

Marcos Malumbres, PhD

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## **Education**

1993 Ph.D., University of León, León, Spain  
1987 B.S., University of Navarra, Pamplona

#### **Research and Professional Experience**

2005-	Head, Cell Division & Cancer Group (CNIO)
2004-	Staff Investigator, Spanish National Council of Research (CSIC), Spain
1999-2004	Staff Scientist, CNIO, Madrid. (Mariano Barbacid, Supervisor)
1995-1998	Assistant Research Scientist, New York University Medical Center, New York, USA, ( Angel Pellicer, Supervisor)
1993-1994	Postdoctoral Fellow, New York University Medical Center, New York, USA, ( Angel Pellicer, Supervisor)
1988-1988	Visiting Scientist, Zentrum für Molekulare Biologie im Heidelberg ( Hermann Bujard, Supervisor)
1988-1993	Graduate Fellow, Dep. of Microbiology, Univ. of León, León, Spain (Juan F. Martín, Supervisor)

### **Awards and Honors**

2007-2008 Council Member, European Association for Cancer Research  
2007- Honorary member, Balkan Union of Oncology  
2005 SEBBM Beckman/Coulter Award 2005  
2001- Honorary Professor, Univ. Autónoma de Madrid  
1994 "Juan Abelló" Prize of the Real Academia de Doctores

## Professional Society Memberships and Committees

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- 2010- American Society of Microbiology  
2009- ISCIII Animal Committee  
2006-2008 CNIO Animal Committee  
2001- European Association for Cancer Research  
1998- Spanish Society of Biochemistry and Molecular Biology (SEBBM)

## Teaching and Administrative Responsibilities

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Post-graduate and doctoral coursers: Universidad Autónoma de Madrid, Universidad Complutense de Madrid, Univ. Del País Vasco (Bilbao), Gulbenkian Institute (Portugal), etc.

## Peer-Review Activities

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### Editorial Boards

- 2007- *J. BUON*  
2008- *Current Medical Chemistry*  
2010- *Molecular Cancer*, Associate Editor  
2010- *Genes & Cancer*, Editorial Board founding member, 2010  
2011- *Frontiers in Molecular and Cellular Oncology*, Associate Editor  
2011- *microRNA*, Editorial Board founding member,

### Journal Peer-Review

*Science*, *Nature Reviews series* (*Nat. Rev. Cancer*, *Nat. Rev. Mol. Cell Biol.*, *Nat Rev. Drug Discover*), *Nat. Med.*, *Nat. Cell Biol.*, *Nat. Communications*, *Nat. Struc. Biol.*, *Mol. Syst. Biol.*, *Trends series*, *Cancer Cell*, *PLoS Biol.*, *EMBO J.*, *EMBO Reports*, *EMBO Mol. Med.*, *J. Clin. Invest.*, *PNAS*, *J. Nat. Cancer Inst.*, *J. Cell Biol.*, *Mol. Biol. Cell*, *Mol. Cell. Biol.*, *Cancer Res.*, *Oncogene*, *Nucleic Acids Res.*, *Carcinogenesis*, *Cell Cycle*, *FEBS Letters*, *Bioessays* and others.

### Grant Review

Ad hoc reviewer for EU 7FP, Cancer Research UK, The Wellcome Trust, Association for International Cancer Research. Dutch Cancer Society, Research Grants Council (Hong Kong), INCA (France) and Spanish National (MICINN, FIS, AECC) and Regional (Cataluña, Junta Andalucía, Xunta Galicia, etc.) Institutions. Staff member of the Spanish Agency of Evaluation of Research Projects (ANEP; since 2012).

### Grant Committees

Spanish Ministry of Innovation and Research (MICINN, 2001-2011), Ministry of Health (FIS 2006-2010), EU 7FP (2008), AECC (2007), Staff Spanish National Agency for Project Evaluation (ANEP; 2012-).

