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**PROSTATIC CANCER OF VIRAL ORIGIN: HOMOLOGU OF HUMAN ONCOGENIC PAPILLOMAVIRUS (HPV) L1 WITH NUCLEOPHOSMIN (NPM1), A CONTROLLER OF ANDROGEN RECEPTOR TRANSCRIPTION**

**Association de la Recherche contre les Tumeurs de la Prostate ARTP 2014, 19 November, Paris**

**Background:** We showed previously that HPV contained **Prostatic Cancer (PC)** related oncogenic proteins:

1°) HPV **E2** (51-112) is homologous to an **Epidermal Growth Factor** (*Tran MKG, 1997*).

2°) **E1** to **PTEN**, **E6** and **L1** to the c-Myc inhibitor **Bin-1** (**Bridging integrator 1** or amphiphysin II), a tumor suppressor deleted in 42% of PC (*Tran GMK, 2008*).

3°) HPV-18 **E2** mimics **Osteoprotegerin** and **ParaTHormone related Protein** (PTHrP) active

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site (explaining bone metastasis). Anwar K (1992) found 80% HPV-18 in metastatic PC in Japan. Our meta-analysis concluded to a frequency of about 30%-40% (21-80%) oncogenic HPV (-16, -18, -33) in PC

(*EuroConf Cancer Pasteur Inst, 2004*).

The most important point is the

### **PCR (E6 primer and fresh tissue**

); L1 primer and formalin-fixed, paraffin-embedded yielded negative results. For example, Terris MK (1997) obtained 21% positivity with E6 and 0% with L1 primer, in the

### **same**

patients. Recent results confirm that use of paraffin-embedded tissues

(*Groom HCT, 2012; Ghasemian E, 2013*)

or L1 primer

(*Sylvestre RV, 2009*)

or both (13/104 versus 8/104

*Aghakhani A, 2011*)

were unsuccessful. Positive results were reported (in 10.5%

*Jalilvand S, 2014;*

worse overall survival,

*Pascale M, 2013*).

Noda S (1975) described **papillomavirus-like particles** in electron microscopy of prostate cancer tissue. Whitaker NJ (2013) found

### **koilocytes**

in HPV-18 infected prostate cancer.

Our **aim** is to link HPV to Androgen Receptor (AR). Another hormonal cancer linked to virus is breast cancer, as the virus integration site is Aromatase, the estrogen synthesizing enzyme (*Tekmal RR, 1995*)

**Methods:** Amino Acid (AA) sequence comparison between HPV (*Lowe J, 2008*) and **NPM1 (nucleophosmin)**

, which controls

### **AR**

transcriptional activity by promoting S-phase entry and hyperproliferation (cyclin switch D1 to E1 and p27kip1 loss)

(*Boudra R, ARTP 2013*)

. Clinically, high

### **p27kip1**

is a correlate of better survival after prostatectomy at 5 years.

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**Results:** HPV L1 chimera (types 16, 18, 31, 33, 44, 56, 66, 115) [type-16, AA 167-219] is homologous to NPM1 chimera (human, duck, alligator, sheep, rhinoceros, turtle,...)[AA 1-48]

**NPM1 MEDSMDMDSMQPLRPQMFLFGC- - - SGAHWARISPCSLGFFAGCELKSD**

**HPV L1 VEDSMDV – SMDPKQIQMFLI GCKPPTGEHWAR-SPCSPVG- --AGDCELKSD**

**Conclusion:** Anti-androgen escape may be explained by AR mutations, but also in a PC subset (about perhaps 30%-40%, depending on the number of HPV serotypes screened) by a viral infection (oncogenic HPV), as HPV L1 is a viral NPM1 mimetic, enhancing AR transcriptional activity and inducing lethal p27kip1 loss. Japanese mushroom **Shii**

**take**

is a non toxic and highly efficient anti-HPV

(*Smith*

*JA,*

*2014)*

**Indole 3 carbinol**

from

**cruciferous vegetables**

(Brussels sprouts, broccoli)

are efficient against HPV-16 by viral transcription inhibition

(*Bradlow HL, 1999; Rieck GC, 2006*).

Anti-cancer drugs discovered by

**HPV-18**

infected KB cells screening

(*Perdue RE Jr, 1982*)

may act,

**by**

**serenpidity**

**, as anti-HPV**

: Taxol (Paclitaxel, Docetaxel, Cabazitaxel), topotecan, Vinca Alkaloids (Vinorelbine, Vinflunine)

**HPV vaccination**

of young men could protect against PC.

**Bibliography Aghakhani A.** *Scand J Infect Dis* 2011, 43: 64-9. **Anwar K** et al. Presence of ras oncogene mutations and human papillomavirus DNA in human prostate carcinomas.

*Cancer Res*

1992, 52: 5991-6.

**Boudra R**

et al. Nucleophosmin overexpression down-regulates p27Kip1 and induces hyperproliferation in the prostate gland of transgenic mice. ARTP 2013.

**Bradlow HL**

. Multifunctional aspects of the action of indole-3-carbinol as an antitumor agent.

*Ann N Y Acad Sci*

1999, 889: 204-13. Review.

**De Villiers EM**

.

*Breast Cancer Res*

2005, 7: R1-R11

**Ghasemian E.**

*Asian Pacific J*

*Cancer Prev*

14: 3305-8

**Groom HCT.**

*PLoS ONE*

2012, 7(3): e34221.

**Jalilvand S**

.

*Asian*

*Pac J Cancer Prev*

15 (17), 7029-35

**Lowe J**

et al. Evolutionary and structural analyses of alpha-papillomavirus capsid proteins yields novel insights into L2 structure and interaction with L1.

*Virology*

2008, 5: 150.

**Noda S.**

*The Kurume Med J*

1975, 22: 261-8

**Pascale M**

*Disease Markers*

2013, 35: 607-13

**Perdue RE Jr**

. KB cell culture I. Role in discovery of antitumor agents from higher plants.

*J Nat Prod*

1982, 45: 418-26.

**Rieck GC**

*Mol Nutr Food Res*

2006, 52: 105-13.

**Smith JA**

et al. Evaluation of active hexose correlated compound (AHCC) for the eradication of HPV infections in women with HPV positive Pap smears.

*Int Conf Soc Integrative Oncology*

, Houston, Oct 28, 2014.

**Tekmal RR**

, Durgam VR. The overexpression of int-5/Aromatase, a novel MMTV integration locus gene, is responsible for D2 mammary tumor cell proliferation.

*Cancer Lett*

1995, 88: 147-55.

**Terris MK**

*Urology* 1997, 50: 150-6.

**Tran GMK**

et al. Role of human papillomavirus type 18 in a subgroup of prostatic cancer with bone metastases: Its protein E2 contains the osteoprotegerin active site.

*EuroConf. Cancer, Pasteur Institute,*

Jan 15-16 2004, Paris.

(Free on: [Positifs.org](http://Positifs.org) C.53)

**Tran MKG**

et al. Human papillomavirus (HPV) E2 protein contains a chimera of epidermal growth factor (EGF) and EGF family mitogens: Heregulin and tumor growth factor.

*6th Europ Conf Clin Aspects Treatment HIV*

*infection*

, Hamburg, Germany, Oct 11-15,1997: P474.

**Tran GMK**

et al. Cancer de la prostate métastatique: Le suppresseur de tumeur Bin1, inhibiteur de c-Myc, est homologue aux protéines E6 et L1 des papillomavirus humains oncogènes et PTEN à la protéine E1.

*Bull Cancer*

2008, 95: 592. P33.

**Whitaker NJ.**

*Prostate*

2013, 73: 236-41

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