

QUIMICA BIOLOGICA II

Programa teórico

Metabolismo del Hemo y enfermedades relacionadas

Biosíntesis del hemo, enzimas involucradas, propiedades y mecanismos de reacción. Regulación de la biosíntesis del hemo en células hepáticas y eritroides. Catabolismo del hemo: rol de la Hemo oxigenasa y sus productos de reacción. Hemoproteínas. Control de los niveles intracelulares de hemo, transportadores. Efectos pleiotrópicos del hemo sobre diferentes eventos biológicos. Hemo como regulador de la homeostasis celular. Enfermedades relacionadas con el camino biosintético del Hemo: Porfirias. Biología Molecular y Farmacogenética de las Porfirias. Sistema metabolizante de drogas: Polimorfismos del citocromo P450 y su relación con el desencadenamiento de las porfirias.. Porfiria experimental: modelos farmacológicos y genéticos. El laboratorio de las porfirinas.

Cinética enzimática

Cinética del estado estacionario. Ecuación de Michaelis-Menten para un sustrato. Reacciones en la que intervienen de 2 ó más sustratos y productos. Mecanismos de reacción. Modelos de velocidades iniciales: intersectantes y paralelos. Estudios de inhibición: por producto, producto alterno, dead-end y sustrato. Estudios de intercambio isotópico. Gráficas obtenidas para los distintos mecanismos enzimáticos. Efecto del pH y la temperatura sobre la actividad enzimática. Predicción y validación de mecanismos de reacción.

Regulación de la actividad enzimática. Enzimas alostéricas: Modelos. Efectos Homotrópicos y Heterotrópicos. Cooperatividad positiva, negativa y mixta. Reactividad de la mitad de los sitios. Mecanismo Flip-Flop. Cooperatividad en enzimas monoméricas. Aplicación a casos reales. Significado fisiológico del comportamiento alostérico.

Cáncer

Bioquímica y biología de la célula tumoral. Angiogénesis. Invasión y metástasis. Oncogenes y genes supresores tumorales. Terapias anticancerígenos:

Quimioterapia, Bioterapia, Terapia fotodinámica. Nuevos tratamientos y perspectivas futuras. Proteómica: técnicas utilizadas y su aplicación en el diagnóstico y tratamiento del cáncer. Estrés oxidativo e inflamación en el proceso tumoral.

Bibliografía

Unidad Hemo

* **“Porfirias y Porfirinas. Aspectos clínicos, bioquímicos y biología molecular”.**

Batlle A *Acta Bioquímica Clínica Latinoamericana*. (1997)

* **Molecular epidemiology and diagnosis of PBG deaminase gene defects in acute intermittent porphyria** Puy H et al (1997) *Am J Hum Genet* 60: 1373-83

* **Familial porphyria cutanea tarda: characterization of seven novel uroporphyrinogen decarboxylase mutations and frequency of common hemochromatosis alleles.** Mendez M, , Lonnie Sorkin, María Victoria Rossetti, Kenneth H. Astrin, Alcira M.del Carmen Batlle, Victoria E Parera, Gerardo Aizencang, Robert J Desnick. (1998) *Am J Hum Genet* 63: 1363-75

* **Identification and characterization of Hydroxymethylbilane Synthase mutations causing Acute Intermittent Porphyria: Evidence for an ancestral founder of the common G111R mutation.** De Siervi A, Rossetti MV, Parera V, Astrin KH, Aizencang GI, Glass IA, Batlle A, Desnick RJ. (1999) *Am J Gen Genet* 86: 366

* **The heme oxygenase pathway and its interaction with nitric oxide in the control of cellular homeostasis.** Foresti R, Motterlini R. (1999) *Free Rad Res* 31: 459-475

* **Dynamics of haem oxygenase-1 expression and bilirubin production in cellular protection against oxidative stress.** Clark JE, Foresti R, Green C, Motterlini R. (2000) *Biochem J* 348: 615-619

* **The heme oxygenase system and its functions in the brain.** Maines MD (2000) *Cell Mol Biol* 46: 573-585

* **The management of porphyria cutanea tarda.** Sarkany RPE (2001) *Clin Exp Dermatol* 26: 225-232

* **Terminal steps of haem biosynthesis.** Dailey HA. *Biochem Soc Trans* (2002) 30: 590-595.

