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## Review Paper

# The Role of Laboratory Animals in Pharmacological Testing and Drug Development

**Dr. Pasupulati Haritha\*, Jarupla Mahika, Ambala Syama, Mogilicherla Archana,  
Dr. Ramadevi Pemmereddy**

*Bharath School of Pharmacy, Hyderabad, Telangana- 501510, India.*

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## ABSTRACT

Animal research has long contributed to advances in medicine and drug development. By observing how diseases progress and how medicines affect living organisms, researchers gain valuable insights into how similar treatments might respond in humans. These studies also help scientists understand biological processes and identify possible safety risks before new drugs move on to clinical testing. Over the years, animal research has played an important role in improving medical treatments, including the development of vaccines and other life-saving therapies. Different animals are chosen for studies depending on how closely they relate to humans, how manageable they are in research settings, how consistent the results tend to be, and whether their use meets accepted ethical standards. Along with describing the roles of different laboratory animals used in experimental pharmacology, this work highlights their importance in drug discovery, disease studies, and broader biomedical research.

## INTRODUCTION

After a therapeutic molecule is discovered, it is crucial to study how animals behave and react to it for human use. The choice of an appropriate animal model is based on physiological, anatomical, and behavioural similarities to humans in order to guarantee the applicability and transferability of the experimental findings. One of

the initial stages in the study and creation of novel drugs is the use of animal models to comprehend how a disease manifests in the body. Any medicinal agent must pass a rigorous preclinical evaluation process, which involves the use of animal models, before it is deemed safe and effective for use in humans. Thus, pharmacokinetics (absorption, distribution, metabolism, and excretion), pharmacodynamics

**\*Corresponding Author:** Dr. Ramadevi Pemmereddy

**Address:** Assistant Professor, Department of Pharmacology, Bharath Institute of Pharmacy, Hyderabad, Telangana-501510, India.

**Email** ✉: [ramadevipemmireddy002@gmail.com](mailto:ramadevipemmireddy002@gmail.com)

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(mechanism of action), and possible toxicological effects that a drug might produce can all be observed in a whole-system setting that animal models offer.

## DISCUSSION

A wide variety of animals are commonly used in pharmacological studies, selected based on their physiological relevance to human systems.

### (1) HAMSTER

*Mesocricetus auratus* is also called as Syrian or golden hamster. In the past 5 yrs hamsters have been increasingly useful in pharmacological and toxicological investigations. The extensive use of hamsters in research can be attributed to their fast reproductive cycle and relatively large litter size, which average around six pups per litter.

**TABLE 1: EXPERIMENTAL USE OF HAMSTERS**

| Area of the study  | Description                                                                                                                                                                                                                                                                                                                                                              | References |
|--------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------|
| Teratogen studies  | Short gestation period of Hamsters allows studies on effects of teratogens and pregnancy drugs. Hamsters are twice more sensitive to strychnine than mice.                                                                                                                                                                                                               | [1,2]      |
| Blood Pressure     | Hamsters show similar BP response to acetylcholine and epinephrine as other mammals; used to study pharmacological effects on BP.                                                                                                                                                                                                                                        | [3]        |
| Amyloidosis        | Syrian hamsters have proven to be a valuable animal model for the study of amyloidosis due to the similarity in disease progression and response to stimuli when compared to humans. The induction of amyloid deposition through subcutaneous injections of casein and lipopolysaccharide (LPS) effectively mimics systemic inflammation and promotes amyloid formation. | [4]        |
| Other applications | Hamsters are used to study different drugs used in human disease conditions such as muscle dystrophy, epilepsy, atrial thrombosis, carcinogenesis, type 2 diabetes mellitus.                                                                                                                                                                                             | [5,6]      |

### (2) MARMOSSET

Marmoset (*Callithrix jacchus*) are numerous species of long-tailed American monkeys, similar in appearance as squirrels. The common

marmosets live in the scrub forest of northeastern Brazil and weighing 400 grams with 15-30 cm height. Their average life span is about 5 to 7 years and a maximum life span of 16.5 years.

