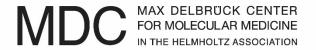


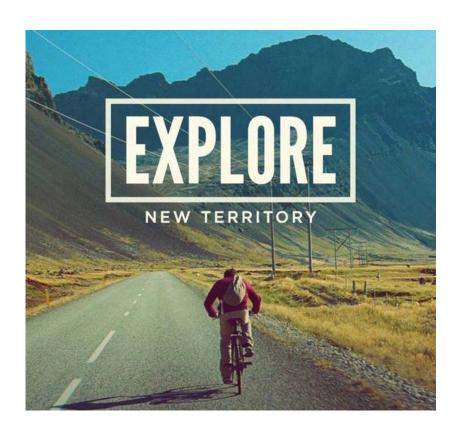
HOW TO ASSESS WELFARE IN GENETICALLY ALTERED ANIMALS?



Anne Zintzsch Animal Welfare Officer





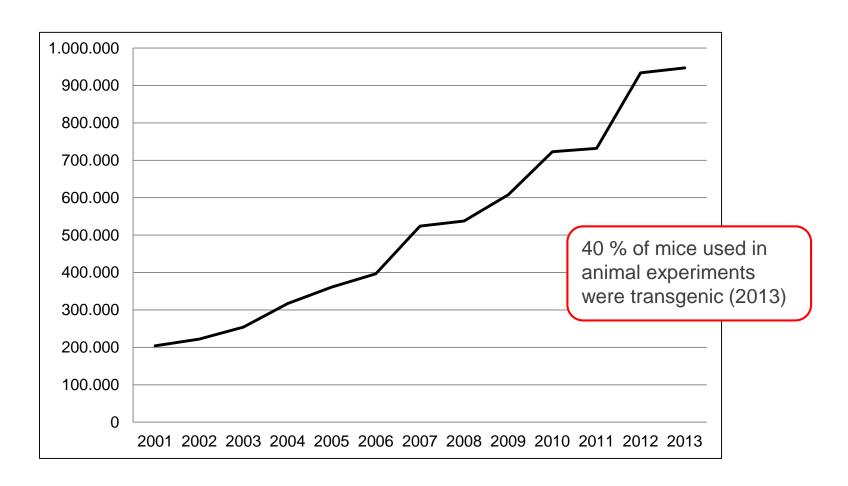


... IN ANIMAL WELFARE LEGISLATION

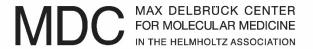




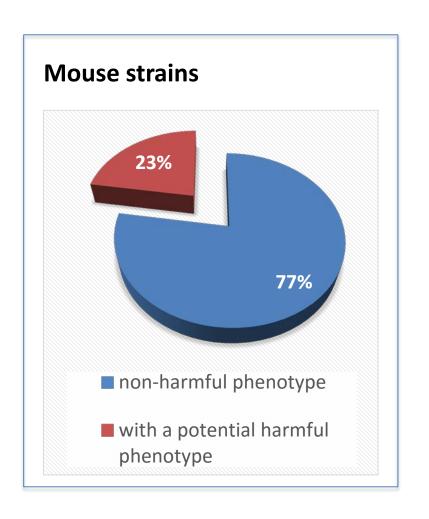
USE OF GENETICALLY ALTERED LAB ANIMALS IN GERMANY

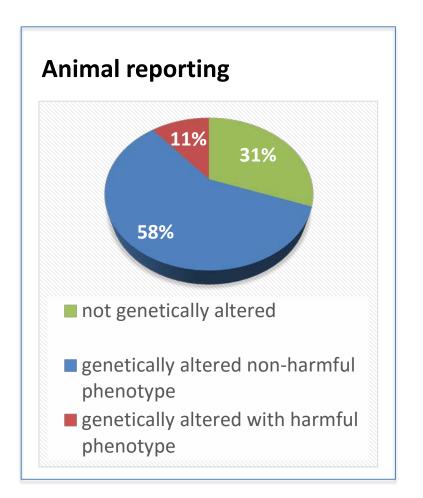




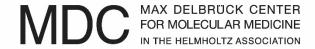


STATUS QUO AT THE MDC

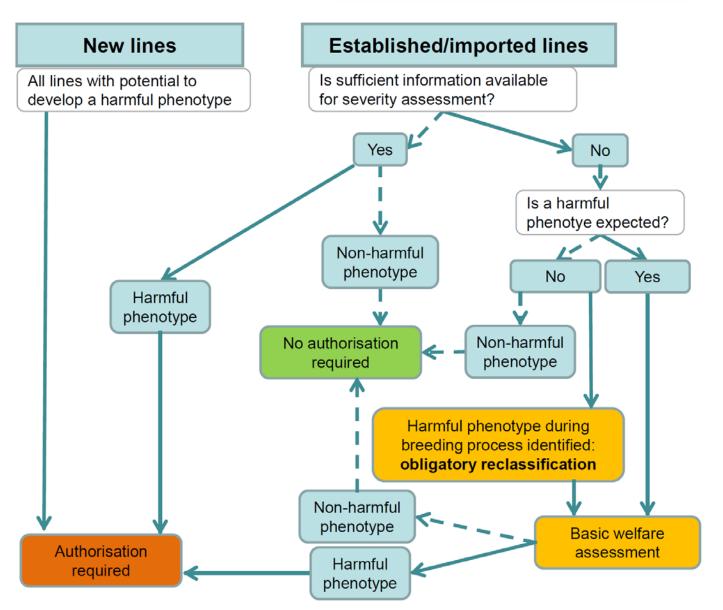




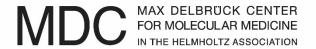




LEGAL
REQUIREMENTS
FOR SEVERITY
ASSESSMENT







RECOMMENDATIONS FOR A BASIC WELFARE ASSESSMENT

- 7 m, 7f
- Neonates, at weaning, adult every two month → line-specific





Form Version 2 of 21 July 2015

"Assessment of newborn litter "*

