

1st ISBS Summer School

St. Petersburg, Russia

May 9th -15th,2008

Definitions

Grooming:

An animal's cleaning and caring for the body surface. This includes preening, the cleaning and oiling hair with the tongue

Two forms of grooming

Hetero-grooming or "allogrooming"

When an animal grooms another animal



Self-grooming or "autogrooming"

When an animal grooms itself



Grooming

- Evolutionarily ancient behavior
- Second (after sleep) in represented behaviors
- in rodents up to 30-50% of their waking time
- Highly sensitive to stress
- Bidirectionally sensitive to many pharmaceutical compounds

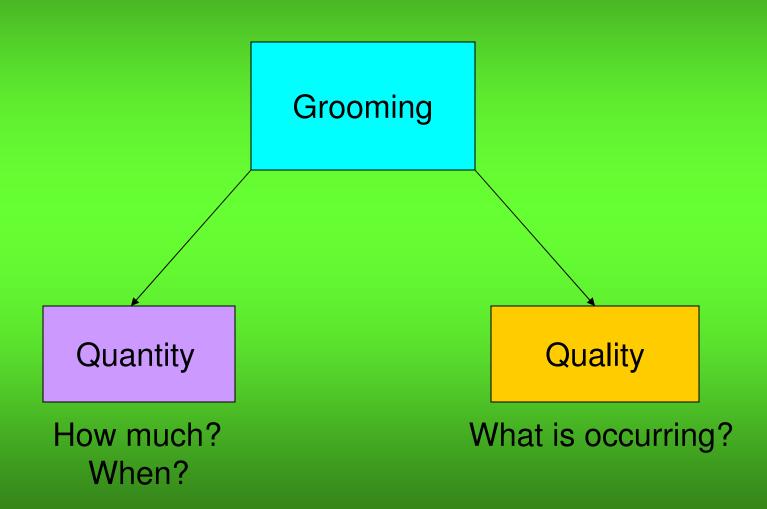
Grooming

- Has strong genetic component
- Sensitive to genetic manipulation
- Emerges as a very important behavioral endpoint for mental health research
- In many published studies, is often the only behavior affected

Biological functions of grooming

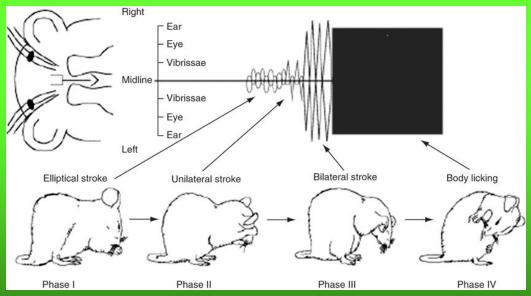
- Removing parasites
- Basic hygiene
- Thermoregulation
- Chemocommunication
- Distribution of pheromones over body surface
- Maternal care
- Establishing social hierarchy
- Displacement behavior
- Dearousal
- Social signaling ("I am not afraid")

Two aspects of grooming



Patterned grooming: "quality"

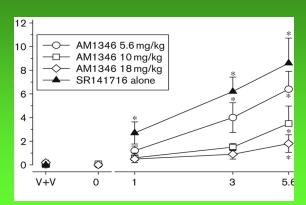
- Four distinct phases in sequential order
- Rodent cephalo-caudal pattern
- Includes more meaningful endpoints of grooming transitions, interruptions, completed sequences
- Sensitive to stress and pharmacological compounds



Elliptical stroke Unilateral stroke Bilateral stroke Body licking

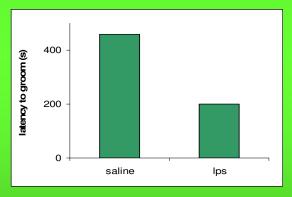
Traditional endpoints: "quantity"

 Number of bouts (frequency)



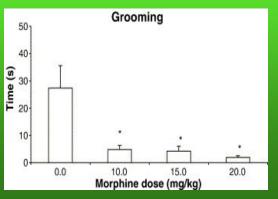
Jarbe et al., 2007

Latency to begin (s)



Spencer et al., 2006

Duration (s)



Patti et al., 2005

"Quality" of grooming

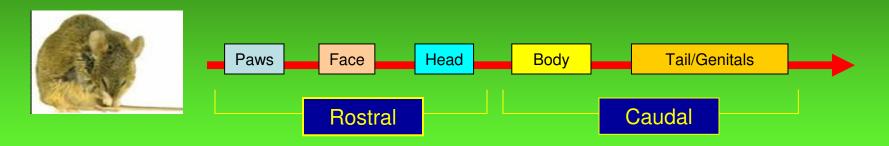
- How performed?
- What stages?
- Transition between stages?
- Regional distribution?
- Interrupted/ incomplete bouts?

