

A CASE OF LOCALIZED, HIGH-RISK PROSTATE CANCER

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CLINICAL CASE: DIAGNOSIS & STAGING

BG, 71 years, Professor of Ingeneering, PS=0

Mild hypertension

Thyroidectomy for goiter

Benign prostatic hyperplasia (PSA around 3.5)

March 2017

PSA 4.59 (F/T 25.7%)

DRE: enlarged prostate, new hardness in the left lobe

What's next?

EVIDENCE FROM LITERATURE: MR of prostate

Is MR indicated before prostate biopsy?

EUROPEAN UNDIOGT 73 (2018) 23-30

available at www.sciencedirect.com journal homepage: www.europeanurology.com





Platinum Priority - Prostate Cancer
Edizated by Jacken Walt on an 31-42 of thir pseu-

Optimising the Diagnosis of Prostate Cancer in the Era of Multiparametric Magnetic Resonance Imaging: A Cost-effectiveness Analysis Based on the Prostate MR Imaging Study (PROMIS)

Rita Faria ^{a.*}, Marta O. Soares ^a, Eldon Spackman ^b, Hashim U. Ahmed ^{c.a}, Louise C. Brown ^{d.*} Richard Kaplan ^a, Mark Emberton ^{e.f}, Mark J. Sculpher ^a

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Oncology

Interpreting Prostate Multiparametric Magnetic Resonance Imaging: Urologists' Guide Including Prostate Imaging Reporting and Data System

Daniel Christidis, Shannon McGrath, Barry Leaney, Richard O'Sullivan, and Nathan Lawrentschuk

OBJECTIVE To review and explain the development of multiparametric MIS and its use in purstate cancer diagnosis while colorating on the implication of certain real/cological findings.
METHODS

The physics of magnetic reconnuct assigning is reviewed befor the explanation of different phase technologies in "multiparametric" scanning. Sample images of the postate are used to display theorems and excluding the properties of the postate are used to display theorems and excluding the properties.

Modalities of multiparametric magnetic resonance imaging (mpMRI) of the prostate were reviewed and the interpretation of certain findings were displayed on sample images to educate clinicism desert their measures and significant.

CONCLUSION Diagnosis, biopsy targeting, surveillance, operative planning and staging has led to endansement of mpMRI and it is imperative that treating undoposts have an understanding of mpMRI to appreciate the power and infinitacism of its indiagn. URCLOY III.116-1182, 2018. 2023 [7] Elsevier

Should MR-guided biopsy become the new standard?

Radiology

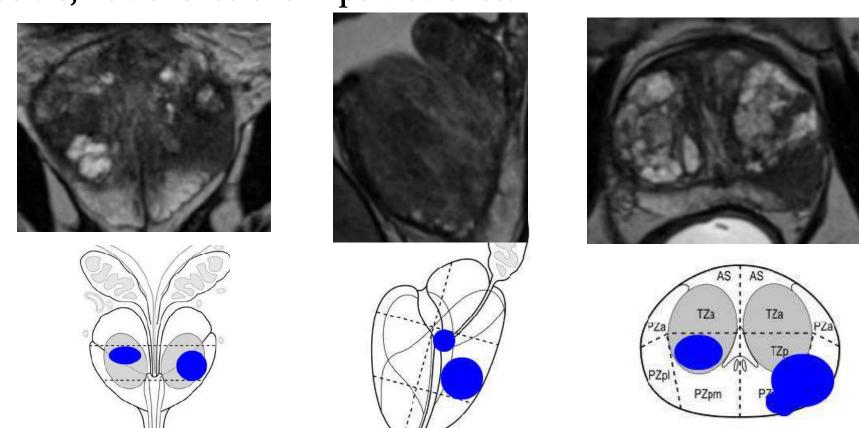
Sadhna Verma, MD Peter L. Choyke, MD Steven C. Eberhardt, MD Aytekin Oto, MD Care M. Tempany, MD Baris Turkbey, MD Andrew B. Rosenkrantz, MD