**TABLE 2: EXPERIMENTAL USE OF MARMOSETS**

| Area of the study       | Description                                                                                                                                           | References |
|-------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------|------------|
| Pharmacodynamic Studies | Marmosets are widely used in pharmacodynamic research, especially in neuroscience, immunology, and infectious diseases.                               | [7]        |
| Neuroscience            | Used as models for Parkinson's disease and Alzheimer's disease, which primarily affect humans and also aid in studying pharmacokinetics of biologics. | [8]        |
| Infectious Diseases     | Marmosets are used as models to study bacterial and viral infections due to their ability to closely mimic human disease progression.                 | [9]        |



|                             |                                                                                                                                                                                     |      |
|-----------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|
| Psychiatric Diseases        | Used as experimental models for understanding symptoms and mechanisms related to anxiety, panic, stress, and obsessive-compulsive disorder (OCD).                                   | [10] |
| Antibiotics & Drug Research | The effects of rapamycin have been studied in marmosets, where oral administration of the drug has been shown to enhance both lifespan and health span in these non-human primates. | [11] |

### (3) OWL

Owls (*Strigiformes*) are birds which are found in all regions, except in polar ice caps and some remote islands. Owls are utilized to investigate the pharmacokinetic properties of Meloxicam, a pain-relieving drug, with studies showing an increase in plasma concentration following oral administration. In general, owls and other bird species are commonly used to examine the pharmacokinetics of analgesic medications [12].

*Drosophila melanogaster* also known as ‘fruit-fly’ or ‘lesser fruit fly’ and less commonly called as ‘vinegar fly’ or ‘banana fly’ or ‘pomace fly’. Their development time differs with the external temperatures. Under optimum temperatures at 25°C, their life span is about 50 days. The shortest development time is 7 days, which is at 28°C. Development time increases at higher temperatures due to heat & stress. Heart rate is of 4-6 beats/min and their body weight is about 1-1.5 mg.

### (4) DROSOPHILA MELANOGASTER

**TABLE 3: EXPERIMENTAL USE OF DROSOPHILA MELANOGASTER**

| Area of the study                         | Description                                                                                                                                                                                                                                                                                                                                                                                                                        | References |
|-------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------|
| Genetics                                  | Used in developmental biology, cell biology, neurobiology, and genetics. First used by Thomas Hunt Morgan in 1910. Helped discover sex-linked inheritance, multiple alleles, and gene mapping.                                                                                                                                                                                                                                     | [13]       |
| Age-related Studies                       | Suitable model for testing anti-aging drugs and studying age-related diseases like Parkinson's, Alzheimer's, Huntington's, cardiovascular disease, muscular dystrophy, and metabolic disorders.                                                                                                                                                                                                                                    | [14,15]    |
| Anti-seizure Medications                  | Fischer et al used bang-sensitive mutants (e.g., parabss1) to test seven anti-seizure drugs. Mutants show seizures after mechanical stimulation, making them ideal for screening seizure treatments.                                                                                                                                                                                                                               | [16]       |
| Toxicity studies and Carcinogenic studies | <i>Drosophila</i> has proven to be an effective <i>in vivo</i> model for investigating the mechanisms of action, toxicity, and bioavailability of drugs used in the treatment of colorectal, thyroid, brain, and various other cancers.                                                                                                                                                                                            | [17,18]    |
| CNS Studies                               | <i>Drosophila</i> are widely used in Central Nervous System (CNS) studies due to their genetic tractability, conserved genes with humans, rapid life cycle, and comprehensive genetic resources. The fruit fly's small yet informative CNS, coupled with advanced genetic tools like the Gal4/UAS system, allows for detailed analysis of neuron function, glial cell interactions, and the implementation of neural computations. | [19]       |

### (5) CHAENOCEPHALUS ACERATUS

*Chaenocephalus aceratus* is commonly known as the blackfin icefish or Scotia Sea fish, is a species



of crocodile icefish. Because it lacks red blood cells and haemoglobin, it is frequently referred to as the "white-blooded fish." It grows best in Antarctic waters that are close to the freezing point of saltwater (about -2°C).