Two approaches to grooming qualitative analysis

 Grooming Analysis Algorithm (GAA) (Kalueff et al., 2004). Examines grooming behavior globally

 Syntactic Chain (Berridge et al., 1987). Examines only selected types of grooming behavior, but more specifically Video clip 1 Video clip 2 Video clip 3

Grooming Analysis Algorithm (GAA)



Gross activity scores:

Frequently unaltered, or change unpredictably Is not stress: can be increased by anxiolytics

Importance of alternative approaches

Grooming patterning (GAA): Kalueff, 2001



- Transitions
- Interruptions
- % completed sequences
- regional distribution

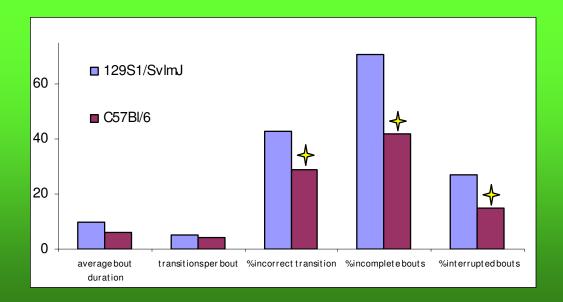


Stress disrupts behavioral organization of grooming:

- ↓ correct transitions, ↓ completed sequences,
- ↑ interrupted bouts, ↑ rostral grooming, ↓ caudal grooming

Grooming Analysis Algorithm (GAA)

- Patterning –Whether of not the grooming continues in the cephalo-caudal progression
- Interruptions Is the grooming cycle interrupted?



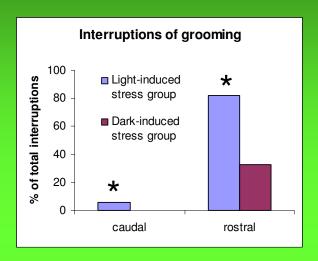
GAA Protocol

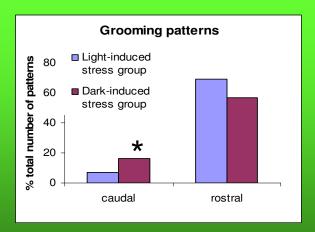
- Transfer rodents to the procedure room for acclimation 1 hr before testing
- 2) Remove the rodent from the cage and expose it to a stress that will induce grooming
- 3) Use a timer to record general cumulative measures of grooming activity, such as the latency to onset, the duration and number of grooming episodes (bouts)

GAA Protocol

- 4) Record grooming patterns for each bout:
- paw licking
- nose/face grooming (strokes along the snout)
- head washing
- body and leg grooming/scratching
- tail/genitals grooming (licking genital area and tail)
- 5) Assess the total number of transitions between grooming stages and the average number of transitions per bout

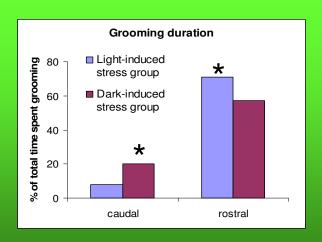
Regional distribution of grooming





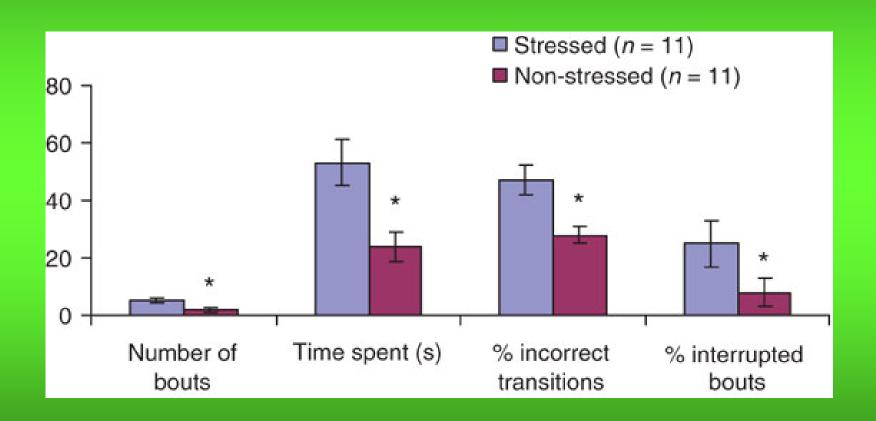
Q: Where?