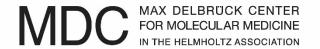
*at the latest during the first cage change

Location (Institute and room):	Husbandry system (e.g. IVC, conventional cage, filter top,
	isolator etc.; hygiene status where applicable):

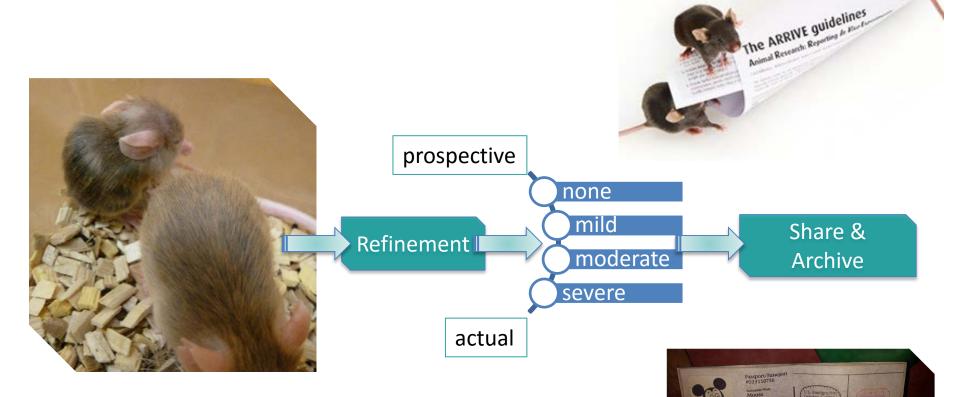
Grune B et al: Animal welfare: Rules for assessing pain in lab animals. Nature 2014; 512: 28.

Bundesinstitut für Risikobewertung: Severity Assessment of genetically altered mice and rats - Version 2: Recommendation no. 002/2016 by the National Committee (TierSchG) 2016.





BASIC WELFARE ASSESSMENT - OBJECTIVES

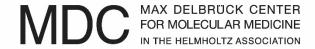




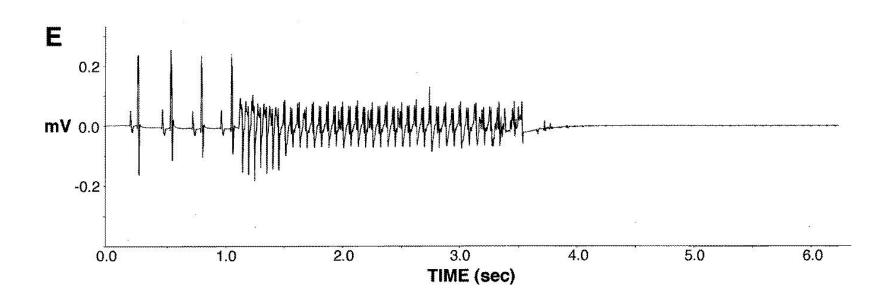


EXAMPLES ON SEVERITY ASSESSMENT AND CLASSIFICATION

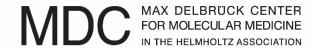




EXAMPLE I: SUDDEN CARDIAC DEATH







Sudden cardiac death - Which severity degree would you assign?