Blackfin ice fish is used as model for examining the drugs like Sodium EDTA, Verapamil and atropine. These drugs shown the inhibition of sustain contraction and increases calcium supply and caused progressive relaxation. Aortic blood flow and cardiac output was studied in this species. The ventral aortic blood flow, measured by electromagnetic flow meter, increased during hypoxia and decreased during the hyperoxia and the cardiac output was well regulated in these species [20].

Zebra fish gives vertebrates and has a backbone like humans and have close relation to humans than commonly used invertebrates' models, such as insects and worms. Due to its clear eggs which can be developed outside the body, allow watching a zebra fish egg grow into a newly formed fish under a microscope. Lifespan of zebra fish is considered to be approximately 5 years and length of the adult fish is about 6 cm. When compared to other animals for drug testing like rats, mice, frog, monkeys, zebrafish is most commonly and easily found and can develop the larvae in-vivo within 24 hrs, and cost effective, easy to handle, shares higher similarities in brain, heart, genes with humans. These benefits made the use of zebrafish as a model made easier.

## (6) ZEBRAFISH

**TABLE 4: EXPERIMENTAL USE OF ZEBRA FISH**

| Area of the study     | Description                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                | References |
|-----------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------|
| Genetics              | Zebrafishes are used as a model for biomedical studies, especially in genomics and developmental biology. Free-swimming larvae are ideal for studying hearing/balance disorders (ototoxicity), locomotion, and social learning.                                                                                                                                                                                                                                                                            | [21]       |
| Neuroscience          | Due to structural similarity to the human brain but with less complexity, zebrafish are used to study brain physiology, anatomy, and behavior, including cognitive and behavioral research.                                                                                                                                                                                                                                                                                                                | [22]       |
| Cancer studies        | Doxorubicin is an antitumor agent belonging to the class of neoplastic agents and is commonly used in the treatment of cancers such as lymphoma, leukemia, and breast cancer. Although therapeutic doses are approximately 500 mg/m <sup>2</sup> of body surface area and have been reported to benefit around 3–4% of patients, its clinical application is limited by dose-dependent cardiotoxicity. Zebrafish has emerged as a valuable model organism for studying doxorubicin-induced cardiotoxicity. | [23]       |
| Antipsychotic Studies | Zebrafish serve as an alternative to rats for evaluating cardiotoxic effects of first-generation antipsychotic drugs (e.g., aripiprazole, risperidone), overcoming limitations of rodent models.                                                                                                                                                                                                                                                                                                           | [24]       |
| CNS Studies           | Ideal for CNS research due to transparent embryos, genetic similarity to humans, and real-time neural imaging. Used to study epilepsy, autism, Parkinson's disease, and spinal cord injuries.                                                                                                                                                                                                                                                                                                              | [25]       |

## (7) AFRICAN GREEN MONKEY

The African green monkey is a vervet monkey native to sub-Saharan Africa. African green



monkey, has emerged as a promising alternative model in pharmacological research, offering several advantages over other commonly used primates. They are medium-sized primates, with males being slightly larger than females. Males typically weigh between 4.5 and 7.5 kg (10 to 17 lb), while females weigh between 4 and 6 kg (9-13 lb). Normal body temperature is 36.0°C – 39.5°C (96.8°F – 103.1°F) and heart rate around 74 beats per min. They exhibit sexual dimorphism, with males generally being larger and heavier than females. African green monkey, has emerged as a promising alternative model in pharmacological research, offering several advantages over other commonly used primates. Areas of research: Pharmacokinetics, Antimicrobial efficacy, Ocular pharmacokinetics, Neurological disorders. African green monkey has a lifespan of about 15 years in captivity and 10-13 years in the wild. They reach sexual maturity at about 4-5 years of age.

### **EXPERIMENTAL USES OF AFRICAN GREEN MONKEY:**

#### **Research on infectious diseases and Vaccine development:**

Along with explaining the roles of different laboratory animals used in experimental pharmacology, this work shows the animals remain important in drug research, disease studies, and biomedical science. [26,27].