## International Conferences Organized

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CNIO Cancer Conference *The Cell Cycle and Cancer*, Centro Nacional de Investigaciones Oncológicas, Madrid, 2002 (co-organizers: Jiri Bartek, Charles J. Sherr)

*Workshop on Mouse Models in Cancer*, Barcelona, 2004.b(co-organizer)  
*Cell Cycle and Cancer Meeting*, Toulouse, March 2008 (organizing Committee).  
*Cell Cycle Regulators/Inhibitors & Cancer*, Vienna, February 2011 (organizing Committee).  
*Aneuploidy and Chromosomal Instability and Cancer*. CNIO, Madrid, 2013 (co-organizer)

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## Invited Conferences

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(Representative conferences in the last 5 years)

- 2008 Cell Cycle and Cancer Meeting, Toulouse, France
- 2008 MRC Research Centre, Cambridge, UK
- 2008 10<sup>th</sup> European Congress of Endocrinology, Berlin, Germany
- 2008 MRC Clinical Sciences Center, London, UK
- 2009 Institute for Molecular Pathology, Vienna, Austria
- 2009 Baylor College of Medicine, Houston, USA
- 2009 AACR Annual Meeting, Denver, USA
- 2009 The First EMBO Meeting, Amsterdam, The Netherlands
- 2009 IRBB, ICC, Barcelona, Spain
- 2009 Gulbenkian Institute, Oeiras, Lisbon
- 2010 Centre de Recherche en Biochimie Macromoléculaire, Montpellier
- 2010 Chromosome segregation and Aneuploidy Symposium, IBMC, Porto, Lisbon
- 2011 Cell Cycle Inhibitors and Cancer Workshop, Vienna
- 2011 The Institute for Cancer Research, London
- 2011 Univ. Of Lisboa, Lisbon
- 2011 Cell Signaling Networks, Merida, México
- 2011 Inproteolysis 2011, Valencia
- 2011 Univ. P. Sabatier, Toulouse
- 2011 Nordisk Mitotic Network, Copenhagen
- 2012 Cancer Research Center (CIC) Salamanca
- 2012 Institute for Research in Immunology and Cancer (IRIC), Université de Montréal
- 2012 Jacques-Monod Conference of the Cell Cycle, Roscoff, 2012
- 2012 EMBO Workshop on Early Mouse Development, Cambridge, UK
- 2012 EMBL Monterontondo, Italy
- 2012 56th Nat. Meeting of the Italian Society of Biochemistry and Molecular Biology, Chieti
- 2013 EMBO Workshop, Cell Death in Mitosis meeting, Obergurgl, Austria
- 2013 AACR Annual Meeting, Washington

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## Patents

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Bueno, M.J., Pérez de Castro, I., Fernández-Piqueras, J., Malumbres, M. (2008) Use of microRNA-203 for the manufacture of a medicament for treating cancer in humans and for

reducing V-abl Abelson murine leukemia viral oncogene homolog 1 (ABL1) and B cell receptor (BCR)-ABL1 expression and tumor cell proliferation.

Patent Number(s): WO2009112625-A1; ES2325726-A1; ES2325726-B1

Alvárez, C., Diéguez, C., García-Lavandeira, M., Malumbres, M. (2008) Isolation of multipotent hypophysary cells and in vitro differentiation thereof.