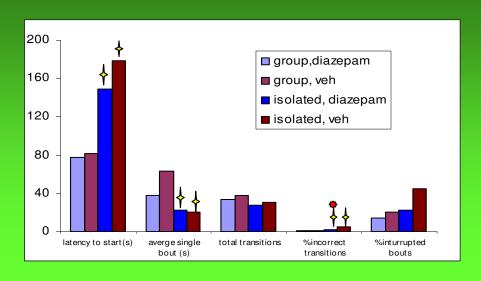
Example: High vs. low stress Wistar rats



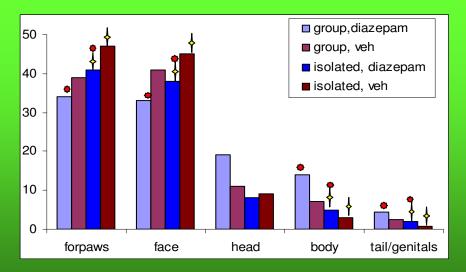
Kalueff et al., 2005

Psychological stress disrupts patterning

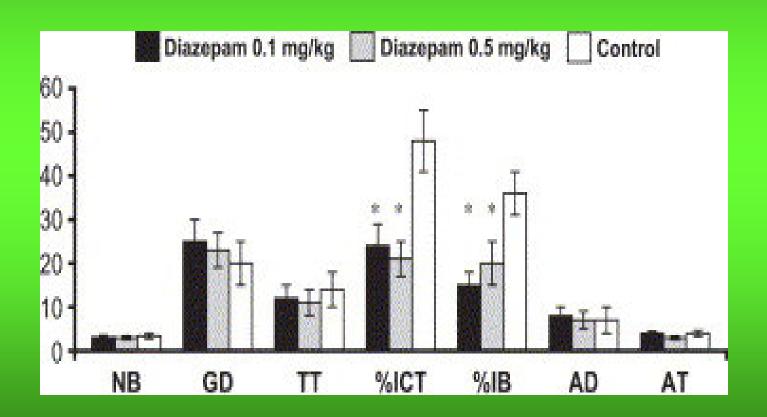




Drugs modulate patterning in rats



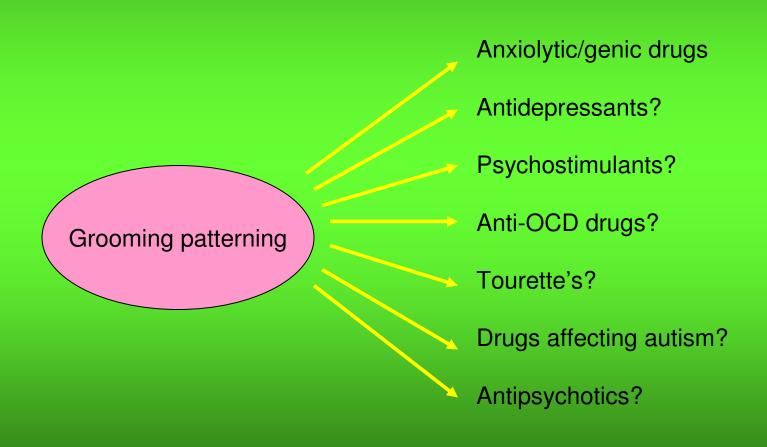
Drugs modulate patterning in mice

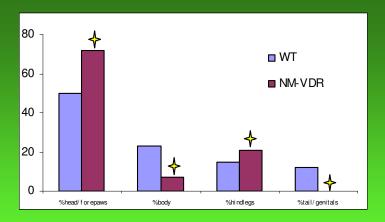


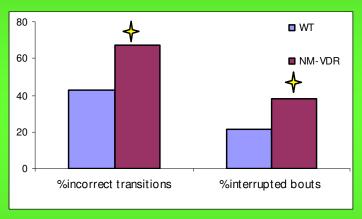
Enginar et al. (2008): Evaluation of the acute effects of amitriptyline and fluoxetine on anxiety using the grooming analysis algorithm in rats. Pharmacol Biochem Behav.

