



Genetically engineered models (GEMS)

p53 knockout rat

Model	p53 knockout rat
Strain	HsdSage:SD-Tp53 ^{tm1Sage SD}
Location	U.S.
Availability	Cryopreserved

Characteristics/husbandry

- + Monoallelic 11 base pair deletion in Tp53
- + Broad tumor spectrum
- + High degree of tumor malignancy
- + Background Strain: Sprague-Dawley

Zygosity genotype

+ Cryopreserved as heterozygous embryos

Research use

- + Xenografts
- + Vaccine development
- + Cancer
- + Autoimmune disease
- + Infectious disease
- + Hematopoiesis

Origin

The p53 knockout rat model was originally created at SAGE Labs, Inc. in St. Louis, MO and distributed out of the Boyertown, PA facility. The line continues to be maintained through the original SAGE Labs animal inventory acquired by Envigo.

Description

Homozygous null Tp53 rats display onset of tumors at ~4 months of age. A high degree of malignancy is observed across a broad spectrum of tumors. Heterozygous rats have a delayed onset of spontaneous tumors, making them valuable for carcinogenicity screening, as well as studying efficacy of chemopreventive and therapeutic treatment.

p53 is a tumor suppressor protein encoded by the Tp53 gene. Its role in cell cycle regulation and stabilization for preventing genome mutation is observable among a wide variety of multicellular organisms, including humans, rodents, frogs, and fish. Heterozygous rats deficient in p53 protein are prone to spontaneous tumors, making them valuable for in vivo screening of carcinogenicity, as well as studying chemopreventive and therapeutic treatment.

Citations

Dai MS, Hall SJ, Vantangoli Policelli MM, Boekelheide K, Spade DJ. Spontaneous testicular atrophy occurs despite normal spermatogonial proliferation in a Tp53 knockout rat. Andrology. 2017 Nov;5(6):1141-1152.

McCoy A, Besch-Williford CL, Franklin CL, Weinstein EJ, Cui X. Creation and preliminary characterization of a Tp53 knockout rat. Dis Model Mech. 2013 Jan;6(1):269-78.

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Figure 1. Weight and age comparison chart

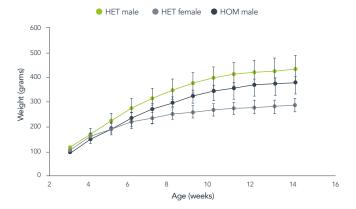


Figure 2. Survival rates of both heterozygous and homozygous p53 null rats (n = 30)

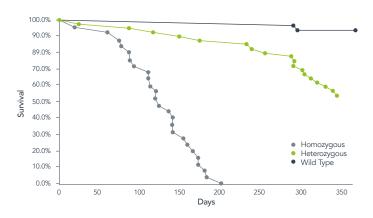


Table 1. Tp53 (+/-) Tumor Spectrum

	Days Survived	Lesions Identified	Metastasis
Sex			
F	21	died and not examined	NA.
M	61	epithelioid hemagiosarcoma – hip	
F	76	lymphoblastic lymphoma of 8-cell origin	multiple organs
F	78	glioblastoma – brain	
M	87	malignant meningioma (meningeal sarcoma) - brain	
M	88	pulmonary carcinoma – lung	kidney
M	93	epithelioid hemangiosarcoma - face	-
M	110	lymphoblastic lymphoma of 8-cell origin	multiple organs
F	112	epithelioid hemangiosarcoma – back	
M	114	rhabdomyosarcoma – thoracic wall	
F	120	diffuse astrocytoma – brain	
M	120	pleomorphic astrocytoma – brain	
M	124	diffuse astrocytoma – brain	
M	136	malignant meningioma (meningeal sarcoma) – brain	-
		osteosarcoma – scapula	lung
M	141	lymphoblastic lymphoma of 8-cell origin	multiple organs
		rhabdomyosarcoma – back	
M	141	osteosarcoma – spine	
F	142	mammary adenocarcinoma - mammary gland	
		epithelioid hemangiosarcoma – pancreas	
F	154	lymphoblastic lymphoma of 8-cell origin	multiple organs
		mammary carcinoma – mammary gland	
		Zymbal's gland carcinoma – ear canal	
M	160	malignant meningioma (meningiosarcoma) - brain	
		osteosarcoma – pelvis	lung
F	166	pleomorphic astrocytoma – brain	
M	172	osteosarcoma – tibia	
M	173	malignant meningioma (meningiosarcoma) – brain	
M	181	osteosarcoma – skull	
F	183	mammary adenocarcinoma – mammary gland	
		malignant meningioma (meningiosarcoma) – brain	
		osteosarcoma – spine pleomorphic sarcoma – back muscle	
	F M F F M M M M M M M F M M M F M M M M	Sex Survived F 21 M 61 F 76 F 78 M 87 M 88 M 93 M 112 M 114 F 120 M 124 M 136 M 141 F 142 F 154 M 160 F 166 M 173 M 181	Sex Survived Lesions identified F 21 died and not examined M 61 epitheloid hemagiosarcoma – hip F 76 lymphoblasts (lymphoma of 8-cell origin F 78 glioblastoma – brain M 87 maligrant meningiona dimeringeal sarcomal – brain M 88 pulmonary carcinoma – lung M 93 epitheloid hemangiosarcoma – face M 110 lymphoblasts (lymphoma of 8-cell origin F 112 epitheloid hemangiosarcoma – btain M 120 diffuse astrocytoma – btain M 124 diffuse astrocytoma – btain M 124 diffuse astrocytoma – btain M 136 maligrant meningioma (meningeal sarcoma) – brain M 141 symphoblasts (lymphoma of 8-cell origin rhadomyosarcoma – back marmany adenocarcinoma – mammary gland F 142 mammary adenocarcinoma – mammary gland F 143 mammary adenocarcinoma – mammary gland Zymbal's gland carcinoma – ear canal

Table 2. Tp53 (-/-) Tumor Spectrum

ID	Sex	Days Survived	Lesions Identified	Metastasis
Het 1	M	25	epithelioid hemangiosarcoma – limb	
Het 2	M	87	malignant mengioma (meningeal sarcoma) – brain	
Het 3	F	117	no cancer detected	
Het 4 M	M	150	epithelioid hemangiosarcoma – thoracic wall	
			diffuse astrocytoma – brain	
Het 5	M	175	liposarcoma – abdomen	lung
Het 6	F	233	pleomorphic astrocytoma – brain	
Het 7	M	239	malignant mengioma (meningeal sarcoma) – brain	
Het 8	M	255	no cancer detected	
Het 9	M	289	olfactory neuroblastoma – brain	
Het 10	M	291	squamous cell carcinoma – hip	
Het 11	M	291	osteosarcoma – spine	
Het 12	F	302	olfactory neuroblastoma – brain	
Het 13	M	304	no cancer detected	
Het 14	M	312	diffuse astrocytoma – brain	
Het 15	F	319	olfactory neuroblastoma – brain	
Het 16	M	330	myxosarcoma – thoracic wall	
Het 17	F	341	sgamous cell carcinoma – dorsal skin	
			adrenal cortical adenoma – adrenal gland	
			complex pheochromocytoma – adrenal gland	
Het 18	M	344	sgamous cell carcinoma – hip	
			Zymbal's gland carcinoma – ear canal	
Het 19	F	375	mammary fibroadenoma - mammary gland	