Pub. No.: WO/2010/061030. International Application No.: PCT/ES2009/070530

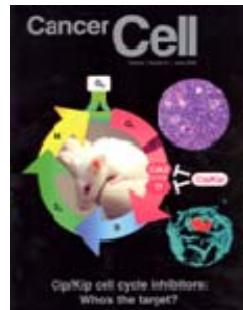
## Publications

1. Malumbres, M., Mateos, L.M., Guerrero, C. and Martín, J.F. (1988) Nucleotide sequence of the threonine synthase (*thc*) gene of *Brevibacterium lactofermentum*. *Nucleic Acids Res.* 16, 9859. [PMID: 3186450]
2. Martín, J.F., Cadenas, R.F., Malumbres, M., Mateos, L.M., Guerrero, C. and Gil, J.A. (1990) Construction and utilization of promoter-probe and expression vectors in corynebacteria. Characterization of corynebacterial promoters. In: *Genetics of Industrial Microorganisms '90*. Heslot, H., Davies, J., Florent, J., Bobichon, L., Durant, G., Penasse, L., eds. Société Française de Microbiologie, Strasbourg, pp. 283-292.
3. Martín, J.F., Mateos, L.M., Cadenas, R.F., Guerrero, C., Malumbres, M., Colina, A. and Gil, J.A. (1990) Molecular genetics of corynebacteria: cloning and characterization of the tryptophan operon and the genes of the threonine biosynthetic pathway. In: *Microbiology Applications in Food Biotechnology*. Nga, B.H., Lee, Y.K., eds. Elsevier, London, pp. 20-26.
4. Guerrero, C., Mateos, L.M., Malumbres, M. and Martín, J.F. (1992) The bleomycin resistance gene from Tn5 is an excellent marker for transformation of corynebacteria. *Appl. Microbiol. Biotechnol.* 36, 759-762. [PMID: 1373065]
5. Pisabarro, A., Malumbres, M., Mateos, L.M., Oguiza, J.A. and Martín, J.F. (1993) A cluster of three genes, *dapA*, *orf2*, and *dapB*, of *Brevibacterium lactofermentum* encodes dihydridipicolinate synthase, dihydridipicolinate reductase and a third polypeptide of unknown function. *J. Bacteriol.* 175, 2743-2749. [PMID: 8478336]
6. Malumbres, L. and Malumbres, M. (1993) Promoter structure recognition in corynebacterial DNA sequences by artificial neural networks. In: *Industrial & Cognitive Applications of Neural Networks*. EC2 Publishing, Nanterre, pp. 155-164.
7. Coque, J.J.R., Malumbres, M., Martín, J.F. and Liras, P. (1993) Analysis of the codon usage of the cephamycin C producer *Nocardia lactamdurans*. *FEMS Microbiol. Lett.* 110, 91-96.
8. Oguiza, J.A., Malumbres, M., Eriani, G., Pisabarro, A., Mateos, L.M., Gangloff, J. and Martín, J.F. (1993) A gene encoding arginyl-tRNA synthetase is located in the upstream region of the *lysA* gene in *Brevibacterium lactofermentum*: Regulation of the *argS-lysA* cluster expression by arginine. *J. Bacteriol.* 175, 7356-7362. [PMID: 8226683]
9. Malumbres, M., Gil, J.A. and Martín, J.F. (1993) Codon preference in corynebacteria. *Gene* 134, 15-24. [PMID: 8244028]
10. Guerrero, C., Mateos, L.M., Malumbres, M. and Martín, J.F. (1994) Directed mutagenesis of a regulatory palindromic sequence upstream from the *Brevibacterium lactofermentum* tryptophan operon. *Gene* 138, 35-41. [PMID: 7510262]

