

LECTURE #1: NUCLEOCYTOPLASMIC TRAFFICKING

Assigned (Required) Reading is (1)-(4):

•General Background (in your Reader):

(1) Tran EJ, Wentz SR (2006) Dynamic nuclear pore complexes: life on the edge. *Cell* 125: 1041-1053.

(2) Lange A, Mills RE, Lange CJ, Stewart M, Devine SE, Corbett AH (2007) Classical nuclear localization signals: definition, function, and interaction with importin- α . *J. Biol. Chem.* 282: 5101-5105.

(3) Cole CN, Scarcelli JJ (2006) Unravelling mRNA export. *Nature Cell Biol.* 8: 645-647.

•Paper for Friday Discussion Session (9 Nov.):

(4) Frey S, Görlich D (2007) A saturated FG-repeat hydrogel can reproduce the permeability properties of nuclear pore complexes. *Cell* 130: 512-523.

Nuclear Envelope and Nuclear Pore:

Stavru F, Nautrup-Pedersen G, Cordes VC, Görlich D (2006) Nuclear pore complex assembly and maintenance in POM121- and gp210-deficient cells. *J Cell Biol.* 173: 477-483.

Stavru F, Hulsmann BB, Spang A, Hartmann E, Cordes VC, Görlich D (2006) NDC1: a crucial membrane-integral nucleoporin of metazoan nuclear pore complexes. *J Cell Biol.* 173: 509-519.

Hetzer MW, Walther TC, Mattaj IW (2005) Pushing the envelope: structure, function, and dynamics of the nuclear periphery. *Annu. Rev. Cell Dev. Biol.* 21: 347-380.

Fahrenkrog B, Koser J, Aebi U (2004) The nuclear pore complex: a jack of all trades? *Trends Biochem. Sci.* 29: 175-182.

Zeitler B, Weis K (2004) The FG-repeat asymmetry of the nuclear pore complex is dispensable for bulk nucleocytoplasmic transport in vivo. *J. Cell Biol.* 167: 583-590.

Strawn LA, Shen T, Shulga N, Goldfarb DS, Wentz SR (2004) Minimal nuclear pore complexes define FG repeat domains essential for transport. *Nature Cell Biol.* 6: 197-206.

Suntharalingam M, Wentz SR (2003) Peering through the pore: nuclear pore complex structure, assembly, and function. *Dev. Cell* 4: 775-789.

Schirmer EC, Gerace L (2002) Organellar proteomics: the prizes and pitfalls of opening the nuclear envelope. *Genome Biol.* 3: Reviews1008.1-1008.4.

Ribbeck K, Görlich D (2002) The permeability barrier of nuclear pore complexes appears to operate via hydrophobic exclusion. *EMBO J.* 21: 2664-2671.

Paschal BM (2002) Translocation through the nuclear pore complex. *Trends Biochem. Sci.* 27: 593-596.

Aitchison JD, Rout MP (2002) A tense time for the nuclear envelope. *Cell* 108: 301-304.

Lyman SK, Gerace L (2001) Nuclear pore complexes: dynamics in unexpected places. *J. Cell Biol.* 154: 17-20.

Rout MP, Aitchison JD (2001) The nuclear pore complex as a transport machine. *J. Biol. Chem.* 276: 16593-16596.

Adam SA (2001) The nuclear pore complex. *Genome Biol.* 2: Revs. 0007.1-0007.8

- Vasu SK, Forbes DJ (2001) Nuclear pores and nuclear assembly. *Curr. Opin. Cell Biol.* 13: 363-375.
- Stewart M, Baker RP, Bayliss R, Clayton L, Grant RP, Littlewood T, Matsuura Y (2001) Molecular mechanism of translocation through nuclear pore complexes during nuclear protein import. *FEBS Lett.* 498: 145-149.
- Ribbeck K, Görlich D (2001) Kinetic analysis of translocation through nuclear pore complexes. *EMBO J.* 20: 1320-1330.
- Rout, M.P., J.D. Aitchison, A. Suprpto, K. Hjertaas, Y. Zhao and B.T. Chait (2000) The yeast nuclear pore complex: composition, architecture, and transport mechanism. *J. Cell Biol.* 148: 635-651.
- Wente, S.R. (2000) Gatekeepers of the nucleus. *Science* 288: 1374-1377.
- Allen TD, Cronshaw JM, Bagley S, Kiseleva E, Goldberg MW (2000) The nuclear pore complex: mediator of translocation between nucleus and cytoplasm. *J. Cell Sci.* 113: 1651-1659.
- Wilson KL (2000) The nuclear envelope, muscular dystrophy and gene expression. *Trends Cell Biol* 10: 125-129.
- Signals for Nucleocytoplasmic Trafficking:
- Erkmann JA, Sanchez R, Treichel N, Marzluff WF, Kutay U (2005) Nuclear export of metazoan replication-dependent histone mRNAs is dependent on RNA length and is mediated by TAP. *RNA* 11: 45-58.
- Kau TR, Way JC, Silver PA (2004) Nuclear transport and cancer: from mechanism to intervention. *Nature Rev. Cancer* 4: 106-117.
- Weis K (2003) Regulating access to the genome: nucleocytoplasmic transport throughout the cell cycle. *Cell* 112: 441-451.
- Cartier R, Reszka R (2002) Utilization of synthetic peptides containing nuclear localization signals for nonviral gene transfer systems. *Gene Ther.* 9: 157-167.
- Gallouzi IE, Steitz JA (2001) Delineation of mRNA export pathways by the use of cell-permeable peptides. *Science* 294(1): 1895-1901.
- Jans DA, Xiao CY, Lam MH (2000) Nuclear targeting signal recognition: a key control point in nuclear transport? *Bioessays* 22: 532-544. Review.
- Michael WM (2000) Nucleocytoplasmic shuttling signals: two for the price of one. *Trends Cell Biol.* 10: 46-50.