* **Homozygous acute intermittent porphyria in a 7-year-old boy with massive excretions of porphyrins and porphyrin precursors.** Hessels, J., Voortman, G., van der Wagen, A., van der Elzen, C., Scheffer, H. and Zuijderhoudt, F.M. (2004) *J. Inherited Metab. Dis.* 27, 19–27.

* **Acute intermittent porphyria: studies of the severe homozygous dominant disease provides insights into the neurologic attacks in acute porphyrias.** Solis, C., Martinez-Bermejo, A., Naidich, T.P., Kaufmann, W.E., Astrin, K.H., Bishop, D.F. and Desnick, R.J. (2004) *Arch. Neurol.*, 61: 1764–1770

* **Porphyrias** Kauppinen R. *Lancet* (2005) 365: 241-252. Review

- * **A Chilean boy with severe photosensitivity and finger shortening: the first case of homozygous variegate porphyria in South America.** Poblete-Gutiérrez P, Wolff C, Farias R, Frank J. *Br J Dermatol* (2006) 154:368-371.
- * **May 2006 update in porphobilinogen deaminase gene polymorphisms and mutations causing acute intermittent porphyria: comparison with the situation in Slavic population.** Hrdinka M, Puy H, Martasek P (2006) *Physiol Res* 55 (supl 2) S119-S136
- * **Genetic and Biochemical Studies in Argentinean Patients with Variegate Porphyria.** María Victoria Rossetti, Bárbara Xoana Granata, Jimena Giudice, Victoria Estela Parera, Alcira Batlle
BMC Med Genet (2008), 9:54.
- * **C-terminal deletions in the ALAS2 gene lead to gain of function and cause X-linked dominant protoporphyria without anemia or iron overload.** Whatley SD, Ducamp S, Gouya L, Grandchamp B, Beaumont C, Badminton MN et al. *Am J Hum Genet* (2008) 83:408-14.
- * **Erythropoietic protoporphyria.** Lecha M, Puy H, Deybach JC. *Orphanet J Rare Dis* (2009) 4:19
- * **Seasonal palmar keratoderma in erythropoietic protoporphyria indicates autosomal recessive inheritance.** Holme SA, Whatley SD, Roberts AG, Anstey AV, Elder GH, Ead RD, et al. *J Invest Dermatol* (2009) 129:599-605.
- * **Novel null-allele mutations and genotype-phenotype correlation in Argentinean patients with Erythropoietic Protoporphyria**
Victoria E. Parera, Rita H. Koole, Gardi Minderman, Maria V. Rossetti, Alcira Batlle & Felix.W.M De Rooij. *Mol Med* (2009)15:425-31.
- * **A homozygous mutation in the ferrochelatase gene underlies erythropoietic protoporphyria associated with palmar keratoderma.** Méndez M, Poblete-Gutiérrez P, Morán-Jiménez MJ, Rodríguez ME, Garrido-Astray MC, Fontanellas A, et al. *Br J Dermatol* (2009) 160:1330-4.
- * **The very first description of a patient with hepatoerythropoietic porphyria in Argentina. Biochemical and molecular studies.** Granata BX, Parera VE, Melito VA, Teijo MJ, Batlle A, Rossetti MV. *Cell Mol Biol (Noisy-le-grand)* 2009 Feb 16; 55(1):61-5.
- * **HFE Gene mutations in patients with altered iron metabolism in Argentina.** MV Rossetti, Méndez M, Afonso SG, Gerez EN, Batlle A, Muñoz A, Parera VE. *Cellular and Molecular Biology* (2009) 55 (2): 31-35.
- * **A homoallelic FECH mutation in a patient with both erythropoietic protoporphyria and palmar keratoderma.** Minder EI, Schneider-Yin X, Mamet R, Horev L, Neuenschwander S, Baumer A, et al. *J Eur Acad Dermatol Venereol* (2010) 24:1349-53.
- * **Feline acute intermittent porphyria: a phenocopy masquerading as an erythropoietic porphyria due to dominant and recessive hydroxymethylbilane synthase mutations.** Clavero S, Bishop DF, Haskins ME, Giger U, Kauppinen R, Desnick RJ. *Hum Mol Genet.* (2010) 19(4):584-96.
- * **Structural insight into human variegate porphyria disease.** Qin X, Tan Y, Wang L, Wang Z, Wang B, Wen X, Yang G, Xi Z, Shen Y. *FASEB J* (2011) 25:653-664.
- * **Identification of CYP3A5 and CYP2B6 Polymorphisms in Porphyria Cutanea Tarda associated to human Immunodeficiency virus.** JV Lavandera, Parera VE,

Rossetti MV, A Batlle, AM Buzaleh. Journal of Clinical and Experimental Dermatology Research (2011) <http://dx.doi.org/10.4172/2155-9554.S4-001>.