The Current State of MR Imaging-targeted Biopsy Techniques for Detection of Prostate Cancer¹

Systematic transrectal ultrasonography (US)-guided biopsy is the standard approach for histopathologic diagnosis of prostate cancer. However, this technique has multiple limitations because of its inability to accurately visualize and target prostate lesions. Multiparametric magnetic resonance (MR) imaging of the prostate is more reliably able to localize significant prostate cancer. Targeted prostate biopsy by using MR imaging may thus help to reduce false-negative results and improve risk assessment. Several commercial devices are now available for targeted prostate biopsy, including in-gantry MR imaging-targeted biopsy and real-time transrectal US-MR imaging fusion biopsy systems. This article reviews the current status of MR imaging-targeted biopsy platforms, including technical considerations, as well as advantages and challenges of each technique.

CLINICAL CASE: MR of the Prostate

April 2017: Multiparametric MR: left lateral-posterior nodule 24 mm with capsule involvement (P-RADS 5) + median nodule 19 mm (PI-RADS 3). No suspicious lymph-nodes in pelvis, no bone lesions in pelvic bones.

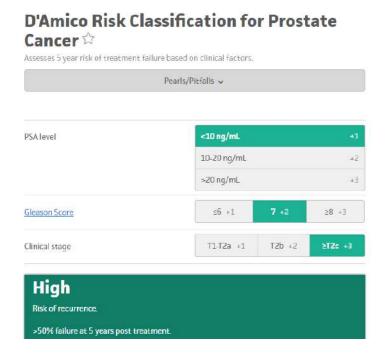


CLINICAL CASE: Prostate biopsy

1st June 2017: ultrasound-guided 12 core biopsies.

Three cores show adenocarcinoma Gleason 4+3

- -paramedian left lobe
- -paramedian right lobe
- -transitional zone



What further staging?

EVIDENCE FROM LITERATURE: staging

Are CT scan + bone scintigraphy



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indicated?

High-risk localised PCa/High-risk locally advanced PCa			
Use prostate mpMRI for local staging.	2b	Α	
Perform metastatic screening including at least cross-sectional abdominopelvic imaging and a bone-scan.	2a	Α	

EUROPEAN UROLOGY 70 (2016) 161-175

Should PET/TAC become the new standard for staging?

Choline or PSMA?

available at www.sciencedirect.com journal homepage: www.europeanurology.com





Collaborative Review - Prostate Cancer

New Clinical Indications for 18F/11C-choline, New Tracers for Positron Emission Tomography and a Promising Hybrid Device for Prostate Cancer Staging: A Systematic Review of the Literature

Laura Evangelista a.*, Alberto Briganti b, Stefano Fanti c, Stephen Joniau d, Sven Reske e, Riccardo Schiavina^f, Christian Stief^g, George N. Thalmann h, Maria Picchio

* Radiotherapy and Nuclear Medicine Unit, Veneto Institute of Oncology IOV-IRCCS, Padua, Italy; b Department of Urology, Vita-Salute University San Raffaele, Milan, Italy; Service of Nuclear Medicine, Policlinico S. Orsola-Malpighi, University of Bologna, Bologna, Italy; Department of Urology, University Hospital, Leuven, Belgium; eKlinik für Nuklearmedizin, Universität Ulm, Ulm, Germany; Department of Urology, University of Bologna, S. Orsola-Malpighi Hospital, Italy: 8 Department of Urology, Ludwig-Maximilians-Universität Munich, Germany; h Department of Urology, University of Bern, Inselspital, Bern, Switzerland; Nuclear Medicine Department, IRCCS San Raffaele Scientific Institute, Milan, Italy

EVIDENCE FROM LITERATURE: neoadjuvant therapy

Neo-adjuvant Endocrine therapy



TUMOUR REVIEW

A systematic review and meta-analysis of randomised trials of neo-adjuvant hormone therapy for localised and locally advanced prostate carcinoma