The Ran GTPase Cycle:

- Seewald MJ, Kraemer A, Farkasovsky M, Korner C, Wittinghofer A, Vetter IR (2003) Biochemical characterization of the Ran-RanBP1-RanGAP system: are RanBP proteins and the acidic tail of RanGAP required for the Ran-RanGAP GTPase reaction? *Mol. Cell. Biol.* 23: 8124-8136.
- Dasso M (2002) The Ran GTPase: theme and variations. *Curr Biol.* 12: R502-R508.
- Bischoff FR, Scheffzek K, Ponstingl H (2002) How Ran is regulated. *Results Probl Cell Differ.* 35: 49-66.
- Künzler M, Hurt E (2001) Targeting of Ran: variation on a common theme? *J Cell Sci.* 114: 3233-3241.
- Moore JD (2001) The Ran GTPase and cell-cycle control. *Bioessays* 23: 77-85.

Azuma Y, Dasso M. (2000) The role of Ran in nuclear function. *Curr Opin Cell Biol.* 12: 302-307.

Nishimoto T (2000) Upstream and downstream of Ran GTPase. *Biol Chem* 381: 397-405.

Overviews of the Biochemical Basis for Nucleocytoplasmic Transport:

Frey S, Richter RP, Gorlich D (2006) FG-rich repeats of nuclear pore proteins form a three-dimensional meshwork with hydrogel-like properties. *Science* 314: 815-817.

Becskei A, Mattaj IW (2005) Quantitative models of nuclear transport. *Curr. Opin. Cell Biol.* 17: 27-34.

Kuersten S, Ohno M, Mattaj IW (2001) Nucleocytoplasmic transport: Ran, *beta* and beyond. *Trends Cell Biol.* 11: 497-503.

Quimby BB, Corbett AH (2001) Nuclear transport mechanisms. *Cell Mol Life Sci.* 58: 1766-1773.

Macara IG (2001) Transport in and out of the nucleus. *Microbiol. Mol. Bio. Rev.* 65, in press.

Chook YM, Blobel G (2001) Karyopherins and nuclear import. *Curr. Opin. Struct. Biol.* 11: 703-715.

Strom AC, Weis K (2001) Importin-beta-like nuclear transport receptors. *Genome Biol.* 2: Revs. 3008.1-3008.9.

Export of Proteins and RNA from the Nucleus:

Behm-Ansmant I, Izaurralde E (2006) Quality control of gene expression: a stepwise assembly pathway for the surveillance complex that triggers nonsense-mediated mRNA decay. *Genes Dev.* 20: 391-398.

Fasken MB, Corbett AH (2005) Process or perish: quality control in mRNA biogenesis. *Nature Struct. Mol. Biol.* 12: 482-488.

Conti E, Izaurralde E. (2005) Nonsense-mediated mRNA decay: molecular insights and mechanistic variations across species. *Curr. Opin. Cell Biol.* 17: 316-325.

Erkman JA, Kutay U (2004) Nuclear export of mRNA: from the site of transcription to the cytoplasm. *Exp. Cell Res.* 296: 12-20.

Bohnsack MT, Czaplinski K, Görlich D (2004) Exportin 5 is a RanGTP-dependent dsRNA-binding protein that mediates nuclear export of pre-miRNAs. *RNA* 10: 185-191.

Jensen TH, Dower K, Libri D, Rosbash M (2003) Early formation of mRNP: license for export or quality control? *Mol Cell* 11: 1129-1138.

Maniatis T, Reed R (2002) An extensive network of coupling among gene expression machines. *Nature* 416: 499-506.

