

3050 Spruce Street, St. Louis, MO 63103 USA
Tel: (800) 521-8956 (314) 771-5765 Fax: (800) 325-5052 (314) 771-5757
email: techservice@sial.com sigma-aldrich.com

Product Information

Anti-p53 antibody, Mouse monoclonal clone DO-2, purified from hybridoma cell culture

Product Number P7874

Product Description

Monoclonal Anti-p53 (mouse IgG1 isotype) is derived from the hybridoma DO-2 produced by the fusion of mouse myeloma cells (SP2 cells) and splenocytes from BALB/c mice immunized with recombinant human wild type p53 (Gene ID: 7157). The isotype is determined by double diffusion immunoassay using Mouse Monoclonal Antibody Isotyping Reagents, Product Number ISO2.

Monoclonal Anti-p53 recognizes human p53.¹ The antibody may be used in various applications including: immunoblotting (~53 kDa), immunoprecipitation,² and DNase I footprint.¹ The antibody epitope resides between amino acids 10-16 of human p53.¹

The p53 gene, located on chromosome 17p, is the most commonly mutated gene in human cancer with more than 500 mutations described. These mutations are found in various types of malignancies, hematologic as well as solid tumors. However, not all mutants are equivalent in terms of biological activity. The p53 protein is highly conserved and expressed in normal tissues. Wild-type p53 has been shown to be a sequence-specific transcription factor, directly interacting with various cellular and viral proteins. Intact p53 function is essential for the maintenance of the non-tumorogenic phenotype of cells. Thus, p53 plays a vital role in suppressing the development of cancer.

The p53 tumor suppressor protein is important in the cellular response to DNA damage and other genomic aberrations. Cells exposed to DNA-damaging agents such as ionizing radiation, UV radiation, and chemical agents initiate a complex response that includes the inhibition of cell cycle progression until damage is repaired. If the DNA damage is beyond repair, cells may enter a prolonged state of arrest or undergo a programmed cell death known as apoptosis, thereby maintaining genetic stability in the organism. In response to DNA damage, p53 is phosphorylated at multiple sites by several protein kinases.

Phosphorylation of p53 at Ser¹⁵ by ATM, ATR, and DNAPK leads to a reduced interaction with its negative regulator MDM2 and accumulation of p53 protein. Chk2 and Chk1 can phosphorylate p53 at Ser²⁰, which enhances its activity, tetramerization, and stability. Elevation of p53 protein induces the transcriptional activation of multiple genes, including p21^{waf1}. p21^{waf1} interacts directly with cyclin dependent kinases, important for cell cycle progression, thereby inhibiting their activity and resulting in cell cycle arrest.³⁻¹¹

Reagent

Supplied as a solution in 0.01 M phosphate buffered saline, pH 7.4, containing 15 mM sodium azide as a preservative.

Antibody concentration: ~2 mg/mL

Precautions and Disclaimer

For R&D use only. Not for drug, household, or other uses. Please consult the Safety Data Sheet for information regarding hazards and safe handling practices.

Storage/Stability

Storage Temperature –20 °C Storage/Stability

For continuous use, store at 2–8 °C for up to one month. For extended storage, freeze in working aliquots. Repeated freezing and thawing, or storage in "frost-free" freezers, is not recommended. If slight turbidity occurs upon prolonged storage, clarify the solution by centrifugation before use. Working dilution samples should be discarded if not used within 12 hours.

Product Profile

Immunoblotting: a working concentration of 1–2 μ g/mL is recommended using total cell extract of A431 cells.

<u>Note</u>: In order to obtain the best results in various techniques and preparations, it is recommended to determine the optimal working dilution by titration.

References

- 1. Cain, C. et al., *J. Biol. Chem.*, **275**, 39944-39953 (2000).
- Menendez, S. et al., J. Biol. Chem., 278, 18720-18729 (2003).
- 3. Hirao, A. et al., Science, 287, 1824-1827 (2000).
- 4. Hung, J. et al., *Acta Orthop. Scand. Suppl.*, **273**, 68-73 (1997).
- 5. Levine, A.J., Cell, 88, 323-331 (1997).

- 6. Milczarek, G.J. et al., Life Sci., 60, 1-11 (1997).
- 7. Milner, J., Pathol. Biol., 45, 797-803 (1997).
- 8. Prives, C., J. Pathol., 187, 122-126 (1999).
- 9. Prokocimer, M. et al., Hum. Mutat., 12, 4-18 (1998).
- 10. Shaw, P.H., *Pathol. Res. Pract.*, **192**, 669-675 (1996).
- 11. Shieh, S.Y., et al., Cell, 91, 325-334 (1997).

DS,EK,KAA,PHC,MAM 01/19-1