M.D. Shelley ^{a.*}, S. Kumar ^{b.f}, T. Wilt ^{c.g}, J. Staffurth ^{d.h}, B. Coles ^{e.j}, M.D. Mason ^{d.i}

Waiting for data with abiraterone and enzalutamide

Neo-adjuvant Docetaxel

Review Article

Neoadjuvant Chemotherapy prior to Radical Prostatectomy for Patients with High-Risk Prostate Cancer: A Systematic Review

Stavros Sfoungaristos, Vasileios Kourmpetis, Eleftherios Fokaefs, and Petros Perimenis

Department of Urology, Patras University Hospital, 26500 Patras, Greece

Correspondence should be addressed to Stavros Sfoungaristos; sfoungaristosst@gmail.com

Received 23 December 2012; Accepted 22 January 2013

Lancet Oncol 2015; 16: 787-94

Androgen deprivation therapy plus docetaxel and estramustine versus androgen deprivation therapy alone for high-risk localised prostate cancer (GETUG 12): a phase 3 randomised controlled trial

GETUG-12

Karim Fizazi, Laura Faivre, François Lesaunier, Remy Delva, Gwenaëlle Gravis, Frédéric Rolland, Frank Priou, Jean-Marc Ferrero, Nadine Houede, Loïc Mourey, Christine Theodore, Ivan Krakowski, Jean-François Berdah, Marjorie Baciuchka, Brigitte Laguerre, Aude Fléchon, Alain Ravaud, Isabelle Cojean-Zelek, Stéphane Oudard, Jean-Luc Labourey, Paule Chinet-Charrot, Eric Legouffe, Jean-Léon Lagrange, Claude Linassier, Gaël Deplanque, Philippe Beuzeboc, Jean-Louis Davin, Anne-Laure Martin, Muriel Habibian, Aqnès Laplanche, Stéphane Culine

413 paz High Risk

Gleason score of 8 or greater, stage T3 or T4 disease, serum PSA concentration of 20 ng/mL or more, or pathological node-positive disease.

ADT for 3 years + 4 cycles

DOCETAXEL /Estramustine

Staging — random

Lymphadenectomy ADT for 3 years

Local treatments within three months from systemic therapy 6% prostatectomy (only if N0) 87% radiotherapy (for N0 or N+)

Lancet Oncol 2015; 16: 787-94

Androgen deprivation therapy plus docetaxel and estramustine versus androgen deprivation therapy alone for high-risk localised prostate cancer (GETUG 12): a phase 3

GETUG-12

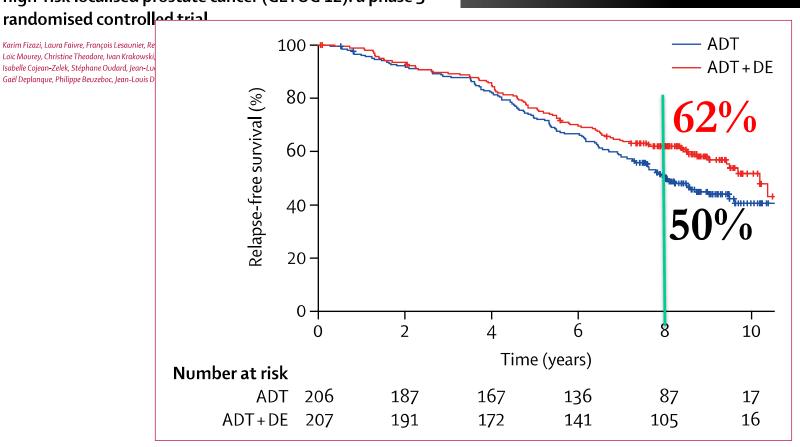


Figure 2: Relapse-free survival

ADT=androgen deprivation therapy. DE=docetaxel and estramustine.

but...Cancer-specific mortality not improved (8 vs 11%)

EVIDENCE FROM LITERATURE: Robotic surgery

Robotic surgery also in high risk patients?