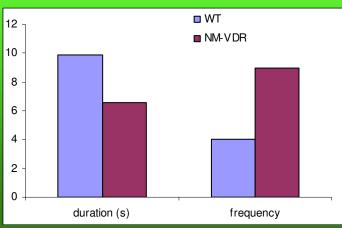
Audet et al. (2006): Subchronic Phencyclidin disorganized grooming sequences in rats, but only under "stress" conditions. The examination of grooming organization is a tool to measure stress-induced behavioral sensitization and motor functions in animal models of schizophrenia.

Developing utility of grooming analysis for drug design and discovery









Genetic mutations

e.g. VDR KO (129S1 background strain)

↑ % of "incorrect" transitions and duration of "incorrect" grooming, interrupted bouts

↑ leg grooming, ↓ body and tail/genital grooming (graphs feature novelty-induced grooming)

Syntax grooming

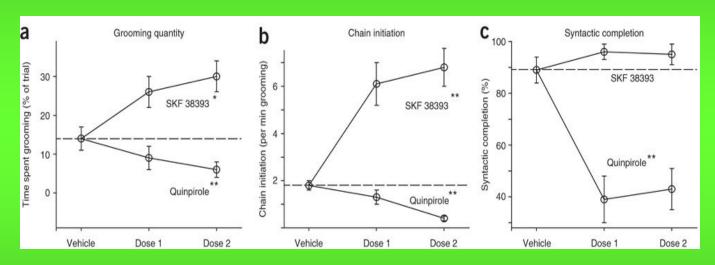
- 1) Handle the animals and put them in the recording chamber for a short period of time (10–30 min) 3–4 d before study, for acclimation to the testing environment and filming procedures
- 2) Remove the rodent from the cage and transfer it to the recording chamber
- 3) Place the rodent in a recording chamber with a clear plastic floor and a video camera to record from below
- 4) Use a frame-by-frame offline analysis of the videotapes, using both a choreographic notation system developed for detailed descriptions of stereotyped grooming sequences

Syntax grooming

- 5) Assess the number of:
- initiated syntactic chains
- the number of fully completed chains (through phase IV)
- the probability of chain initiation (the number of initiated syntactic chains per minute of grooming time)
- the probability of pattern completion once initiated (the percentage of fully completed grooming chains of total number of initiated chains)

A different approach: syntactic chain analysis

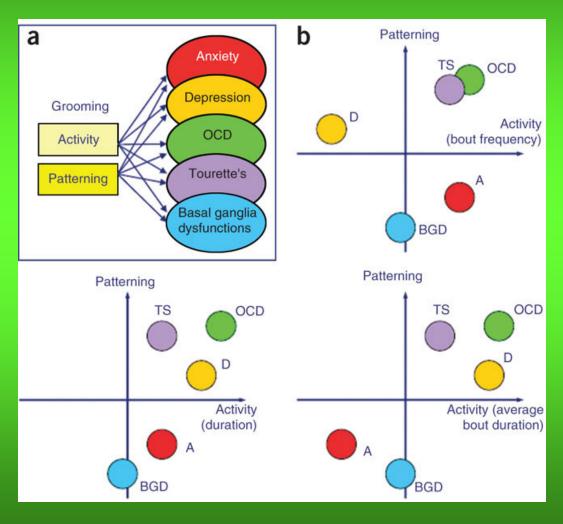
Effects of D1 (SKF 38393) and D2 (quinpirole) agonists



- (a) Amount of grooming as a percentage of total time
- (b) Syntactic chain initiation
- (c) Syntactic completion

Dose 1: SKF 38393 = 10 mg/kg, qunipirole = 5 mg/kg Dose 2: SKF 38393 = 20 mg/kg, quinpirole = 10 mg/kg

Relevance of grooming parameters to modeling human brain disorders



Genetic models: relevance to OCD

Increased grooming in SAPAP3-/- mice



Welch et al., 2007

Other OCD/grooming models:

- BTBR mice: Yang et al. 2007
- Hox8-/- mice: Greer and Capecchi, 2002

Barbering



Wurbel and Garner, 2007

Barbering: The removal or trimming of an animal's hair or whiskers by a conspecific