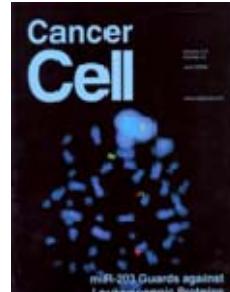
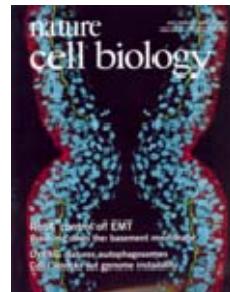
11. Malumbres, M., Mateos, L.M., Lumbreras, M.A., Guerrero, C. and Martín, J.F. (1994) Analysis and expression of the *thrC* gene of *Brevibacterium lactofermentum* and characterization of the encoded threonine synthase. *Appl. Environ. Microbiol.* 60, 2209-2219. [PMID: 8074505]
12. Mateos, L.M., Pisabarro, A., Pátek, M., Malumbres, M., Guerrero, C., Eikmanns, B.J., Sahm, H. and Martín, J.F. (1994) Transcriptional analysis and regulatory signals of the *hom-thrB* cluster of *Brevibacterium lactofermentum*. *J. Bacteriol.* 176, 7362-7371. [PMID: 7961509]
13. Malumbres, M., Mateos, L.M. and Martín, J.F. (1995) Microorganisms for amino acid production: *Escherichia coli* and corynebacteria. In: *Food Biotechnology: Microorganisms*. Hui, Y.H., Khachatourians, G.G., eds. VCH Publishers, Inc, pp. 423-469.
14. Malumbres, M., Mateos, L.M., Guerrero, C. and Martín, J.F. (1995) Molecular cloning of the *hom-thrC-thrB* cluster from *Bacillus* sp. ULM1: Expression of the *thrC* gene in *Escherichia coli* and corynebacteria, and evolutionary relationships of the threonine genes. *Folia Microbiol.* 40, 595-606. [PMID: 8768250]
15. Oguiza, J.A., Marcos, A.T., Malumbres, M. and Martín, J.F. (1996) Multiple sigma factor genes in *Brevibacterium lactofermentum*: characterization of *sigA* and *sigB*. *J. Bacteriol.* 178, 550-553. [PMID: 8550480]
16. Malumbres, M. and Martín, J.F. (1996) Molecular control mechanisms of lysine and threonine biosynthesis in amino acid-producing corynebacteria: Redirecting carbon flow. *FEMS Microbiol. Lett.* 143, 103-114. [PMID: 8837462]
17. Oguiza, J.A., Marcos, A.T., Malumbres, M. and Martín, J.F. (1996) Sequence and transcriptional analysis of the *galE* gene encoding the UDP-galactosidase of *Brevibacterium lactofermentum*. *Gene* 177, 103-107. [PMID: 8921853]
18. Malumbres, M., Pérez de Castro, I., Santos, J., Meléndez, B., Mangues, R., Serrano, M., Pellicer, A. and Fernández-Piqueras, J. (1997) Inactivation of the cyclin-dependent kinase inhibitor p15<sup>INK4b</sup> by deletion and *de novo* methylation with independence of p16<sup>INK4a</sup> alterations in murine primary T-cell lymphomas. *Oncogene* 14, 1361-1370. [PMID: 9178896]
19. Malumbres, M., Mangues, R., Ferrer, N., Lu, S. and Pellicer, A. (1997) Isolation of high molecular weight DNA for reliable genotyping of transgenic mice. *BioTechniques* 22, 1114-1119. [PMID: 9187761]
20. Malumbres, M., Perez de Castro, I., Santos, J., Perez-Olle, R., Fernandez-Piqueras, J. and Pellicer, A. (1998) An AC-repeat adjacent to mouse Cdkn2B allows the detection of specific allelic losses in the p15<sup>INK4b</sup> and p16<sup>INK4a</sup> tumor suppressor genes. *Mamm. Genome* 9, 183-185. [PMID: 9501299]
21. Mangues, R., Corral, T., Kohl, N.E., Symmans, W.F., Lu, S., Malumbres, M., Gibbs, J.B., Oliff, A. and Pellicer, A. (1998) Antitumor effect of a farnesyl-protein transferase inhibitor in mammary and lymphoid tumors overexpressing N-ras in transgenic mice. *Cancer Res.* 58, 1253-1259. [PMID: 9515813]
22. Malumbres, M. and Pellicer, A. (1998) Ras pathways to cell cycle control and cell transformation. *Front. Biosci.* 3, d887-912. [PMID: 9696882]
23. Malumbres, M., Pérez de Castro, I., Santos, J., Fernández-Piqueras, J. and Pellicer, A. (1999) Hypermethylation of the cell cycle inhibitor p15<sup>INK4b</sup> 3'-untranslated region interferes with its transcriptional regulation in primary lymphomas. *Oncogene* 18, 385-396. [PMID: 9927195]
24. Malumbres, M. and Pellicer, A. (1999) Ras signaling in cell cycle regulation and its role in tumor development. *Rev. Oncología* 1, 66-76.