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European Association of Urology

Platinum Priority - Review - Prostate Cancer Editorial by Francesco Montorsi on pp. 928-930 of this issue

The Role of Robot-assisted Radical Prostatectomy and Pelvic Lymph Node Dissection in the Management of High-risk Prostate Cancer: A Systematic Review

Bertram Yuh ^{a,*}, Walter Artibani ^b, Axel Heidenreich ^c, Simon Kimm ^d, Mani Menon ^e, Giacomo Novara ^f, Ashutosh Tewari ^g, Karim Touijer ^d, Timothy Wilson ^a, Kevin C. Zorn ^h, Scott E. Eggener ⁱ

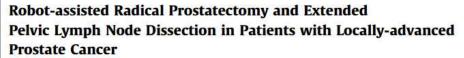
EUROPEAN UROLOGY 71 (2017) 249-256

available at www.sciencedirect.com journal homepage: www.europeanurology.com



European Association of Urology

Surgery in Motion



Giorgio Gandaglia ^{a,b,c,*}, Elisa De Lorenzis ^d, Giacomo Novara ^c, Nicola Fossati ^{a,b,c}, Ruben De Groote ^e, Zach Dovey ^c, Nazareno Suardi ^{a,b}, Francesco Montorsi ^{a,b}, Alberto Briganti ^{a,b}, Bernardo Rocco ^d, Alexandre Mottrie ^c

Conclusions: RARP represents a well-standardized, safe, and oncological effective option in patients with locally advanced PCa. Pathologic stage, Gleason score, and positive margins should be considered to select patients for multimodal approaches.

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JOURNAL OF
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Scand J Urol, 2015; Early Online: 1–7 DOI: 10.3109/21681805.2015.1042038



ORIGINAL ARTICLE

A comparative study of erectile function and use of erectile aids in high-risk prostate cancer patients after robot-assisted laparoscopic prostatectomy

Marie Østby-Deglum¹, Bjørn Brennhovd ², Karol Axcrona^{2,3}, Sophie D. Fosså^{4,5} and Alv A. Dahl^{4,5}

Conclusions. Nearly half of the sample used erectile aids, which significantly increased the proportion with sufficient erection in all risk groups after RALP. With and without the use of erectile aids, the proportions of patients with sufficient erection were 30% or less, with non-significant differences between groups.

¹Student, Faculty of Medicine, University of Southern Denmark, Odense, Denmark, ²Department of Urology, Oslo University Hospital, Radiumhospitalet, Oslo, Norway, ³Department of Urology, St. Olav's University Hospital, Trondheim, Norway, ⁴National Advisory Unit for Late Effects after Cancer Therapy, Oslo University Hospital, Radiumhospitalet, Oslo, Norway, and ⁵Faculty of Medicine, University of Oslo, Oslo, Norway

EVIDENCE FROM LITERATURE: Radiotherapy

Clinical Genitourinary Cancer, Vol. 15, No. 3, 376-83 © 2017 High-Risk Prostate Cancer and Radiotherapy: The Past and the Future. A Benchmark for a New Mixed Beam Radiotherapy Approach

Giulia Marvaso,¹ Barbara A. Jereczek-Fossa,^{1,6} Giulia Riva,⁶ Camilla Bassi,² Cristiana Fodor,¹ Delia Ciardo,¹ Raffaella Cambria,³ Floriana Pansini,³ Dario Zerini,¹ Paolo De Marco,³ Federica Cattani,³ Ottavio De Cobelli,^{4,6} Roberto Orecchia^{5,6}

Survival Outcomes of Dose-Escalated External Bean Radiotherapy versus Combined Brachytherapy for Intermediate and High Risk Prostate Cancer Using the National Cancer Data Base

Arya Amini, Bernard Jones, Matthew W. Jackson, Norman Yeh, Timothy V. Waxweiler, Paul Maroni, Brian D. Kavanagh and David Raben*