Two forms:

- 1) self-barbering
- 2) hetero-barbering

Barbering

- Behavior-associated hair loss
- A complex behavior
- Hair trimming or plucking
- Frequent in laboratory rodents
- Has several distinct contexts
- Has strong genetic component
- Common in some mouse strains
- Sensitive to various manipulations: stress, aggression
- Represents a useful index in behavioral research



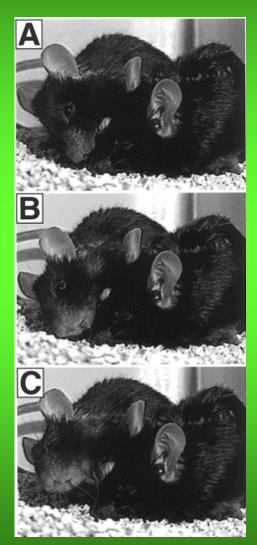
thefunmouse.com

Normal whiskers

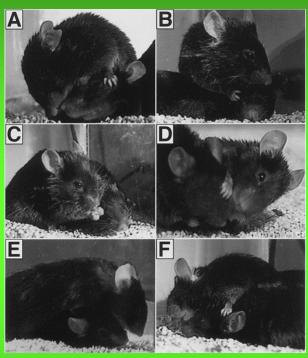
- Whiskers (importance for senses and balance) are chewed off by conspecifics
- Commonly done by the dominant mouse
- Every animal in cage may participate in barbering
- Socially transmitted (can be evoked in non-barbering cages)



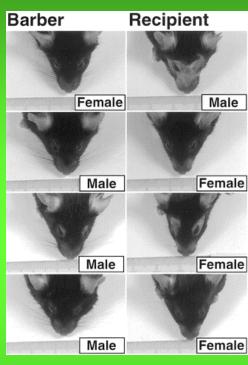
Barbering: How?



A typical barbering act

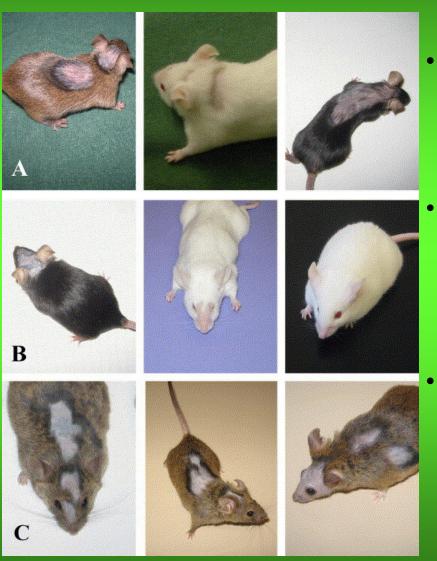


Barbering postures: plucking and trimming



Examples of facial barbering

Mouse strain differences



(A) Sexual barbering by female barbers, left to right: 129S1, NMRI, C57BI/6 males.

(B) Social (dominant)
barbering in same-sex cages,
left to right: C57BL/6 females,
NMRI females;

(C) Barbering in male mice of F1 hybrid strains, left to right: NMRI-129S1, C57BL/6-129SvJ mice

Genetic nature of barbering

Balb/c	F1 Balb/c x C57	C57
Aggressive	Aggressive	Not aggressive
No barbering	No barbering	Active barbering



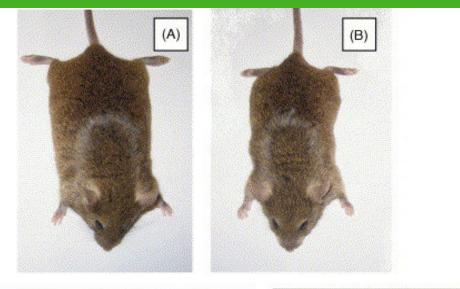


Balb/c "no barbering" genes expressed >> "Barbering" genotype of C57



(Balb/c has stronger genetic effect)