25. Pérez de Castro, I., Malumbres, M., Santos, J., Pellicer, A. and Fernández-Piqueras, J. (1999) Cooperative alterations of Rb-pathway regulators in mouse primary T-cell lymphomas. *Carcinogenesis* 20, 1675-1682. [PMID: 10469610]
26. García-España, A., Biria, S., Malumbres, M., Levin, B., Meruelo, D. and Pellicer, A. (1999) Targeted gene transfer system using a streptavidine-transforming growth factor-a chimeric protein. *DNA Cell Biol.* 18, 743-749. [PMID: 10541433]
27. Pellicer, A. and Malumbres, M. (2000) Bases moleculares de la transformació neoplásica. In: *Llicons de Patologia Molecular*, González-Sastre, F. and Guinovart, J.J., eds. Springer-Verlag, Heidelberg, Barcelona.
28. Meléndez, B.\*, Malumbres, M.\* , Pérez de Castro, I., Santos, J., Pellicer, A. and Fernández-Piqueras, J. (2000) Characterization of the murine p19<sup>ARF</sup> promoter CpG island and its methylation pattern in primary lymphomas. *Carcinogenesis* 21, 817-821. [PMID: 10753221]
29. Malumbres, M., Pérez de Castro, I., Hernández, M.I., Jiménez, M., Corral, M.T. and Pellicer, A. (2000) Cellular response to oncogenic Ras involves induction of the Cdk4 and Cdk6 inhibitor p15<sup>INK4b</sup>. *Mol. Cell. Biol.*, 20, 2915-2925. [PMID: 10733595]
30. Hernández-Muñoz, M.I., Malumbres, M., Leonardi, P. and Pellicer, A. (2000) The Rgr oncogene (homologous to RalGDS) induces transformation and gene expression by activating Ras, Ral and Rho mediatedpathways. *Oncogene*, 19, 2745-2757. [PMID: 10851075]
31. Latres, E., Malumbres, M., Sotillo, R., Martín, J., Ortega, S., Martín-Caballero, J., Flores, J.M., Cordon-Cardo, C. and Barbacid. M. (2000) Limited overlapping roles of P15<sup>INK4b</sup> and P18<sup>INK4c</sup> cell cycle inhibitors in proliferation and tumorigenesis. *EMBO J.*, 19, 3496-3506.
32. Malumbres, M., Ortega, S. and Barbacid, M. (2000) Genetic analysis of cyclin-dependent kinases and their inhibitors. *Biol. Chem.*, 381, 827-838.
33. Esteban, L.M., Vicario-Abejon, C., Fernandez-Salguero, P., Fernandez-Medarde, A., Swaminathan, N., Yienger, K., Lopez, E., Malumbres, M., McKay, R., Ward, J.M., Pellicer, A., Santos, E. (2001) Targeted genomic disruption of H-ras and N-ras, individually or in combination, reveals the dispensability of both loci for mouse growth and development. *Mol. Cell. Biol.* 21, 1444-1452.
34. Sotillo, S., Dubus, P., Martín, J., de la Cueva, E., Ortega, S., Malumbres, M. and Barbacid, M. (2001) Wide spectrum of tumors in knock in mice carrying a Cdk4 protein insensitive to INK4 inhibitors. *EMBO J.* 20, 6637-6647.
35. Sotillo, R., García, J. F., Ortega, S., Martín, J., Dubus, P., Barbacid, M and Malumbres M. (2001) Invasive melanoma in Cdk4 targeted mice. *Proc. Natl. Acad. Sci. USA* 98, 13312-13317.
36. Malumbres, M. and Barbacid, M. (2001) To cycle or not to cycle: a critical decision in cancer. *Nature Reviews Cancer* 1, 222-231.
37. Ortega, S., Malumbres, M. and Barbacid, M. (2002) Cdk4 and their INK4 inhibitors in tumor biology. *Biochim. Biophys. Acta* 87513, 1-15.
38. Diaz, R., Ahn, D., Lopez-Barcons, L., Malumbres, M., Pérez de Castro, I., Lue, J., Ferrer, N., Mangues, R., Tsong, J., García, R., Pérez-Soler, R. and Pellicer, A. (2002) The N-ras protooncogene can suppress the malignant phenotype in the presence or absence of its oncogene. *Cancer Res.* 62, 4514-4518.
39. Ortega, S., Malumbres, M. and Barbacid, M. (2002) Cell cycle and Cancer: The G1 restriction point and the G1/S transition. *Curr. Genomics* 3, 245-263.