From the Department of Radiation Oncology and Division of Urology, Department of Surgery (PM), University of Colorado School of Medicine, Aurora, Colorado

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http://dx.doi.org/10.1016/j.juro.2015.11.005 Vol. 195, 1453-1458, May 2016 Printed in U.S.A. Research Open Access

Outcomes of hypofractionated stereotactic body radiotherapy boost for intermediate and high-risk prostate cancer

Mekhail Anwar 🕍 9 , Vivian Weinberg , Zachary Seymour , I. Joe Hsu , Mack Roach III and Alex R. Gottschalk

Radiation Oncology 2016 11:8

https://doi.org/10.1186/s13014-016-0585-y © Anwar et al. 2016

Received: 12 October 2015 | Accepted: 7 January 2016 | Published: 21 January 2016







Cancer-specific mortality of high-risk prostate cancer after carbon-ion radiotherapy plus long-term androgen deprivation therapy

Goro Kasuya, 1 D Hitoshi Ishikawa, 2 Hiroshi Tsuji, 1 Yasuo Haruyama, 3 Gen Kobashi, 3 Daniel K. Ebner, 1.4 Koichiro Akakura, 5 Hiroyoshi Suzuki, 6 Tomohiko Ichikawa, 7 Jun Shimazaki, 7 Hirokazu Makishima, 1 Takuma Nomiya, 8 Tadashi Kamada, 1 and Hirohiko Tsujii 1 the Working Group for Genitourinary Tumors

EVIDENCE FROM LITERATURE: Multidisciplinary Team discussion



BJUI

The 6-year attendance of a multidisciplinary prostate cancer clinic in Italy: incidence of management changes

Tiziana Magnani*, Riccardo Valdagni*[†], Roberto Salvioni[†], Sergio Villa[†], Lara Bellardita[§], Simona Donegani[§], Nicola Nicolai[†], Giuseppe Procopio[†], Nice Bedini[†], Tiziana Rancati[†] and Nadia Zaffaroni^{††}

*Prostate Cancer Programme, Scientific Director's Office, [†]Division of Radiation Oncology 1, [†]Division of Urology, *Prostate Cancer Program, Psychology Service, [‡]Division of Medical Oncology 2, ^{††}Division of Molecular Pharmacology, Fandazione IRCCS Istituto Nazionale dei Tumori, Milan, Italy Accepted for publication 9 November 2011

Enhancing Prostate Cancer Care Through the Multidisciplinary Clinic Approach: A 15-Year Experience

By Leonard G. Gomella, MD, Jianqing Lin, MD, Jean Hoffman-Censits, MD, Patricia Dugan, RN, Fran Guiles, RHIA, CTR, Costas D. Lallas, MD, Jaspreet Singh, DO, Peter McCue, MD, Timothy Showalter, MD, Richard K. Valicenti, MD, Adam Dicker, MD, and Edouard J. Trabulsi, MD

Kimmel Cancer Center, Thomas Jefferson University, Philadelphia, PA



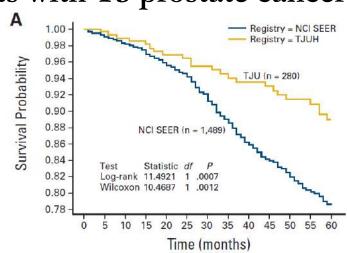
Critical Reviews in Oncology/Hematology 95 (2015) 133-143



Prostate Cancer Unit Initiative in Europe: A position paper by the European School of Oncology