* **Structural analysis of heme proteins: implications for design and prediction.** T. Li, H. Bonkovsky, J. Guo BMC Structural Biology (2011) 11:1-13.

* **Correia MA, Sinclair P and De Matteis F.) Cytochrome P450 regulation: The interplay between its heme and apoprotein moieties in synthesis, assembly, repair and disposal.** Drug Metab Rev. (2011) 43(1): 1–26.
doi:10.3109/03602532.2010.515222.

* **Computational Prediction of Heme-Binding Residues by Exploiting Residue Interaction Network.** Liu R and Hu J. (2011) PLoS ONE 6: e25560

* **Heme oxygenase-1 comes back to endoplasmic reticulum.** Hong Pyo Kim, Hyun-Ock Pae, Sung Hun Back, Su Wol Chung, Je Moon Woo, Yong Son. Biochem. Biophys. Res. Comm. (2011) 404: 1-5.

* **Molecular analysis of the UROD gene in 17 Argentinean patients with familial porphyria cutanea tarda: Characterization of four novel mutations.** Manuel Méndez, María Victoria Rossetti , Sara Gómez-Abecia , María Josefa Morán-Jiménez, Victoria Parera, Alcira Batlle, Rafael Enríquez de Salamanca. Molecular Genetics and Metabolism (2012) 105(4):629-33

* **Functional associations of genetic variants involved in the clinical manifestation of erythropoietic protoporphyria in the Argentinean Population.** FP Colombo, MV Rossetti, M Mendez, JE Martínez, R Enriquez de Salamanca, A Batlle, VE Parera. Journal of European of the European Dermatology and Vennereology (2012) DOI: 10.1111/j.1468-3083.2012.04566.x

Unidad Cinética

* **The kinetics of enzyme-catalyzed reactions with two or more substrates or products. I. Nomenclature and rate equations.** 1963. Cleland, WW. *Biochim Biophys Acta*. (1989) 1000: 213-20

* **The kinetics of enzyme-catalyzed reactions with two or more substrates or products. II. Inhibition: nomenclature and theory.** Cleland, WW (1963) *Biochim Biophys Acta* 67: 173-87

* **The kinetics of enzyme-catalyzed reactions with two or more substrates or products. III. Prediction of initial velocity and inhibition patterns by inspection.** Cleland, WW (1963) *Biochim Biophys Acta* 67: 188-96

* **Allosteric proteins and cellular control systems.** Monod J, Changeux JP, Jacob F. (1963) *J Mol Biol* 6: 306-29

* **Kinetics of regulatory enzymes. kinetic order of the yeast diphosphopyridine nucleotide isocitrate dehydrogenase reaction and a model for the reaction.** Atkinson DE, Hathaway JA, Smith EC (1965) *J Biol Chem* 240: 2682-90.

* **Kinetics of regulatory enzymes: effectors for yeast phosphofructokinase do not alter the apparent kinetic order of the reaction.** Atkinson DE, Hathaway JA, Smith EC (1965) *Biochem Biophys Res Commun* 18:1-5

* **Biological feedback control at the molecular level.** Atkinson DE (1965) *Science* 150 : 851-857. Review