Mentimeter



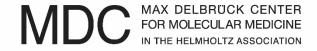




GUIDELINES ON SEVERITY ASSESSMENT AND CLASSIFICATION

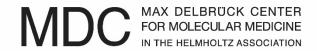
No.	Symptom/disease	Non-harmful phenotype	Mild severity	Moderate severity	Severe severity	Monitoring, Refinement, special housing requirements
7	Cardiovascular and haematolo	ogical diseases				
7.1	Cardiac arrhythmia, e.g. asymptomatic cardiac channelopathies with		Short-term arrhythmia with sudden cardiac			





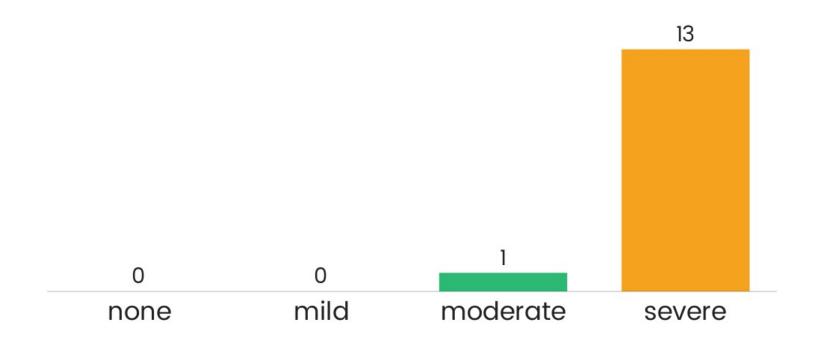
DECOMPENSATED HEART INSUFFICIENCY



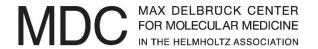


Decompensated heart insufficiency - Which severity degree would you assign?

■ Mentimeter







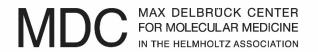
GUIDELINES ON SEVERITY ASSESSMENT AND CLASSIFICATION

No.	Symptom/disease	Non-harmful phenotype	Mild severity	Moderate severity	Severe severity	Monitoring, Refinement, special housing requirements
-----	-----------------	--------------------------	---------------	-------------------	-----------------	---

7 Cardiovascular and haematological diseases

7.4	Dilated or hypertrophic cardiomyopathy	Transient short-term intensified breathing normal ac the home permanen	respiratory distress and impairment general condition, classification dep on expression of symptoms tivity in cage; no	
		impairmer general co	nt of	

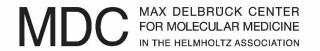




EXAMPLE III: MICROPHTHALMIA, ANOPHTHALMIA





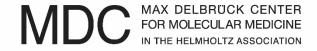


Blindness - Which severity degree would you assign?

Mentimeter







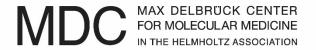
GUIDELINES ON SEVERITY ASSESSMENT AND CLASSIFICATION

No.	Symptom/disease	Non-harmful phenotype	Mild severity	Moderate severity	Severe severity	Monitoring, Refinement, special housing requirements	
4 Di	4 Diseases of the sensory organs ^{IV}						
4.1	Eyes						
4.1.3	Microphthalmia, anophthalmia	Blindness ^V (e.g., small or no eyes) without impairment of normal behaviour				House animals in constant environment	

IV The lack of more than one sense is considered to cause an impairment that should be classified as harmful phenotype

^V If the animals are kept in a constant environment.