Riccardo Valdagni ^{a,b,c,*}, Hendrik Van Poppel ^d, Michael Aitchison ^e, Peter Albers ^f,
Dominik Berthold ^g, Alberto Bossi ^h, Maurizio Brausi ^l, Louis Denis ^{j,k},
Lawrence Drudge-Coates ^l, Maria De Santis ^{m,n}, Günther Feick ^{j,o}, Chris Harrison ^p,
Karin Haustermans ^d, Donal Hollywood ^{r,1}, Morton Hoyer ^s, Henk Hummel ^l, Malcolm Mason ^u,
Vincenzo Mirone ^v, Stefan C. Müller ^w, Chris Parker ^x, Mahasti Saghatchian ^y,
Cora N. Sternberg ^z, Bertrand Tombal ^{aa}, Erik van Muilekom ^{bb}, Maggie Watson ^{cc},
Simone Wesselmann ^{dd}, Thomas Wiegel ^{ce}, Tiziana Magnani ^b, Alberto Costa ^a

Patients with T3 prostate cancer



CLINICAL CASE: Prostatectomy

No MDS discussion, no additional radiological staging. 27 July 2017: open prostatectomy with extended external iliac and obturator lymph node dissection

Adenocarcinoma 4+5=9, grade group 5, pT3b pN1

Left seminal vescicle invasion

Left margin positive

Lymphatic and perineural invasion present

Two positive (left and right) out of 12 total lymphnodes



EVIDENCE FROM LITERATURE:

New Prognostic Tools

Genomic Test	Oncotype Dx [™] (GPS)	Prolaris™ (CCP)	Decipher™	
Company:	Genomic Health	Myriad Genetics	Genome Dx Biosciences	
mRNA signature:	12 genes + 5 reference	31 genes + 15 housekeeping	22 genes	
Assay:	Quantitative –RT PCR	qPCR	Microarray	
Class of genes:	Stromal response, androgen signaling, proliferation, cellular organization	Cell cycle progression	Proliferation, migration, adhesion, androgen signaling, immune system	
Score range:	0 to 100	-3 to 7	0 to 1	
Initially developed to predict:	Risk of adverse pathology on RP	Risk of progression post- RP and in untreated cohorts	Risk of recurrence and mets post-RP with high specificity	
NCCN guidelines:	Post-biopsy for low- and very low-risk w. 10-20 year life exp.	Post-biopsy for low- and very low-risk w. 10 year life exp.	Patients treated w. RP and adverse pathology	

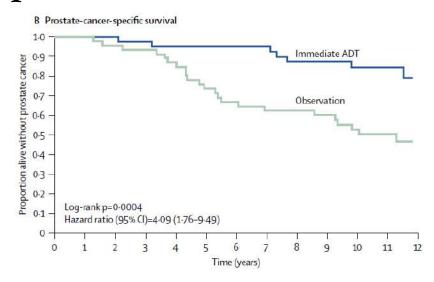
EVIDENCE FROM LITERATURE: Androgen Ablation

Post-operative ADT in pN+ patients

Lancet Oncol 2006: 7: 472-79

• Immediate versus deferred androgen deprivation treatment in patients with node-positive prostate cancer after radical prostatectomy and pelvic lymphadenectomy

Edward M Messing, Judith Manola, Jorge Yao, Maureen Kiernan, David Crawford, George Wilding, P Anthony di Sant Agnese, Donald Trump, on behalf of the Eastern Cooperative Oncology Group study EST 3886





Do not perform a limited LND.				
Up	on detection of nodal involvement during RP:			
	offer adjuvant androgen deprivation therapy (ADT);	1b	А	
•	discuss adjuvant ADT with additional radiotherapy (see Section 6.2.6.3);	2b	А	
•	offer observation (expectant management) to a patient after eLND with < 2 nodes with microscopic involvement, and a PSA < 0.1 ng/mL and absence of extranodal extension.	2b	В	

EVIDENCE FROM LITERATURE: Radiotherapy

Post-operative radiation therapy in pN+ patients

JOURNAL OF CLINICAL ONCOLOGY ORIGINAL REPORT

Impact of Adjuvant Radiotherapy on Survival of Patients With Node-Positive Prostate Cancer

