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concentration (1% - 2%) ("%" = g/100 mL).

Regular water bottles are removed from the cage and replaced with the DSS-dosed bottles.

DSS is administered in drinking water for 5 days, followed by a return to regular water for 5 days. This cycle is repeated 3 times.

Non-standard water bottles are used for the DSS rats, a special request is required and must be submitted to ACS.

Monitoring:

Ensure that cages are flagged with specific ACS cage alerts that animals are on a DSS study.

Daily monitoring: cages are checked for DSS-water consumption (volume drop in bottle, urine output) and bottles topped up/cages replaced as necessary. Daily weight, appearance, and behavioural changes must be recorded at room level.

In the first two (more severe) DSS cycles: All rats in a DSS-treated group must receive a daily subcutaneous injection of 5ml lactated Ringer's solution. Each and every animal requires a new sterile syringe and needle. Refer to SOP 10.9.1 "Subcutaneous Injection Techniques in Rats". Rats must be weighed before fluid therapy administration.

In the third DSS cycle, when clinical symptoms are expected to be less severe, only rats failing to gain weight over **a**

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Date	New Version
April 27 th , 2018	Updated
February 25 th , 2021	Triennial Update
October 17 th , 2023	Triennial Update and Format Change