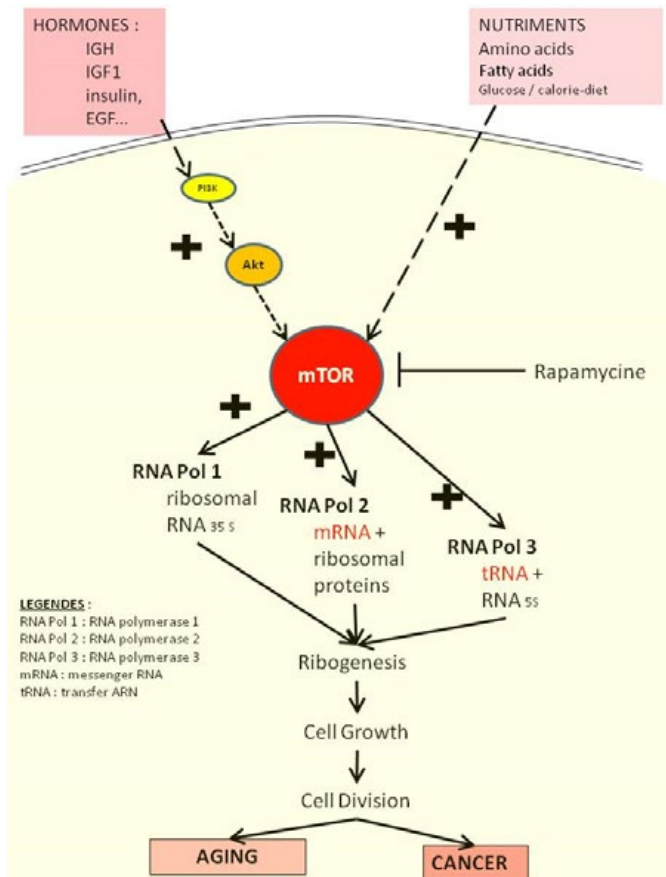


mTOR

masterchief of cell growth, mitosis and aging

Determination of Urinary Modified Nucleosides give us informations the about mTOR profile of the patient.

mTOR (mammalian Target of Rapamycin) is a cytoplasmic kinase which integrates hormonal and nutria stimuli and traduces them in cell growth. mTOR, once activated, induces the constitution and activity of protein synthesis industry of the cell.



Except the period from birth to adult age, growth is not profitable to cells and organisms physiology and longevity, and this for several reasons:

1. Growth of cytoplasmic mass induced transition from G0/G1 phase which the period of prosperity and longevity of the cell, to S phase of duplication of chromosomic material.

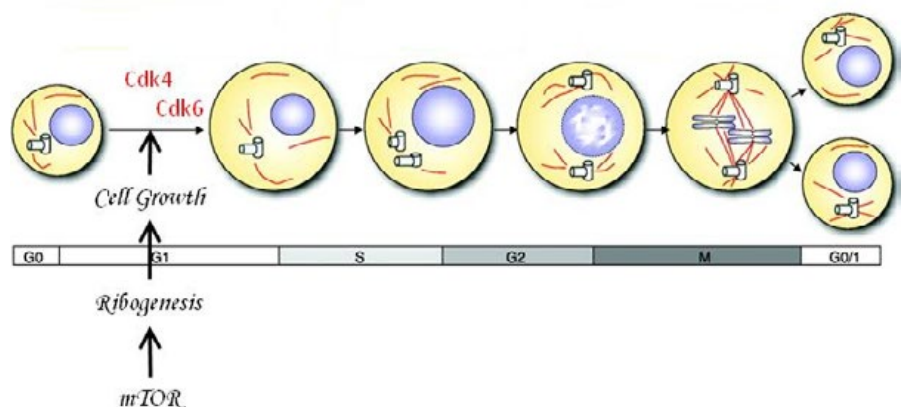
2. Thus, because Mitosis is systematically coupled to cytoplasmic mass expansion, growth leads to shortening telomeres, reduction of replication potential and accelerates replicative senescence.

3. Excessive and uncontrolled growth exhausts our niches of stem cells by accelerating their maturation and leading them to apoptosis before they can express their specific regenerative activity.

4. Cell growth is the hallmark of growth hormone and factors hGH, IGF1/2, insulin, that in addition promote inflammation, oxidative stress, metabolic overload, which in turn enhance proliferative mechanism.

5. In aging cells and organisms prone to cancerous diseases, high degree of growth may contribute to their development.

mTOR induces G0/G1 -> S transition through their activation of protein synthesis and therefore cytoplasmic mass expansion which promote the expression of CDK4 & 6. CDK 4 & 6 engage the cell in the S phase.



