

Behavioral Core Protocols and Training

Rotarod

Basics

This is a test of motor coordination and motor learning [1-4]. In addition, it can be used to assess intoxication [5], sedation and strength / stamina [6].



The latency to fall from a rotating rod is scored automatically with infrared sensors in a Rotamex 5 rotarod (Columbus Inst; Columbus, Ohio). Motor coordination can be tested by comparing the latency to fall on the very first trial between treatment groups [7]. Motor learning can also be assessed by comparing the first trial with subsequent trials and is evident as an increased latency to fall over time [3, 4, 8-13].

Motor coordination can be tested by comparing the latency to fall on the very first trial between treatment groups [7, 9]. Motor learning can also be assessed both within and between subjects by comparing the first trial with subsequent trials and is evident as an increased latency to fall over time [3, 4, 8-13].

Procedure

The acceleration step and time should be determined empirically. A rough starting point is : increased by 0.5 cm./sec. every 5 sec. If the step is too fast and too soon, there will be a floor effects (the controls will not be able to stay up or get better) and if it is too slow and too long between steps you will have a ceiling effect (deficits will not be detected because the task is too easy).

In general, the mice require 4-6 trials per day for 3-6 days to see significant improvement in controls.

Data Analysis and Illustration

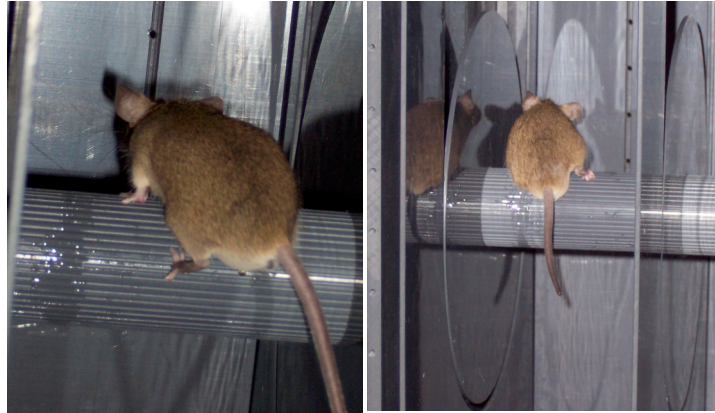
There are numerous ways of dealing with these data – some common ones are below

- 1) mean latency to fall (all trials) per day
- 2) best (or worst) latency to fall per day

3) All trials represented individually

Variants

There are numerous procedural variations to those stated above [11]. It is also possible to test reference motor memory (long term, skill or procedural memory) by stopping the trials for a period of time and then testing the animals to see if the latency to fall when re-introduced is significantly lower than it was on



the last trial. In some instances, a single speed (non-accelerating) protocol is also used [14].

Data Analysis

The most common form of analysis for these data is repeated measures ANOVA. Although this test is very powerful, it can be complicated (or impossible) to perform the test if all subjects have not completed all tests (i.e. missing variables). The basis of a repeated measures (within subject design) test is that the data are essentially paired and thus statistical programs cannot handle missing variable in such tests without interpolation.

Useful References

Variations

[9, 14-16]

Other

[8, 17-23]