Reed R, Hurt E (2002) A conserved mRNA export machinery coupled to pre-mRNA splicing. *Cell* 108: 523-531.

Izaurralde E (2002) A novel family of nuclear transport receptors mediates the export of messenger RNA to the cytoplasm. *Eur. J. Cell Biol.* 81: 577-584.

Lei EP, Silver PA (2002) Protein and RNA export from the nucleus. *Dev Cell.* 2: 261-272.

Reed R, Hurt E (2002) A conserved mRNA export machinery coupled to pre-mRNA splicing. *Cell* 108: 523-531.

Zenklusen D, Stutz F (2001) Nuclear export of mRNA. *FEBS Lett.* 498: 150-156.

- Moore MJ, Rosbash M (2001) Cell biology. TAPping into mRNA export. *Science* 294: 1841-1842.
- Kruse C, Hartmann RK, Muller PK (2001) Nuclear-cytoplasmic translocation of tRNA. *Exp Cell Res*. 262: 3-7.
- Grosshans H, Simos G, Hurt E. (2000) Transport of tRNA out of the nucleus-direct channeling to the ribosome? *J. Struct. Biol.* 129: 288-294.
- Cole CN (2000) mRNA export: the long and winding road. *Nature Cell Biol.* 2: E55-E58.
- Daneholt B. (1999) Pre-mRNP particles: From gene to nuclear pore. *Curr Biol.* 9: R412-415.
- Structural Analysis at Atomic Resolution of Nuclear Transport Factors:
- Devos D, Dokudovskaya S, Williams R, Alber F, Eswar N, Chait BT, Rout MP, Sali A. (2006) Simple fold composition and modular architecture of the nuclear pore complex. *Proc. Natl. Acad. Sci. USA* 103: 2172-2177.
- Weirich CS, Erzberger JP, Berger JM, Weis K. (2004) The N-terminal domain of Nup159 forms a beta-propeller that functions in mRNA export by tethering the helicase Dbp5 to the nuclear pore. *Mol. Cell* 16: 749-760.
- Conti E (2002) Structures of importins. *Results Probl Cell Differ* 35: 93-113.
- Peters R, Coutavas E, Siebrasse JP (2002) Nuclear transport kinetics in microarrays of nuclear envelope patches. *J. Struct. Biol.* 140: 268-278.
- Conti E, Izaurralde E (2001) Nucleocytoplasmic transport enters the atomic age. *Curr. Opin. Cell Biol.* 13: 310-319.
- Bayliss R, Littlewood T, Stewart M (2000) Structural basis for the interaction between FxFG nucleoporin repeats and importin-beta in nuclear trafficking. *Cell* 102: 99-108.
- Mattaj IW, Conti E (1999) Cell biology. Snail mail to the nucleus. *Nature* 399: 208-210.
- Chook YM, Cingolani G, Conti E, Stewart M, Vetter I, Wittinghofer A. (1999) Pictures in cell biology. Structures of nuclear-transport components. *Trends Cell Biol.* 9: 310-311.
- Macara IG. (1999) Nuclear transport: Ran-dy couples. *Curr Biol.* 9: R436-R439.
- Regulation of Nucleocytoplasmic Trafficking by Protein Phosphorylation:
- Schuller C, Ruis H (2002) Regulated nuclear transport. *Results Probl Cell Differ.* 35: 169-189.
- Komeili A, O'Shea EK (2001) New perspectives on nuclear transport. *Annu Rev Genet* 35: 341-364.
- Sweitzer TD, Love DC, Hanover JA. (2000) Regulation of nuclear import and export. *Curr Top Cell Regul.* 36: 77-94.
- Zhu J, McKeon F (2000) Nucleocytoplasmic shuttling and the control of NF-AT signaling. *Cell Mol Life Sci* 57: 411-420.
- Stein GS, van Wijnen AJ, Stein JL, Lian JB, Montecino M, Choi J, Zaidi K, Javed A. (2000) Intranuclear trafficking of transcription factors: implications for biological control. *J Cell Sci.* 113: 2527-2533. Review.
- Ferrigno P, Silver PA. (1999) Regulated nuclear localization of stress-responsive factors: how the nuclear trafficking of protein kinases and transcription factors contributes to cell survival. *Oncogene* 18: 6129-6134.

Turpin P, Ossareh-Nazari B, Dargemont C. (1999) Nuclear transport and transcriptional regulation. *FEBS Lett.* 452: 82-86.

Reiser V, Ammerer G, Ruis H (1999) Nucleocytoplasmic traffic of MAP kinases. *Gene Expr.* 7: 247-254.

Wang H, Clapham DE (1999) Calcium-dependent conformational changes of the in situ nuclear pore complex. *Biophys J.* 77: 241-247.

-