#### **Neurological research:**

These animals are often used in research on Parkinson's disease and other disorders of the nervous system. When African green monkeys and other non-human primates are involved in studies, researchers follow strict rules to ensure the animals are treated humanely and kept safe.

### **(8) GOAT**

Goats have distinctive features including rectangular pupils for wide-angle vision, four chambered stomach for digestion. Goats are mostly diurnal animals. Goats normal body temperature falls between 101.3°F-103.5°F (38.5°C- 39.7°C). Respiration rate is 10-30 breaths per min and heart rate is 70-90 beats per min. The life expectancy of goats is generally between 15 and 18 years. Goats have been used in experimental for a variety of purposes, such as drug metabolism and absorption studies, testing drug efficacy and safety and nutritional studies.

### **EXPERIMENTAL USE OF GOAT**

Goats are employed in many experimental contexts, most notably as animal models in agricultural and biomedical research. They act as models for researching human illnesses, creating novel treatments, and comprehending the behaviour of animals.

#### **Research in Biomedicine:**

**Models of animals:** Human diseases such as paratuberculosis, tuberculosis, and other infectious diseases are studied using goats as animal models [28,29]

**Stem Cell Research:** They are employed in research on stem cells for regenerative medicine and tissue engineering, especially in the musculoskeletal system [30].

**Transgenic Goats:** Genetically modified goats can be used as bioreactors to produce valuable proteins in their milk. Recombinant proteins are produced in transgenic goats for medical applications [31].

**Reproductive Biotechnology:** Reproduction biotechniques, such as embryo cryopreservation, sperm sexing, and genome editing, are studied using goats to improve livestock production [32]





**Behavioral Studies:** Goats are used in studies to understand their social behaviour, communication patterns, and cognitive abilities. Goats can be used in therapy programs to improve the well-being of individuals in hospitals, nursing homes, and other settings [33].

## (9) GERBIL

Mongolian gerbil is a small laboratory rodent. Gerbils are phenotypically related to rat and also known as “jirds” or “sand rat”. They are preferred in laboratory because of easy in handling, mild and quiet nature. The average lifespan of a Gerbil is 3 to 5 years, though some gerbils can live up to 8 years or even longer. Normal body temperature of gerbils is 98°F to 102°F (37°C -39°C), respiration rate is 70-120 bpm with heart rate 250-500 per min.

### EXPERIMENTAL USE OF GERBIL

Gerbils are used in experimental studies due to their genetic predisposition to convulsions and their suitability.

Due to their propensity for convulsions and their suitability as models for conditions like otitis media and epilepsy, gerbils are frequently employed in experimental research. Additionally, they are useful in research on neurological conditions, brain development, and the functioning of hearing and vision.

## Epilepsy

Gerbils, particularly some strains, exhibit a genetically determined propensity for seizures, making them useful for studying epilepsy patterns and testing antiepileptic drugs [34].

## Parasitic Diseases and Viral infections

Gerbils are used to study parasitic diseases like schistosomiasis and filariasis, allowing researchers to understand the infection process and develop treatments. Gerbils can be infected with viruses like Rift Valley fever virus to study encephalitis and other related diseases [35,36]

## Behavioural Studies

Gerbils can be used in behavioural studies, such as exploring their responses to environmental changes or their social interactions. Gerbils are used in research on how they process auditory and visual information, which can provide insights into the development and function of sensory systems [37].

## (10) Rats

The Laboratory rats have widely been used for studying various diseases in the field of neurology, metabolic diseases, cardiovascular diseases, cancer biology, toxicology, reproductive biology etc.