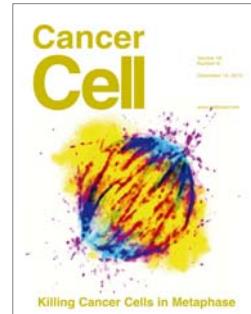
40. Malumbres, M. and Carnero, A. (2003) Cell cycle deregulation: a common motif in cancer. *Progress Cell Cycle Res.* 5, 5-18.
41. Pérez de Castro, I., Diaz, R., Malumbres, M., Hernández, M.I., Jagirdar, J., Jiménez, M., Ahn, D. and Pellicer A. (2003) Mice deficient for N-ras: Impaired antiviral immune response and T-cell function. *Cancer Res.* 63, 1615-1622.
42. Malumbres, M. and Barbacid, M. (2003) *RAS* oncogenes: the first 30 years. *Nature Rev. Cancer* 3, 459-465.
43. Malumbres, M., Hunt, S.L., Sotillo, R., Martín, J., Odajima, J., Martín, A., Dubus, P., Ortega, S., Barbacid, M. (2003) Driving the cell cycle to cancer. *Adv. Exp. Med. Biol.* 532, 1-11.
44. Martín, J., Hunt, S.L., Dubus, P., Sotillo, R., Néhmé-Pelluard, F., Magnuson, M.A., Parlow, A.F., Malumbres, M., Ortega, S. and Barbacid, M. (2003) Genetic rescue of Cdk4 *null* mice restores pancreatic β-cell proliferation but not homeostatic cell number. *Oncogene* 22, 5261-5269.
45. Ortega, S., Prieto, I., Odajima, J., Martín, A., Dubus, P., Sotillo, R., Barbero, J.L., Malumbres, M. and Barbacid, M. (2003) Cyclin dependent kinase 2 is essential for meiosis but not for mitotic cell division in mice. *Nature Genet.* 35, 25-31.
46. Wolff, L., Garin, M.T., Koller, R., Bies, J., Liao, W., Malumbres, M., Tessarollo, L., Powell, D. and Perella, C. (2003) Hypermethylation of the *Ink4b* Locus in Murine Myeloid Leukemia and Increased Susceptibility to Leukemia in p15<sup>Ink4b</sup>-deficient Mice. *Oncogene* 22, 9265-9274.
47. Malumbres, M.\*, Sotillo, R., Santamaría, D., Galán, J., Cerezo, A., Ortega, S., Dubus, P. and Barbacid, M.\* (2004) Mammalian cells cycle without the D-type cyclin-dependent kinases Cdk4 and Cdk6. *Cell* 118, 493-504. [\*Co-corresponding authors]  
*News & Views:* Murray, A.W. (2004) Recycling the cell cycle: cyclins revisited. *Cell* 116, 221-234. Cycling without cyclins. *The Scientist*, August 20, 2004.
48. Auwerx, J., Avner, P., Baldock, R., Ballabio, A., Balling, R., Barbacid, M., Berns, A., Bradley, A., Brown, S., Carmeliet, P., Chambon, P., Cox, R., Davidson, D., Davies, K., Duboule, D., Forejt, J., Granucci, F., Hastie, N., Hrabé de Angelis, M., Jackson, I., Kioussis, D., Kollias, K., Lathrop, M., Lendahl, U., Malumbres, M., Melchner, H. von, Müller, W., Partanen, J., Ricciardi-Castagnoli, P., Rigby, P., Rosen, B., Rosenthal, N., Skarnes, B., Stewart, A.F., Thornton, J., Tocchini-Valentini, G., Wagner, E., Wahli, W. and Wurst, W. (2004) The European dimension for the mouse genome mutagenesis program. *Nat. Genet.* 36, 925-927.
49. Malumbres, M. (2005) Revisiting the “Cdk-centric” view of the mammalian cell cycle. *Cell Cycle* 4, 206-210.
50. Pérez de Castro, I., Benet, M., Jiménez, M., Alzabin, S., Malumbres, M. and Pellicer, A. (2005) Mouse p10, an alternative spliced form of p15INK4b, inhibits cell cycle progression and malignant transformation. *Cancer Res.* 65, 3249-3256.
51. Sotillo, R., Renner, O., Dubus, P., Ruiz-Cabello, J., Martín-Caballero, J., Barbacid, M., Carnero, A. and Malumbres, M. (2005) Cooperation Between Cdk4 and p27<sup>Kip1</sup> in Tumor Development: a Preclinical Model to Evaluate Cell Cycle Inhibitors with Therapeutic Activity. *Cancer Res.*, 65, 3846-3852.
52. Martín, A., Odajima, J., Hunt, S.L., Dubus, P., Ortega, S., Malumbres, M.\* and Barbacid, M.\* (2005) Cell Cycle Inhibition and Tumor Suppression by p21<sup>Cip1</sup> and p27<sup>Kip1</sup> are Independent of Cdk2. *Cancer Cell* 7, 591-598. [\*Co-corresponding authors]