- * **Comparison of experimental binding data and theoretical models in proteins containing subunits.** Koshland DE , Nemethy G & Filmer D(1966) *Biochemistry* 5: 365-85
- * **Negative cooperativity in enzyme action. The binding of diphosphopyridine nucleotide to glyceraldehyde 3-phosphate dehydrogenase.** Conway A & Koshland DE (1968) *Biochemistry* 7: 4011-23
- * **Ligand induced half-of-the-sites reactivity in rabbit muscle glyceraldehyde-3-phosphate dehydrogenase.** Levitzki A (1973) *Biochem Biophys Res Commun* 54: 88-9.
- * **Letter: Kinetic negative co-operativity in the allosteric model of Monod, Wyman and Changeux.** Goldbeter A (1974) *J Mol Biol* 90: 185-90
- * **Letter: The physiological significance of negative co-operativity.** Cornish-Bowden A (1975) *J Theor Biol* 51: 233-5
- * **Co-operativity in monomeric enzymes.** Cornish Bowden A & Cárdenas ML (1987) *J Theor Biol* 124: 1-23
- * **Co-operative and allosteric enzymes: 20 years on.** Ricard J & Cornish Bowden A (1987) *Eur J Biochem* 166: 255-72
- * **Structure and control of pyridoxal phosphate dependent allosteric threonine deaminase.** Travis Gallagher D, Gilliland GL et al (1998) *Structure* 6:465-75
- * **Intramolecular signal transmission in enterobacterial aspartate transcarbamylases II. Engineering co-operativity and allosteric regulation in the aspartate transcarbamylase of *Erwinia herbicola*.** Cunin R, Rani CS, Van Vliet F, Wild JR & Wales M (1999) *J Mol Biol* 294: 1401-11
- * **Divergent allosteric patterns verify the regulatory paradigm for aspartate transcarbamylase.** Wales ME, Madison LL, Shannon SG & Wild JR (1999) *J Mol Biol* 294: 1387-400
- * **The structure of bovine glutamate dehydrogenase provides insights into the mechanism of allostery.** Petersen PE & Smith TJ (1999) *Structure* 7: 769-82
- * **Allosteric signal transmission involves synergy between discrete structural units of the regulatory subunit of aspartate transcarbamoylase.** Liu L, Wales ME & Wild JR (2000) *Arch Biochem Biophys* 373: 352-60
- * **Enzimología.** Núñez de Castro, I (2001) Ediciones Pirámide, Madrid, España
- * **Delineation of the allosteric mechanism of a cytidyltransferase exhibiting negative cooperativity.** Stevens SY, Sanker S, Kent C & Zuiderberg ER (2001) *Nat Struct Biol* 8: 947-52
- * **Direct structural evidence for a concerted allosteric transition in *Escherichia coli* aspartate transcarbamoylase.** Macol CP, Tsuruta H, Stec B & Kantrowitz ER (2001) *Nat Struct Biol* 8: 423-6
- * **Review: allostery in chaperonins.** Horovitz A, Fridmann Y, Kafri G, Yifrach O. *J Struct Biol.* (2001) 135(2):104-14. Review.
- * **Regulation of mammalian acetyl-CoA carboxylase.** Munday MR. *Biochem Soc Trans.* (2002) 30(Pt 6):1059-64. Review.
- * **Implications of enzyme kinetics.** McDonald AG. *Biochem Soc Trans.* (2003) 31(Pt 3):719-22. Review

- * **Reaction mechanism and regulation of cystathionine beta-synthase.** Banerjee R, Evande R, Kabil O, Ojha S, Taoka S. *Biochim Biophys Acta.* (2003) 1647(1-2):30-5. Review
- * **"Allosterism" in the elementary steps of the cytochrome P450 reaction cycle.** Yoon MY, Campbell AP, Atkins WM. *Drug Metab Rev.* (2004) 36(2):219-30. Review
- * **Implications of the allosteric kinetics of cytochrome P450s.** Atkins WM. *Drug Discov Today.* (2004) 9(11):478-84. Review.
- * **Characterization of noncompetitive regulators of proteasome activity.** Gaczynska M, Osmulski PA. *Methods Enzymol.* (2005) 398:425-38. Review
- * **Morpheins--a new structural paradigm for allosteric regulation.** Jaffe EK. *Trends Biochem Sci.* (2005) 30(9):490-7. Review
- * **Non-Michaelis-Menten kinetics in cytochrome P450-catalyzed reactions.** Atkins WM. *Annu Rev Pharmacol Toxicol.* (2005) 45:291-310. Review
- * **Current views on the fundamental mechanisms of cytochrome P450 allosterism.** Atkins WM et al. *Expert Opin Drug Metab Toxicol.* (2006) 2(4):573-9. Review
- * **Screening for positive allosteric modulators of biological targets.** Groebe DR. *Drug Discov Today.* (2006) 11(13-14):632-9. Review
- * **Application of a generalized MWC model for the mathematical simulation of metabolic pathways regulated by allosteric enzymes.** Najdi TS, Yang CR, Shapiro BE, Hatfield GW, Mjolsness ED. *J Bioinform Comput Biol.* (2006) Apr;4(2):335-55
- * **Ligand binding and allostery can emerge simultaneously.** Liang J, Kim JR, Boock JT, Mansell TJ, Ostermeier M. *Protein Sci.* (2007) May; 16(5):929-37
- * **Heterotropic cooperativity in oxidation mediated by cytochrome P450.** Niwa T, Murayama N, Yamazaki H. *Curr Drug Metab.* (2008) 9(5):453-62. Review
- * **Surface sites for engineering allosteric control in proteins.** Lee J, Natarajan M, Nashine VC, Socolich M, Vo T, Russ WP, Benkovic SJ, Ranganathan R. *Science.* (2008) 322(5900):438-42. doi: 10.1126/science.1159052.
- * **Allostery and cooperativity revisited.** Cui Q, Karplus M. *Protein Sci.* (2008) 17(8):1295-307
- * **Kinetics of allosteric activation.** Di Cera E. *Methods Enzymol.* (2009) 466:259-71. doi: 10.1016/S0076-6879(09)66011-0
- * **Jean-Pierre Changeux: allostery as neuro-enlightenment.** Changeux JP. *Mol Interv.* (2009) (2):61-7. doi: 10.1124/mi.9.2.2
- * **The apparent cooperativity of some GPCRs does not necessarily imply dimerization.** Chabre M, Deterre P, Antonny B. *Trends Pharmacol Sci.* (2009) 30(4):182-7. Review.
- * **Biochemistry. An ensemble view of allostery.** Hilser VJ. *Science.* (2010) 327(5966):653-4. doi: 10.1126/science.1186121.ç