**TABLE 5: EXPERIMENTAL USES OF RATS**

| Area of the study    | Description                                                                                                                                                                                                                                                                                                                                                                                           | References |
|----------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------|
| Genetics             | A large number of inbred strains that exhibit a variety of phenotypes and serve as numerous models of human characteristics and illness have been identified. There is now an array of disease models available because more than 350 rat genes have been found to be involved in the underlying causes of diseases or to be crucial in important biological processes that are modified in diseases. | [38]       |
| Neurological studies | Rats are used for neuroscience studies including Parkinsons disease, stress disorders, Schizophrenia, anxiety, depression, epilepsy. Various evaluation disease models such as open field test, swimming                                                                                                                                                                                              | [39]       |



|                        |                                                                                                                                                                                                                                                                                                                                                                              |         |
|------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------|
|                        | behavior, tail suspension test, elevated plus maze test can be best studies in rats.                                                                                                                                                                                                                                                                                         |         |
| Cardiovascular studies | Rats are widely used to study cardiovascular diseases such as myocardial infarction, hypertension, coronary artery disease, congestive heart failure. Larger size compared with mice, physiological similarity with the human heart makes the rats suitable for conducting surgical procedures like coronary ligation method, ECG analysis, chemical induced disease models. | [40,41] |
| Metabolic studies      | Genetically engineered rat models like Zucker rats, Goto-kakizaki rats are used for studying obesity and diabetes. These models are helpful in studying pathophysiology of pancreatic diseases, obesity, diabetes, insulin resistance and to test various drugs against these diseases.                                                                                      | [42]    |
| Toxicological research | Various strains of rats have been used in toxicological studies such as acute, sub-acute and chronic toxicological studies. The metabolic pathways of these animals are similar to humans, thus making these animals suitable for studying dose response relationships, pharmacokinetic and toxicokinetic studies.                                                           | [43,44] |
| Oncology               | Rat carcinogenicity studies are an essential component of determining the possible human cancer risk, especially for chemicals that might be present in the environment or in novel medications. A number of transgenic rat models have been developed to aid in the study of in vivo mutagenesis and carcinogenesis.                                                        | [45]    |

**(11) MICE**

Mice are extensively used in research due to their small size, short reproductive cycles, biological similarities with the humans.

**TABLE 6: EXPERIMENTAL USE OF MICE**

| Area of the study      | Description                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                | References |
|------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------|
| Genetics               | Mice are among the most genetically tractable animal models. Gene editing tools such as CRISPR/Cas9, as well as a huge number of transgenic, knockout, and humanised mice models, make them excellent for studying human genetic illnesses.                                                                                                                                                                                                                                                                | [46]       |
| Immunology             | Mice have played a crucial role in immunology research, including vaccine development, autoimmune diseases, transplant rejection, and cancer immunotherapy due to the widespread availability of immunodeficient and humanised strains.                                                                                                                                                                                                                                                                    | [47]       |
| Cancer Studies         | Mice serve as an excellent animal for developing cancer models, genetically engineered cancer models, xenograft models, spontaneously induced models are widely used.                                                                                                                                                                                                                                                                                                                                      | [48]       |
| Neurological Disorders | Mice are used to study various neurodegenerative diseases like Alzheimer's, Parkinson's disease, autism etc. In addition to genetic models, chemical and lesion-based models are utilised to simulate neurological illnesses by creating specific brain damage or neurochemical imbalances. Furthermore, mouse models offer an excellent platform for testing possible neuroprotective and medical therapies, including as small-molecule medicines, gene therapy methods, and stem cell-based treatments. | [49,50]    |



|                               |                                                                                                                                                                                                                                         |      |
|-------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|
| Metabolism & Diabetes         | Ob/ob and db/db mouse models are useful for researching obesity, diabetes, and metabolic syndrome. Their small size and rapid reproduction make them excellent for long-term nutritional studies.                                       | [51] |
| Cardiovascular Studies        | Transgenic mice are used to explore genetic contributions to atherosclerosis, hypertension, and heart failure. ApoE <sup>-/-</sup> and LDLR <sup>-/-</sup> mice are popular models for studying lipid metabolism and vascular function. | [52] |
| Infectious Disease & Vaccines | Mice are used to study the pathogenesis of viral, bacterial, and parasitic infections. Their immune responses are well-characterized, and many vaccine candidates are first tested in murine models.                                    | [53] |

## (12) FERRET

Ferrets are native to North America; they live in prairie dog burrows. These animals are used as a model to study the illness, severity and effects in lower & upper respiratory tracts caused by influenza [54]. During the covid period, several vaccines were introduced, and the specified vaccine is used to treat coronavirus i.e., SARS-CoV, where nearly 10 vaccines were approved by WHO, but after the administration of the vaccine, many have developed SARS (severe acute respiratory syndrome). The study conducted by Gough et al confirms that ferrets are a suitable model for studying mild or asymptomatic SARS-CoV-2 infection, reflecting many aspects of early-stage or subclinical human COVID-19. Ferrets are especially useful for investigating viral transmission, immune response, and testing of intranasal vaccines or antivirals [55].