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54. Malumbres, M. and Barbacid, M. (2005) Mammalian cyclin-dependent kinases. *Trends Biochem. Sci.*, 30, 630-641.
55. Barbacid, M., Ortega, S., Sotillo, R., Odajima, J., Martín, A., Santamaría, D., Dubus, P. and Malumbres, M. (2005) Cell cycle and cancer: genetic analysis of the role of cyclin-dependent kinases. *Cold Spring Harb. Symp. Quant. Biol.* 70, 233-240.
56. Steitz, J., Büchs, S., Tormo, D., Ferrer, A., Wenzel, J., Huber, C., Wölfel, T., Barbacid, M., Malumbres, M. and Tüting, T. (2006) Evaluation of genetic melanoma vaccines in cdk4-mutant mice provides evidence for immunological tolerance against autochthonous melanomas in the skin. *Int. J. Cancer* 118, 373-380.
57. Duensing, A., Liu, Y., Tseng, M., Malumbres, M., Barbacid, M. and Duesing, S. (2006) Cyclin-dependent kinase 2 is dispensable for normal centrosome duplication but required for oncogene-induced centrosome overduplication. *Oncogene* 25, 2943-2949.
58. Malumbres, M., Dubus, P. and Ortega, S. (2006) Mouse models to study the *in vivo* function of Cyclin-dependent kinases in normal homeostasis and tumor development. In: Inhibitors of cyclin-dependent kinases as anti-tumor agents (Smith, P.J. and Yue, E.W., eds.) CRC Press, Boca Raton. pp. 55-83.
59. Malumbres, M. and Barbacid, M. (2006) Is Cyclin D1/Cdk4 kinase a bona-fide cancer target? *Cancer Cell* 9, 2-4.
60. Hacker, E., Muller, H.K., Irwin, N., Gabrielli, B., Lincoln, D., Pavéy, S., Powell, M.B., Malumbres, M., Barbacid, M., Hayward, N. and Walker, G. (2006) Spontaneous and UV radiation-induced multiple metastatic melanomas in *Cdk4<sup>R24C/R24C</sup>/TPras* mice. *Cancer Res.* 66, 2946-2952.
61. Malumbres, M. (2006) Therapeutic opportunities to control tumor cell cycles. *Clin. Transl. Oncol.* 8, 399-408.
62. Tormo, D., Ferrer, A., Gaffal, E., Wenzel, J., Basner-Tschakarjan, E., Steitz, J., Heukamp, L.C., Gutgemann, I., Buettner, R., Malumbres, M., Barbacid, M., Merlini, G., Tuting, T. (2006). Rapid Growth of Invasive Metastatic Melanoma in Carcinogen-Treated Hepatocyte Growth Factor/Scatter Factor-Transgenic Mice Carrying an Oncogenic CDK4 Mutation. *Am. J. Pathol.* 169, 665-672.
63. Blazquez, C., Carracedo, A., Barrado, L., Real, P.J., Fernández-Luna, J.L., Velasco, G., Malumbres, M. and Guzman, M. (2006) Cannabinoid receptors as novel targets for the treatment of melanoma. *FASEB J.* 20, 2633-2635.
64. Malumbres, M. (2006) Preclinical models for cell cycle-targeted therapies. *Adv. Exp. Med. Biol.* 587, 139-147.
65. Malumbres, M. and Barbacid, M. (2007) Cell cycle kinases in cancer. *Curr. Opin. Genet. Dev.* 17, 60-65.
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