- * **The folding cooperativity of a protein is controlled by its chain topology.** Shank EA, Cecconi C, Dill JW, Marqusee S, Bustamante C. *Nature*.(2010) 3;465(7298):637-40. doi: 10.1038/nature09021
- * **Scalable rule-based modelling of allosteric proteins and biochemical networks.** Julien F. Ollivier, Vahid Shahrezaei, and Peter S. Swain *PLoS Comput Biol*. (2010) 6(11): e1000975.
- * **Allostery in GPCRs: 'MWC' revisited.** Canals M, Sexton PM, Christopoulos A. *Trends Biochem Sci*. (2011) 36(12):663-72.
- * **Conformational selection or induced fit? 50 years of debate resolved.** Changeux JP, Edelstein S. *F1000 Biol Rep*. (2011) 3:19.
- * **50th anniversary of the word "allosteric".** Changeux JP. *Protein Sci*. (2011) 20(7):1119-24. doi: 10.1002/pro.658
- * **Conformational selection or induced fit? 50 years of debate resolved.** Changeux JP, Edelstein S. *F1000 Biol Rep*. (2011) 3:19. doi: 10.3410/B3-19
- * **Allostery and the Monod-Wyman-Changeux model after 50 years.** Changeux JP. *Annu Rev Biophys*. (2012) 41:103-33. doi: 10.1146/annurev-biophys-050511-102222
- * **The Monod-Wyman-Changeux allosteric model accounts for the quaternary transition dynamics in wild type and a recombinant mutant human hemoglobin.** Levantino M, Spilotros A, Cammarata M, Schirò G, Ardiccioni C, Vallone B, Brunori M, Cupane A. *Proc Natl Acad Sci U S A*. (2012) 109(37):14894-9.
- * **A Monod-Wyman-Changeux mechanism can explain G protein-coupled receptor (GPCR) allosteric modulation.** Canals M, Lane JR, Wen A, Scammells PJ, Sexton PM, Christopoulos A. *J Biol Chem*. (2012) 287(1):650-9.
- * **Hemoglobin Allostery: New Views on Old Players.** Miele AE, Bellelli A, Brunori M [J Mol Biol](#). (2012) pii: S0022-2836(12)00951-5. doi: 10.1016/j.jmb.2012.12.018
- * **Protein function and allostery: a dynamic relationship.** Kalodimos CG. *Ann N Y Acad Sci*. (2012) 1260:81-6.
- * **Structure and mechanisms of Escherichia coli aspartate transcarbamoylase.** Lipscomb WN, Kantrowitz ER. *Acc Chem Res*. (2012) 45(3):444-53. doi: 10.1021/ar200166p

Unidad Cáncer

- * **Introducción a la Oncología Molecular.** Gomez DE y Alonso DF. Ed Universidad nacional de Quilmas (1998)
- * **Bases De Oncología Molecular.** Bonfil D, Scharovsky G. Ed Dunken, Buenos Aires. (1998)
- * **Mechanisms in photodynamic therapy: Part one- photosensitizers, photochemistry and cellular localization.** Castano AP, Demidova TN, Hamblin MR. *Photodiagn Photodyn Ther* (2004) 1: 279-93