## (13) SEA HORSE

Common names of sea horse are estuary seahorse, yellow seahorse, spotted seahorse. Seahorses are uniquely used in experimental research for studying male pregnancy and reproduction. Male seahorses possess a brood pouch, functionally similar to a mammalian uterus, which supports fetal development. This pouch contains beneficial microorganisms such as *Marinomonas*, *Holomonas*, and *Aeribacillus* that aid in fetal

nutrition and immune support. Ongoing research focuses on understanding the development, function, and immunological role of the brood pouch in male pregnancy [56].

## (14) CHICKEN

Chickens are commonly employed in biomedical research because of their vulnerability to different viruses, making them useful for studying viral infections, immunological responses, and vaccine development [57]. Their retinas are used as models in neuroscience to investigate visual system development and neuronal plasticity [58]. The embryos of chicken are used in studying the growth of tumor cells, metastasis, and angiogenesis, and also to test new cancer drugs and therapies [59]. Chicken's are also used as experimental model for studying atherosclerosis due to their similar lipoprotein levels to humans. Chicken embryos are useful in developing models of heart conditions in atrial septal defect, or hole in the heart [60]. The first cancer-causing virus, Rous sarcoma virus, was identified in chickens. The isolated chicken intestine (often the duodenum or ileum) is a classic *in vitro* model used in pharmacology and physiology to study the effects of cholinergic and anticholinergic agents on smooth muscle contraction [61].

## (15) MEDAKA





Medaka is also known as Japanese rice fish or *Oryzias latipes* or killfish. These are small (1.4—3.6 cm) and are native to East Asia (Japan). They are euryhaline which are present in both brackish and freshwater. They can survive at wide range of temperature like 0°C or till 42°C.

## EXPERIMENTAL USE OF MEDAKA

### Toxicology & Ecotoxicology

Medaka is used to evaluate or check the effects of chemicals, pollutants, and other environmental stressors on aquatic life [62].

### Space Biology

Medaka is also been used in space missions to study the effects of microgravity on their development and reproduction [63].

### Cancer Research

Medaka is used in carcinogenesis studies to understand the development of tumors and the effects of various substances on tumor formation [64].

### Nanotoxicology

Medaka is used to study the toxicity of nanomaterials, such as graphene oxide.

### Cardiovascular studies

Medaka used for the ECG analysis and these organisms are also used for the testing the effect of Verapamil, a calcium channel blocker drug. Adult Medaka used to evaluate the acute toxicity of drugs like Diclofenac, Triclosan, Carbamazepine in a semi-static water and we observed there is no toxicity. [66]

## (16) XENOPUS LAEVIS

*Xenopus laevis* is an amphibian and also called as clawed frog. It is a genus of highly aquatic frog native from Sub-Saharan Africa. *Xenopus laevis* is an inactive creature and is a dimorphic organism. Their life span is about 15 yrs and with the body weight of 60gms. These species are entirely aquatic and are usually found in lakes, Ponds, Sea, rivers and also in man-made reservoirs. They can survive without food even for a year.

