improvement of the quality of laboratory rodents. Works of Flynn [31] and Klement et al. [32] revealed that pinworm infection is difficult to control because anthelmintics like ivermectin eliminate adult worms but have no effect on ova, which can survive *ex vivo* for prolonged periods. Proposed much earlier by Wescott et al. [33] rigid sanitary procedures using for example filter hoods to prevent aerosol transmission and regular ova examinations which may control the parasitism did not find the practical application. Among many recommended methods for elimination of helminths one involve mixing anthelmintic with food or spraying the animals or bedding with anthelmintic as well as cleaning the colony using embryo transplant method [29]. But still prevention and control of parasitic infections in laboratory animals remains a challenge and it is another topic for review and discussion.

References

- [1] Clough G. 1982. Environmental effects on animals used in biomedical research. *Biology Research* 57: 487–523.
- [2] Pakes S.P., Lu Y.S., Meunier P.C. 1984. Factors that complicate animal research. In: *Laboratory animal medicine*. (Eds. J.G. Fox, B.J. Cohen, F.M. Loew). New York, Academic: 649–665.
- [3] Homberger F.R., Thomann P.E. 1994. Transmission of murine viruses and mycoplasma in laboratory mouse colonies with respect to housing conditions. *Laboratory Animals* 2: 113–120.
- [4] Baird S.M., Beattie G.M., Lannom R.A., Lipsick J.S., Kaplan N.O. 1982. Induction of lymphoma in antigenically stimulated athymic mice. *Cancer Research* 42: 198–206.
- [5] Ito A. 1982. *Hymenolepis nana*: immunogenicity of a lumen phase of the direct cycle and failure to auto-infection in BALB/c mice. *Experimental Parasitology* 54: 113–120.
- [6] Hsu C.K. 1980. Parasitic diseases: how to monitor them and their effects on research. *Laboratory Animals* 14: 48–53.
- [7] Baumans V. 1998. Environmental enrichment: practical applications. *Proceedings of the 2nd World Congress on Alternatives, Utrecht*: 187–191.
- [8] Pinto R.M., Vicente J.J., Noronha D., Goncalves L., Gomes D.C. 1994. Helminth parasites of conventionally maintained laboratory mice. *Memórias do Instituto Oswaldo Cruz, Rio de Janeiro* 89: 33–40.
- [9] Pinto R.M., Goncalves L., Noronha D., Gomes D.C. 2001. Worm burdens in outbred and inbred laboratory rats with morphometric data on *Syphacia muris* (Yamaguti 1935) Yamaguti, 1941 (Nematoda, Oxyuroidea). *Memórias do Instituto Oswaldo Cruz, Rio de Janeiro* 96: 133–136.
- [10] Baker D.G. 1998. Natural pathogens of laboratory mice, rats, and rabbits and their effects on research. *Clinical Microbiology Reviews* 11: 231–266.
- [11] Percy D.H., Barthold S.W. 1993. Pathology of Laboratory Rodents and Rabbits. *Iowa State University Press*.
- [12] National Research Council. 1991. Infectious diseases of mice and rats: a report of the Institute of Laboratory Animal Resources Committee on Infectious Diseases of Mice and Rats. National Academy Press, Washington, D.C.
- [13] Jacobson R.H., Reed N.D. 1974. The thymus dependence of resistance to pinworm infection in mice. *Journal of Parasitology* 60: 976–979.
- [14] McNair D.M., Timmons E.H. 1977. Effects of *Aspiculuris tetraptera* and *Syphacia obvelata* on exploratory behavior of an inbred mouse strain. *Laboratory Animal Science* 27: 38–42.
- [15] Wagner M. 1988. The effect of infection with the pinworm (*Syphacia muris*) on rat growth. *Laboratory Animal Science* 38: 476–478.
- [16] Lübcke R., Hutchenson F.A.R., Barbezat G.O. 1992. Impaired intestinal electrolyte transport in rats infested with the common parasite *Syphacia muris*. *Digestive Diseases Sciences* 37: 60–64.
- [17] Mohn G., Philip E.M. 1977. Effects of *Syphacia obvelata* on the microsomal monooxygenase system in mouse liver. *Laboratory Animals* 15: 89–95.
- [18] Sato Y., Ooi Monaka O.K., Nonaka N., Oku Y., Kamiya M. 1995. Antibody production in *Syphacia obvelata* infected mice. *Journal of Parasitology* 81: 559–562.
- [19] Beattie G.M., Braid S.M., Lipsick J.S., Lannom R.A., Kaplan N.O. 1981. Induction of T- and B-lymphocyte responses in antigenically stimulated athymic mice. *Cancer Research* 41: 2322–2327.