Firas Abdollah, R. Jeffrey Karnes, Nazareno Suardi, Cesare Cozzarini, Giorgio Gandaglia, Nicola Fossati, Damiano Vizziello, Maxine Sun, Pierre I. Karakiewicz, Mani Menon, Francesco Montorsi, and Alberto Briganti

We evaluated 1,107 patients with pN1 prostate cancer treated with radical prostatectomy and anatomically extended pelvic lymph node dissection between 1988 and 2010 at two tertiary care centers. All patients received adjuvant hormonal therapy with or without aRT. Regression tree

Conclusion

The beneficial impact of aRT on survival in patients with pN1 prostate cancer is highly influenced by tumor characteristics. Men with low-volume nodal disease (≤ two PLNs) in the presence of intermediate- to high-grade, non–specimen-confined disease and those with intermediate-volume nodal disease (three to four PLNs) represent the ideal candidates for aRT after surgery.

Original Article

The Role of Adjuvant Radiotherapy in Pathologically Lymph Node Positive Prostate Cancer

Naresh Jegadeesh, MD¹; Yuan Liu, PhD^{2,3,4}; Chao Zhang, PhD^{2,3,4}; Jim Zhong, MD^{1,2}; Richard J. Cassidy, MD^{1,2}; Theresa Gillespie, PhD^{2,5}; Omer Kucuk, MD^{2,6}; Peter Rossi, MD^{1,2}; Viraj A. Master, MD, PhD^{2,6}; Mehrdad Alemozaffar, MD^{2,7}; and Ashesh B. Jani, MD^{1,2}

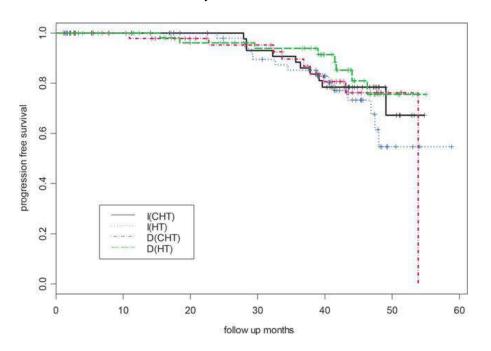
Retrospective analysis in National Cancer Database on > 2,500 pN1M0 patients

n. CONCLUSIONS: RT plus ADT was associated with im-

proved OS after RP in patients with LNI. These results may help guide therapy in the absence of randomized evidence. *Cancer* 2016;000:000-000. © 2016 American Cancer Society.

EVIDENCE FROM LITERATURE: Docetaxel?

Phase III adjuvant TAX 3501 study



Schweizer Cancer 2013



Early interruption for poor accrual 23% febrile neutropenia

Addition of docetaxel or bisphosphonates to standard of care in men with localised or metastatic, hormone-sensitive prostate cancer: a systematic review and meta-analyses of aggregate data Lancet Oncol 2016; 17: 243–56

META-ANALYSIS

Claire L Vale*, Sarah Burdett*, Larysa H M Rydzewska, Laurence Albiges, Noel W Clarke, David Fisher, Karim Fizazi, Gwenaelle Gravis, Nicholas D James, Malcolm D Mason, Mahesh K B Parmar, Christopher J Sweeney, Matthew R Sydes, Bertrand Tombal, Jayne F Tierney, for the STOpCaP Steering Group

High-risk, non metastatic patients

D				_
	Control	Treatment	Relapse-free Survival	Hazard ratio (95% CI)
GETUG-12 ²⁶	111/206	88/207 —		0.71 (0.54–0.94)
RTOG 0521 ²⁸	123/281	98/281		0.76 (0.58-0.99)
STAMPEDE ⁸ (SOC+/–Doc)	176/460	63/230 —	■	0.60 (0.45-0.80)
STAMPEDE ⁸ (SOC+ZA+/-Doc)	88/227	63/228 —	-	0.69 (0.47–1.01)
TAX 3501 ²⁷ (immediate ADT)	14/55	10/55 —	<u> </u>	0.79 (0.34–1.84)
TAX 3501 ²⁷ (delayed ADT)	8/62	9/56 —		— 1·34 (0·39 – 4·59)
Overall				0.70 (0.61-0.81)
Heterogeneity: $\chi^2 = 2.63$; df=5; p=	=0·757; <i>l</i> ² =0%	0·5	<u> </u>	7
			— · —	2
		Favours SO	C + docetaxel Favours SOC	