Abnormally low barbering



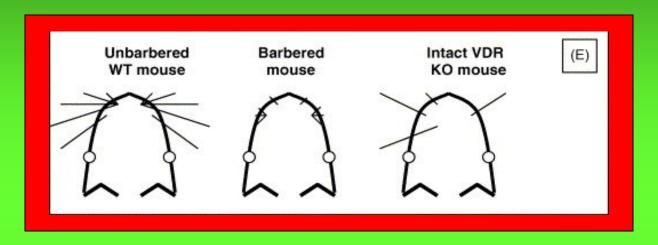


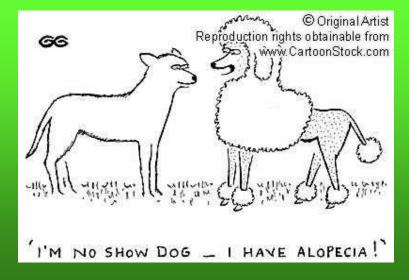


Example: VDR KO mice

- (A) WT 129S1 mouse with normal fur and whiskers
- (B) Barbered WT mouse without whiskers.
- (C) Intact 4-month old VDR KO mouse, with pronounced progressing alopecia
- (D) Nude 1-year old VDR KO mouse, with no fur and whiskers

Alopecia vs. and barbering-evoked whisker loss

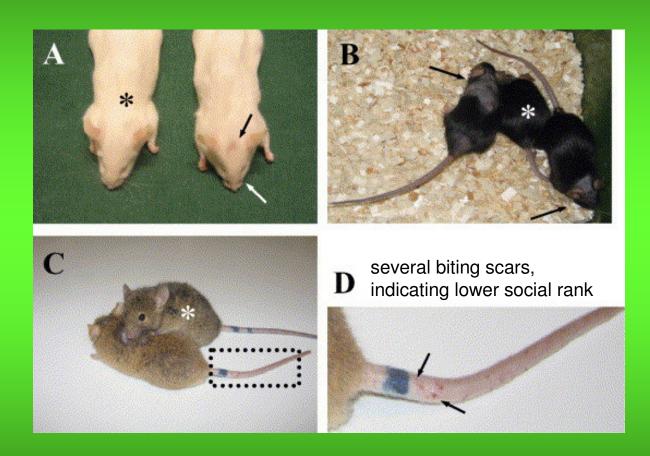




Kalueff et al., 2005

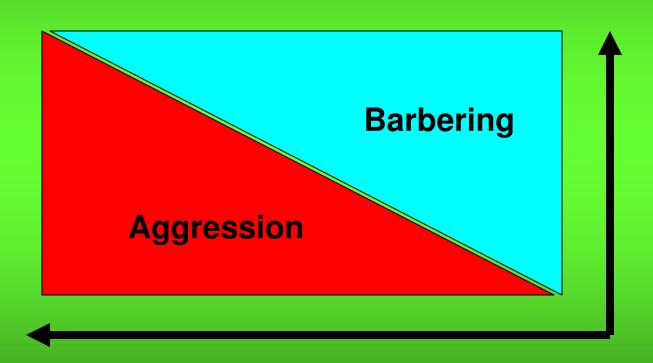
Social/dominant barbering

(* = dominant barber)



(A) NMRI mice. (B) C57Bl/6 mice. (C) Barbering in two F1 C57Bl/6-129S1 mice

Aggression and barbering



Sexual barbering

- Usually performed by females
- Males undergo this barbering willingly
- Some males may also groom females as well

A very human behavior?



Photo; Deluca, 1997



Applicability to human behavior?



Applicability to human behavior?

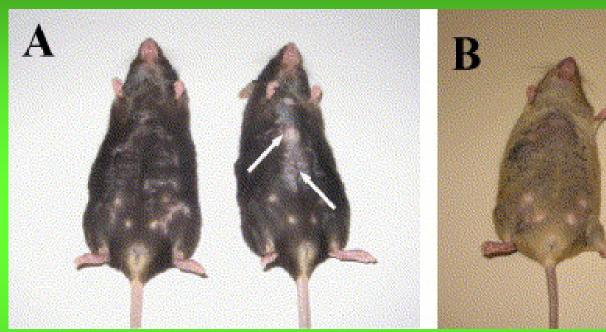








"Maternal" barbering





Produced by suckling pups in lactating mice of C57BL/6 (A) and 129S1 (B) strains, photographed immediately after weaning

Summary

- It is important to assess of grooming and barbering phenotypes in neurobehavioral research in animals
- New techniques and acquisition methods are needed (e.g. 24-hr monitoring systems able to recognize grooming and barbering)
- Applicability for translational research in humans
- Particular relevance for neuropsychiatric disorders (OCD, Trichotillomania, aggression, etc.)