- * **Mechanisms of photodynamic therapy: Part two- cellular signaling, cell metabolism and modes of cell death.** Castano AP, Demidova TN, Hamblin MR. *Photodiagn Photodyn Ther* (2005) 2: 1-23
- * **Mechanisms of photodynamic therapy: Part three-photosensitizer pharmacokinetics, biodistribution, tumor localization and modes of tumor destruction.** Castano AP, Demidova TN, Hamblin MR. *Photodiagn Photodyn Ther* (2005) 2: 91-106.
- * **5-Aminolevulinic acid derivatives in photomedicine: Characteristics, application and perspectives.** Fotinos N, Campo MA, Popowycz F, Gurny R, Lange N. *Photochem Photobiol* (2006) 82 (4): 994-1015. Review
- * [Aminolevulinic acid photodynamic therapy for skin cancers.](#) Blume JE, Oseroff AR. *Dermatol Clin.* (2007) 25(1):5-14. Review.
- * **Inflammation in prostate carcinogenesis**
De Marzo AM, Platz EA, Sutcliffe S, Xu J, Grönberg H, Drake CG, Nakai Y, Isaacs WB, Nelson WG
Nature Reviews Cancer 7: 256-269 (2007) Review
- * **Monoclonal antibodies as therapeutic agents in oncology and antibody gene therapy. Review** °
Qi Zhang, Guihua Chen, Xinyuan Liu, Qijun Qian *Cell Research* (2007) 17: 89–99.
- * **Engineering targeted viral vectors for gene therapy**
R Waehler, SJ Russell, DT Curiel
Nature Reviews Genetics (2007) 8: 573-587. Review
- * **Immunosuppressive Strategies that are Mediated by Tumor Cells**
Gabriel A. Rabinovich, Dmitry Gabilovich & Eduardo M. Sotomayor. *Annu. Rev. Immunol.* (2007). 25, 267–296
- * [ALA and its clinical impact, from bench to bedside.](#) Krammer B, Plaetzer K. *Photochem Photobiol Sci.* (2008) 7(3): 283-9.
- * **The tumor microenvironment and its role in promoting tumor growth. TL Whiteside.** *Oncogene* (2008) 27, 5904–5912
- * **Advances in Chemical Carcinogenesis: A Historical Review and Prospective**
Lawrence A. Loeb & Curtis C. Harris. *Cancer Res* (2008) 68, 6863-6873
- * **Telomeres, Telomerase and Cancer (reprint).** Greider CW, Blackburn EH.
Scientific American (2009)
- * **MicroRNAs — the micro steering wheel of tumour metastases**
Milena S. Nicoloso, Riccardo Spizzo, Masayoshi Shimizu, Simona Rossi & George A. Calin. *Nat Rev Cancer* (2009)9, 293-302

* **Porphyrin and nonporphyrin photosensitizers in oncology: preclinical and clinical advances in photodynamic therapy.** O'Connor AE, Gallagher WM, Byrnie AT. *Photochem Photobiol Sci.* (2009): 85(4):934-42

* **Heme oxygenase 1 (HO-1) challenges the angiogenic switch in prostate cancer.** Ferrando M, Gueron G, Elguero B, Giudice J, Salles A, Leskow FC, Jares-Erijman EA, Colombo L, Meiss R, Navone N, De Siervi A, Vazquez E. *Angiogenesis.* (2011) 14(4):467-79.

* **Key questions in metastasis: new insights in molecular pathways and therapeutic implications.** Gueron G, De Siervi A, Vazquez E. *Curr Pharm Biotechnol.* (2011) 12(11):1867-80. Review.

* **Telomerase Targeted Therapy in Cancer and Cancer Stem Cells.** Yucheng Xu, PhD, Kaijie He, BS, and Amir Goldkorn, MD. *Clinical Advances in Hematology & Oncology.* (2011) 9(6)

* **State-of-the-art gene-based therapies: the road ahead.** MA Kay. *Nature Reviews Genetics* (2011) 12: 316-328

* **Advanced prostate cancer: reinforcing the strings between inflammation and the metastatic behavior.** Gueron G, De Siervi A, Vazquez E. *Prostate Cancer Prostatic Dis.* (2012);15(3):213-21. Review.