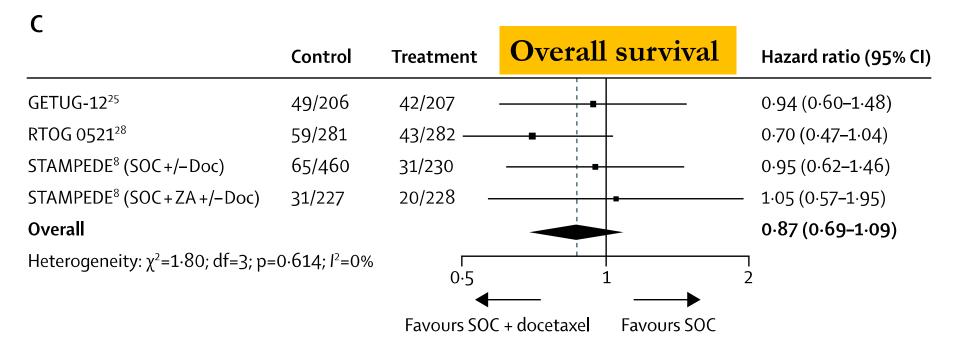
HR=0.7 (95% CI 0·61–0·81, p<0·0001 absolute increase +8% at 4 years

Addition of docetaxel or bisphosphonates to standard of care in men with localised or metastatic, hormone-sensitive prostate cancer: a systematic review and meta-analyses of aggregate data Lancet Oncol 2016; 17: 243-56

META-ANALYSIS

Claire L Vale*, Sarah Burdett*, Larysa H M Rydzewska, Laurence Albiges, Noel W Clarke, David Fisher, Karim Fizazi, Gwenaelle Gravis, Nicholas D James, Malcolm D Mason, Mahesh K B Parmar, Christopher J Sweeney, Matthew R Sydes, Bertrand Tombal, Jayne F Tierney, for the STOpCaP Steering Group

High-risk, non metastatic patients



HR=0.87 95% CI 0.69–1.09; p=0.218) Absolute increase +2% at 4 years

CLINICAL CASE: referral to IOV

September: first MDS consultation @IOV for adjuvant treatments.

Mild incontinence, impotence (not interested in rehabilitation), no bone pain, fully active.

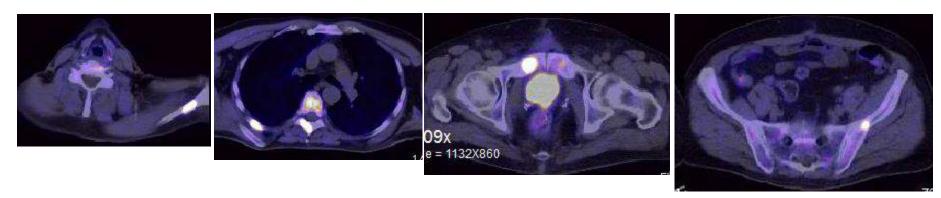
Multidimensional Geriatric Assessment: fit

Pelvic radiotherapy + LH-RH analogue programmed, preceded by physiatric consultation + new PSA

PSA @ 7 weeks from surgery: 4.8 ng/dL

CLINICAL CASE: Re-staging

Staging PET/CT with choline: multiple bone metastases (left scapula, vertebrae, ribs, pubis, ileus)



The patient has now high-risk Castration-sensitive metastatic disease (Charteed definition) and starts Androgen Deprivation + Docetaxel x 6 cycles.

Prato della Valle in Padova in a (very rare!!) snowy day