EXAMPLE IV: DYSTROPHIC EPIDERMOLYSIS BULLOSA

neonatal









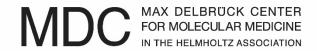






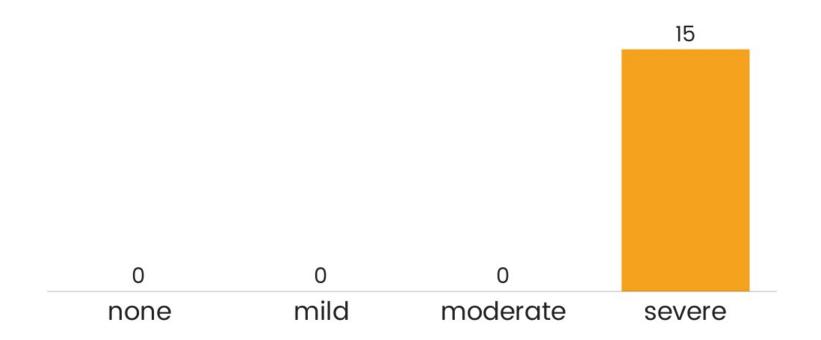
Nyström et al: Rat Model for Dominant Dystrophic Epidermolysis Bullosa: Glycine Substitution Reduces Collagen VII Stability and Shows Gene-Dosage Effect., *PLoS ONE*. 2013;8(5):e64243





Dystrophic epidermolysis bullosa - Which severity degree would you assign?

Mentimeter







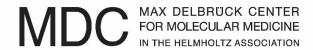
GUIDELINES ON SEVERITY ASSESSMENT AND CLASSIFICATION

No.	Symptom/disease	Non-harmful phenotype	Mild severity	Moderate severity	Severe severity	Monitoring, Refinement, special housing requirements
-----	-----------------	--------------------------	---------------	-------------------	-----------------	---

3 Alterations of the skin and the coat

3.3	Inflammatory skin diseases						
3.4	Dystrophic epidermolysis bullosa				Severe, extensive alterations of the skin (blisters), even limbs may be lost, changes of mucous membranes with compromised food uptake, hyperalgesia ³³		

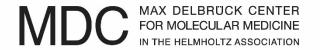




EXAMPLE V: OBESITY







Obesity - Which severity degree would you assign?

Mentimeter







GUIDELINES ON SEVERITY ASSESSMENT AND CLASSIFICATION

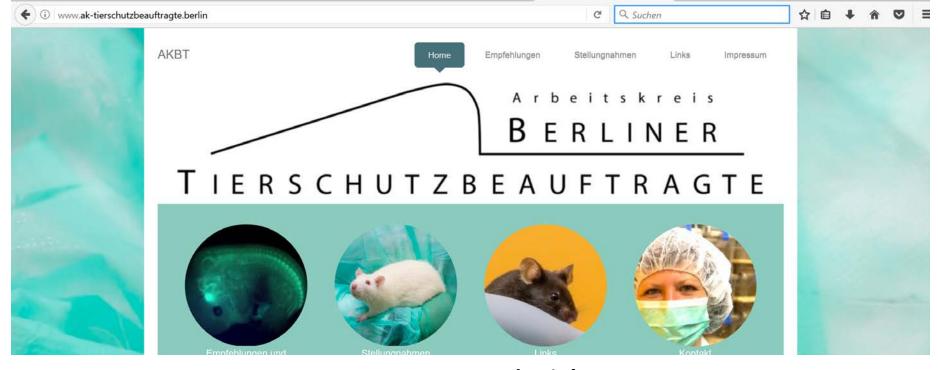
No.	Symptom/disease	Non-harmful phenotype	Mild severity	Moderate severity	Severe severity	Monitoring, Refinement, special housing requirements
10 N	letabolic diseases					
10.3	Obesity ⁵⁷	Bred for obesity without impairment of normal behaviour or general condition	lipid metabolism dis	nents of the metabolic order, elevated levels sification depends on t eneral condition	of blood glucose,	Dietetic food on cage floor, soft bedding when movement is impaired, monitoring the genital health, rat: normal-weight "grooming mate", more frequent change of cages





TAKE HOME MESSAGE

- ➤ Welfare assessment is **line-specific** according to prospective severity assessment and should be adapted to observations
- ➤ Team approach scientist, animal caretaker, AWO/AWB
- ➤ Guidelines on severity assessment and classification as starting point and will be reviewed regularly according to your feedback
- ➤ Transfer and archive severity assessments to support animal welfare and science!



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(now MDC)

HU

Dr. Monika Reißmann

Charité

Dr. Hannah Nickles Dr. Kristina Ullmann

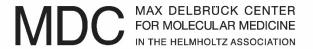
MDC

Dr. Claudia Gösele Dr. Boris Jerchow (now UKE)

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ANY QUESTIONS?



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