References

1. Jones, B.J. and D.J. Roberts, *The quantiative measurement of motor inco-ordination in naive mice using an acelerating rotarod*. J Pharm Pharmacol, 1968. 20(4): p. 302-4.
2. Jones, B.J. and D.J. Roberts, *A rotarod suitable for quantitative measurements of motor incoordination in naive mice*. Naunyn Schmiedebergs Arch Exp Pathol Pharmacol, 1968. 259(2): p. 211.
3. Pritchett, K. and G.B. Mulder, *The rotarod*. Contemp Top Lab Anim Sci, 2003. 42(6): p. 49.
4. Watzman, N., et al., *Semiautomatic System For Timing Rotarod Performance*. J Pharm Sci, 1964. 53: p. 1429-30.
5. Boehm, S.L., 2nd, et al., *Sensitivity to ethanol-induced motor incoordination in 5-HT(1B) receptor null mutant mice is task-dependent: implications for behavioral assessment of genetically altered mice*. Behav Neurosci, 2000. 114(2): p. 401-9.
6. Perrine, J.W. and E.I. Takesue, *Use of the rotarod in determining grip strength in rats with adjuvant-induced arthritis*. Arch Int Pharmacodyn Ther, 1968. 174(1): p. 192-8.
7. Lalonde, R., et al., *Motor coordination in mice with hotfoot, Lurcher, and double mutations of the Grid2 gene encoding the delta-2 excitatory amino acid receptor*. Physiol Behav, 2003. 80(2-3): p. 333-9.
8. Kuhn, P.L., et al., *Motor function analysis of myelin mutant mice using a rotarod*. Int J Dev Neurosci, 1995. 13(7): p. 715-22.
9. Buitrago, M.M., et al., *Short and long-term motor skill learning in an accelerated rotarod training paradigm*. Neurobiol Learn Mem, 2004. 81(3): p. 211-6.
10. Watzman, N., et al., *Influence of certain parameters on the performance of mice on the rotarod*. Arch Int Pharmacodyn Ther, 1967. 169(2): p. 362-74.
11. Rustay, N.R., D. Wahlsten, and J.C. Crabbe, *Influence of task parameters on rotarod performance and sensitivity to ethanol in mice*. Behav Brain Res, 2003. 141(2): p. 237-49.
12. Rozas, G., M.J. Guerra, and J.L. Labandeira-Garcia, *An automated rotarod method for quantitative drug-free evaluation of overall motor deficits in rat models of parkinsonism*. Brain Res Brain Res Protoc, 1997. 2(1): p. 75-84.
13. Ogura, T., et al., *Impaired acquisition of skilled behavior in rotarod task by moderate depletion of striatal dopamine in a pre-symptomatic stage model of Parkinson's disease*. Neurosci Res, 2005. 51(3): p. 299-308.
14. Bogo, V., T.A. Hill, and R.W. Young, *Comparison of accelerod and rotarod sensitivity in detecting ethanol- and acrylamide-induced performance decrement in rats: review of experimental considerations of rotating rod systems*. Neurotoxicology, 1981. 2(4): p. 765-87.
15. Cartmell, S.M., L. Gelgor, and D. Mitchell, *A revised rotarod procedure for measuring the effect of antinociceptive drugs on motor function in the rat*. J Pharmacol Methods, 1991. 26(2): p. 149-59.

16. Plotnikoff, N., D. Reinke, and J. Fitzloff, *Effects of stimulants on rotarod performance of mice*. J Pharm Sci, 1962. 51: p. 1007-8.
17. Carter, R.J., et al., *Characterization of progressive motor deficits in mice transgenic for the human Huntington's disease mutation*. J Neurosci, 1999. 19(8): p. 3248-57.
18. Hamm, R.J., et al., *The rotarod test: an evaluation of its effectiveness in assessing motor deficits following traumatic brain injury*. J Neurotrauma, 1994. 11(2): p. 187-96.
19. Hunter, A.J., et al., *Functional assessments in mice and rats after focal stroke*. Neuropharmacology, 2000. 39(5): p. 806.
20. Ivens, I., *Neurotoxicity testing during long-term studies*. Neurotoxicology and Teratology, 1990. 12(6): p. 637.
21. McIlwain, K.L., et al., *The use of behavioral test batteries: effects of training history*. Physiol Behav, 2001. 73(5): p. 705-17.
22. Mizoguchi, K., et al., *Chronic stress impairs rotarod performance in rats: implications for depressive state*. Pharmacol Biochem Behav, 2002. 71(1-2): p. 79-84.
23. Monville, C., E.M. Torres, and S.B. Dunnett, *Comparison of incremental and accelerating protocols of the rotarod test for the assessment of motor deficits in the 6-OHDA model*. J Neurosci Methods, 2006.