TABLE 7: EXPERIMENTAL USE OF ZEBRA FISH

| Area of the study                       | Description                                                                                                                                                                                                                                                  | References |
|-----------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------|
| Embryonic Development and Organogenesis | <i>Xenopus</i> embryos are widely used to study fundamental processes such as gastrulation, neurulation, and organogenesis, including the development of the heart and nervous system                                                                        | [67]       |
| Cell and Molecular Biology              | Oocytes of <i>Xenopus</i> serve as a powerful tool for the heterologous expression of membrane proteins, ion channels, and G-protein-coupled receptors, enabling functional characterization of these molecules                                              | [68]       |
| Human Disease Modeling                  | <i>Xenopus</i> represents a powerful and versatile model system for investigating the mechanisms underlying congenital heart diseases, left-right patterning defects, and other developmental abnormalities                                                  | [69]       |
| Regeneration Research                   | <i>Xenopus</i> serves as a key model for studying tissue and organ regeneration, particularly in the spinal cord, brain, and limbs                                                                                                                           | [70]       |
| Cancer Studies                          | Allegrucci C et al studied the effects of various anticancer studies on the embryonic development of <i>Xenopus laevis</i> . All the anticancer drugs except cisplatin have shown malformations like abnormal edema, head, eye and abnormal heart at highest | [71]       |



|                                |                                                                                                                                                                                                        |      |
|--------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|
|                                | concentration. This indicates anticancer drug may affect embryogenesis.                                                                                                                                |      |
| Epilepsy Research              | Valuable for studying seizures, early brain development, and GABAergic signaling; supports high-throughput and developmental neuroscience research.                                                    | [72] |
| Hepatotoxicity Studies         | Paracetamol-induced liver injury was examined in <i>Xenopus</i> embryos, demonstrating effects comparable to those observed in humans experiencing hepatotoxicity due to paracetamol overdose.         | [73] |
| Genetic and Molecular Research | <i>Xenopus</i> oocytes are widely utilised to investigate the biological and molecular mechanisms of genetic material, enhancing the comprehension of the pathophysiology of numerous human disorders. | [74] |

### (17) PIGEON

Pigeons are birds belonging to family Columbidae, which also includes Doves. Pigeons are most frequently used in the experimental research, particularly in Psychology and behavioural science, to study various cognitive studies including learning, discrimination, and categorization. They are also used as animal model in research studies like study aspects of vision and social communication. Additionally, pigeons are also used in operant conditioning experiments, demonstrating their ability to learn through reinforcement and punishment. In recent studies, by using animated digital models of pigeons it is concluded that pigeons can categorize different locomotive animal gaits and types of complex human behaviour [75]. Pigeons are also used in microbiological and genetic research, especially to look into the genetic factors that make *Escherichia coli* resistant to antibiotics. This research is important to understand the underlying mechanisms of resistance to antibiotics [76].

### (18) LLAMAS

Llamas are domesticated South American camelids. A healthy llamas have heart rate ranging between 60-90 beats per minute and Crias (baby llamas) heart rate ranging from 80-120 bpm. Normal respiration rate of llamas at rest is between

10-30 breaths per min. Llamas are induced ovulators, meaning ovulation occurs after mating, typically from 24-36 hours later, which allows for breeding at any season of year. Females reach puberty around 12 months, while males often don't become reproductively functional until 3 years old. Llamas are increasingly being used in medical research, mainly in virology, due to their unique characteristics of antibodies and the ability to produce "Nano bodies" that can be engineered into therapies.

### EXPERIENTIAL USE OF LLAMAS

The use of llamas in biomedical research has increased recently, especially in virus-related studies. This is due to the fact that they create unique antibodies called nanobodies, which are more stable and smaller than ordinary antibodies. They are helpful for creating novel therapies and diagnostic instruments for infectious diseases and other medical conditions because of these characteristics.

#### HIV Research:

Antibodies produced by llamas have been shown to work against different strains of HIV, suggesting new possibilities for vaccine research and targeted therapies. Due to their strength and accuracy, llama-derived nanobodies are also increasingly being used in diagnostic testing [78].