Sources:

- Marcos Malumbres, Physiological Relevance of Cell Cycle Kinases, Physiological Reviews July 2011 Vol. 91 no. 3, 973-1007
- Stefan M. Schieke, Toren Finkel, TOR and Aging: Less Is More, Cell Metabolism, Vol. 5, Issue 4, p233-235
- Kapahi P, Zid BM, Harper T, Koslover D, Sapin V, Benzer S. Regulation of lifespan in Drosophila by modulation of genes in the TOR signaling pathway. Curr Biol. 2004 May 25;14(10):885-90.
- Kaeblerlein M, Kennedy BK. Hot topics in aging research: protein translation and TOR signaling, 2010. Aging Cell. 2011 Apr;10(2):185-90.
- Jeng LB, Lo WY, Hsu WY, Lin WD, Lin CT, Lai CC, Tsai FJ, Analysis of urinary nucleosides as helper tumor markers in hepatocellular carcinoma

- diagnosis. Rapid Commun Mass Spectrom. 2009 Jun;23(11):1543-9.
- Jeng LB, Lo WY, Hsu WY, Lin WD, Lin CT, Lai CC, Tsai FJ. Analysis of urinary nucleosides as helper tumor markers in hepatocellular carcinoma diagnosis. Rapid Commun Mass Spectrom. 2009 Jun;23(11):1543-9
- Li S, Jin Y, Tang Z, Lin S, Liu H, Jiang Y, Cai Z. A novel method of liquid chromatography-tandem mass spectrometry combined with chemical derivatization for the determination of ribonucleosides in urine. Anal Chim Acta. 2015 Mar 15;864:30-8.
- Zhang YR, Shi L, Wu H, Tang DD, Wang SM, Liu HM, Zhang LR, Song DK. Urinary modified nucleosides as novel biomarkers for diagnosis

- and prognostic monitoring of urothelial bladder cancer. Tumori. 2014 Nov-Dec;100(6):660-6.
- Borland KM, AbdulSalam SF, Solivio MJ, Burke MP, Wolfkiel PR, Lawson SM, Stockman CA, Andersen JM, Smith S, Tolstolutskaia JN, Gurjar PN, Bercz AP, Merino EJ, Litosh VA. Base-modified thymidines capable of terminating DNA synthesis are novel bioactive compounds with activity in cancer cells. Bioorg Med Chem. 2015 Apr 15;23(8):1869-81.
- Rita Horvath, Patrick F. Chinnery, Modifying Mitochondrial tRNAs: Delivering What the Cell Need, Cell Metabolism, March 2015 Volume 21, Issue 3, p351-352



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M. MTOR

Dossier n° B170330001
Résultats de M. MTOR
Né(e) le 01/01/1970 (47 ans)

MAIN PROPRE

Dossier enregistré le : 30/03/2017 à 09:48

Compte-rendu complet
Edition le Vendredi 31 Mars 2017 à 10:41

OXIDATIVE STRESS

Urinary modified nucleosides

LC-MS/MS -SHIMADZU -deuterated standards

Nucleosides en µmoles/gr Creatinin except m6A et m7G in nanomoles / gr Creatinin

↗ Pseudouridine	239	µM/gCr	(130-220)
Methyl-1-guanosine	9.0	µM/gCr	(6.5-13.5)
Methyl-2-guanosine	9.2	µM/gCr	(4.7-9.3)
↑ Methyl-2-O-guanosine	3.52	µM/gCr	(0.30-1.70)
Di-methyl-2-guanosine	14.4	µM/gCr	(8.0-20.0)
↓ Tri-methyl-2-guanosine	0.71	µM/gCr	(0.80-1.20)
Methyl-7-guanosine	140.9	nM/gCr	(0.0-400.0)
↗ Methyl-1-adenosine	37.2	µM/gCr	(16.0-36.0)
Methyl-6-adenosine	11	nM/gCr	(0-36)
Acetyl-4-cytidine	4.2	µM/gCr	(2.0-6.0)
Methyl-thio-adenosine	0.77	µM/gCr	(0.30-1.70)
↑ Sum of nucleosides	454	µM/gCr	(170-300)

Mildly increased "whole body" proliferative index.
Mildly increased protein synthesis / ribogenesis rate.
Mildly increased mTOR activity.

Urinary Creatinin
Urate/Creatratio

1000 mg/l (1 200-2 000)
0.42-0.7