- [20] Bazzano T., Restel T.I., Pinto R.M., Gomes D.C. 2002. Patterns of infection with the nematodes *Syphacia obvelata* and *Aspicularis tetraptera* in conventionally maintained laboratory mice. *Memórias do Instituto Oswaldo Cruz, Rio de Janeiro* 97: 1–5.
- [21] Perek A. 2005. Udział antygenów nicieni *Syphacia obvelata* (*Oxyuridae*) w powstawaniu odpowiedzi immunologicznej u myszy laboratoryjnej (szczep BALB/c). *Wiadomości Parazytologiczne* 51: 169–170.
- [22] Taffs L.F. 1976. Pinworms infections in laboratory rodents: A review. *Laboratory Animals* 10: 1–13.
- [23] Szymańska H., Piasecki A., Krysiak E., Lis A., Czarnomorska A. 1986. Zakażenia pasożytnicze i bakteryjne u myszy laboratoryjnych utrzymywanych w warunkach hodowli konwencjonalnej otwartej i zamkniętej. *Zwierzęta Laboratoryjne* 23: 65–71.
- [24] Obruśnik A., Stankiewicz D. 1990. Kontrola parazytologiczna myszy laboratoryjnych w hodowli konwencjonalnej otwartej. *Zwierzęta Laboratoryjne* 27: 71–75.
- [25] Klaus E., Złotorzycka J. 1979. Pasożyty zewnętrzne i wewnętrzne myszy (*Mus musculus* L.) hodowanych i dziko żyjących. *Wiadomości Parazytologiczne* 25: 295–299.
- [26] Okulewicz A., Perek A. 2003. Parasite infections in laboratory mice colonies. *Polish Journal of Veterinary Sciences* 6: 51–53.
- [27] Pacoń J., Piekarska J. 2004. Pasożyty myszy w wybranych hodowlach zwierząt laboratoryjnych. *Wiadomości Parazytologiczne* 50: 93–94.
- [28] Gilioli R., Sakurada J.K., Andrade L.A.G., Kraft V., Meyer B., Rangel H. A. 1996. Virus infection in rat and mouse colonies reared in Brazilian animal facilities. *Laboratory Animal Science* 46: 582–584.
- [29] Shibahara T. 1999. Necessity of reexamining the pathogenicity and elimination of parasites in rats and mice. Microbial status and genetic evaluation of mice and rats. *Proceedings of the 1999 US/Japan Conference*: 21–26.
- [30] Casebolt D.B., Lindsey J.R., Casell G.H. 1998. Prevalence rates of infectious agents among commercial breeding populations of rats and mice. *Laboratory Animal Science* 38: 327–329.
- [31] Flynn R.J. 1973. Nematodes, parasites of laboratory animals. Iowa State University Press, Ames Iowa: 203–320.
- [32] Klement P., Augustine J.M., Delaney K.H., Klement G., Weitz J.I. 1996. An oral ivermectin regimen that eradicates pinworms (*Syphacia* spp.) in laboratory rats and mice. *Laboratory Animal Science* 46: 286–290.
- [33] Wescott R.B., Malczewski A., Van Hoosier G.L. 1976. The influence of filter top caging on the transmission of pinworm infections in mice. *Laboratory Animal Science* 26: 742–745.

Wpłynęło 23 listopada 2005,
Zaakceptowano 6 marca 2006