### **COVID-19 Research and infectious diseases:**

During the COVID-19 pandemic, nanobodies derived from llamas were found to bind strongly to the SARS-CoV-2 virus and prevent it from entering human cells. This discovery helped speed up research into llama-based treatments for severe coronavirus infections. [79]

### **Other Infectious Diseases:**

The use of llama antibodies in the treatment and prevention of other bacterial and viral infections is being investigated in ongoing studies, increasing their significance in the field of global health research. [80]

## **(19) HORSE**

The horse (*Equus ferrus caballus*) is a large, herbivorous mammal that has been domesticated for thousands of years for work, transport, companionship, and research. Horses typically have a lifespan of 30–35 years, with mature adults weighing between 300–1000 kg, depending on breed and nutrition. Normal physiological parameters include a body temperature of 99–105°F (37.2–40.5°C), heart rate of 28–48 beats per minute, and a respiration rate of 8–16 breaths per minute.

### **EXPERIMENTAL USE OF HORSE**

Horses have historically played a significant role in biomedical, pharmacological, and veterinary research, owing to their size, physiology, and unique biological characteristics. They continue to be valuable both in human medicine and veterinary sciences.

### **Equine-Assisted Services**

Research has explored the effects of equine-assisted services on human stress levels, including the ability of horses to perceive and respond to

human emotions and stress. Such studies provide insight into animal–human interactions and their therapeutic potential [81].

### **Antitoxin Development:**

Horses contributed to the development of therapies for diphtheria and dysentery by producing antitoxins, which were highly effective in neutralizing bacterial toxins [82].

## **(20) CAENORHABDITIS ELEGANS**

*Caenorhabditis elegans* is a eukaryotic, multi-organ, transparent nematode (Hermaphrodite worm) which lives in the interstitial water of soil and survives by feeding on microbes. It is a free-living of 1mm in length that lives in temperate soil environment. *C.elegans* has two natural genders female(XX) and male (XO). The female – hermaphrodite can produce up to 300 progenies via self fertilization which produces easy generation of genetically identical progeny. This soil nematode offered great potential for genetic analysis, partly because of its rapid (3-day) life cycle, small size (1.5-mm-long adult), and ease of laboratory cultivation.

### **EXPERIMENTAL USE OF C.ELEGANS**

*C. elegans* is an invaluable model organism in biomedical research due to its small size, short life cycle, transparent body, and fully sequenced genome. It can be easily cultured in Petri dishes with *Escherichia coli* as a food source, and large populations can be maintained with minimal resources. These advantages have made *C. elegans* a cornerstone in experimental biology.

### **Parkinson's studies**

**Parkinson's** disease is studied using *C. elegans* as a model. Despite the absence of the human alpha-synuclein (PARK1) gene in its genome, transgenic strains that overexpress this protein have been



created. Researchers can examine the effects of pathological alpha-synuclein aggregation on neuronal health and degeneration using these models [83].

### Disease Modeling

*C. elegans* has been extensively used to study various human diseases, including Alzheimer's disease, polycystic kidney disease, and cancer. By replicating disease-related genetic and molecular processes in the worm, researchers gain valuable insights into the mechanisms underlying these conditions [84].

### Developmental Biology and neurobiology

Research on *C. elegans* has greatly advanced our understanding of important developmental processes like cell differentiation, organogenesis, and the formation of complex tissues. Its transparent body and invariant cell lineage allow scientists to track cell development remarkably accurately. The simple but well-mapped nervous system of *C. elegans* provides an excellent framework for researching neuronal development, synaptic function, and the effects of drugs or toxins on the nervous system. These investigations have improved our knowledge of both fundamental neurobiology and neurodegeneration associated with disease [85].

### CONCLUSION

Pharmacology's study of animal models serves as a link between laboratory discoveries and appropriate human treatment. Every species offers a different perspective on physiology, the course of disease, and drug response, from the tiny fruit fly to the sophisticated primates. Their diversity allows researchers to answer different aspects, like zebrafish and medaka provides an illuminate study of genetics and developmental biology, rodents provide robust models for neurology and

metabolism, while the other primates like marmosets and African green monkeys bring a closer understanding of human -specific conditions. Although unconventional models such as llamas, seahorse and pigeons expand the horizons of biomedical research, a novel perspective in immunology, reproduction and behavioral science. When taken as a whole, these species offer insights into developing safer, more potent treatments that eventually enhance human health. Animal models continue to be essential on the path from molecules to pharmaceuticals.

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