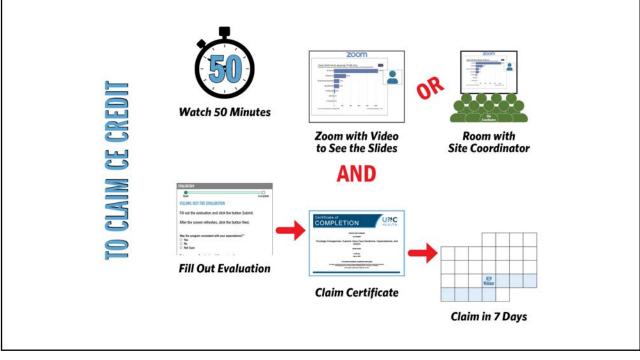
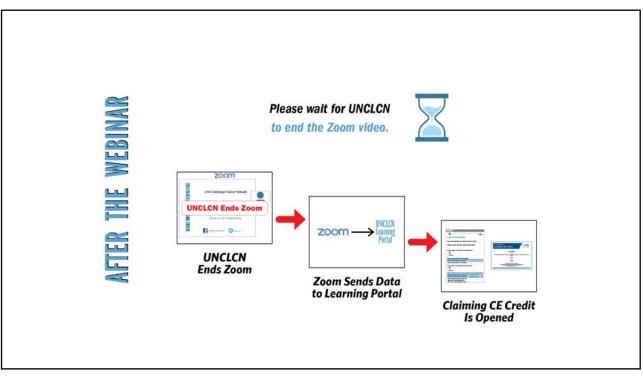
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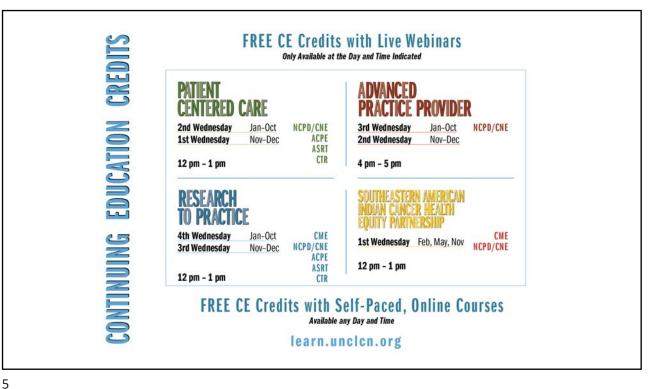




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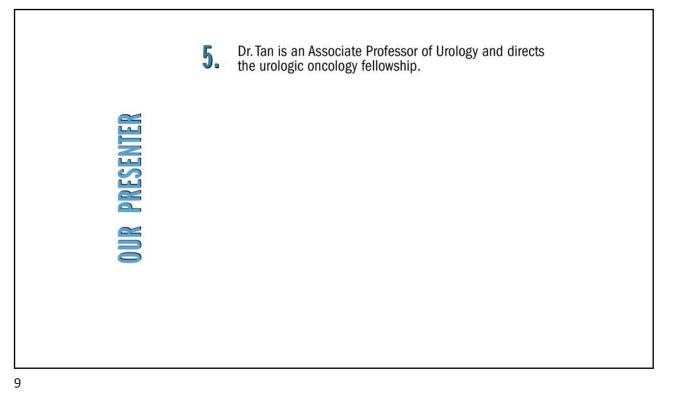


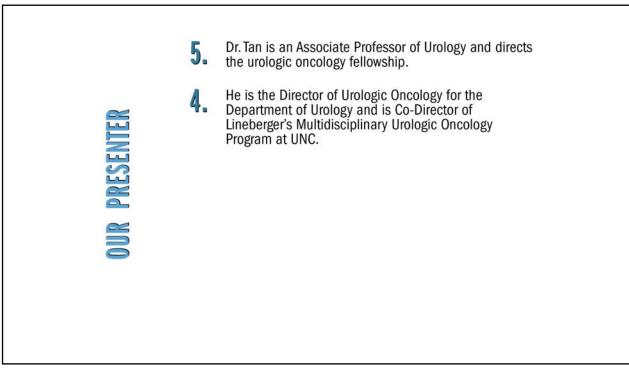
Dr. Tan is an Associate Professor of Urology, Director of Urologic Oncology for the Department of Urology, and Co-Director of the Lineberger Comprehensive Cancer Center Multidisciplinary Urologic Oncology Program at UNC. He also directs the urologic oncology fellowship.

Raised in North Carolina, received his medical degree from the University of Michigan where he also completed his residency in urology. Afterwards, he obtained advanced fellowship training in urologic oncology and health services research through the Institute of Urologic Oncology and the Robert Wood Johnson Foundation Clinical Scholars Program at UCLA.

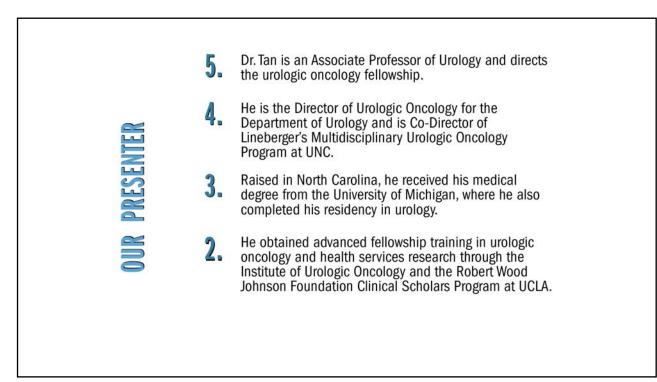
Dr. Tan specializes in the management of prostate, kidney, bladder, and testicular cancer. He performs complex open and robotic surgeries for cancer as well as specialized procedures (e.g., MRI-US fusion & transperineal prostate biopsy, retzius-sparing prostatectomy, robotic RPLND). To further meet the health needs of his patients, Dr. Tan leads a robust research program focuses on decision-making, risk communication, and survivorship with funding support from the American Cancer Society, the Department of Defense, and the National Institutes of Health.



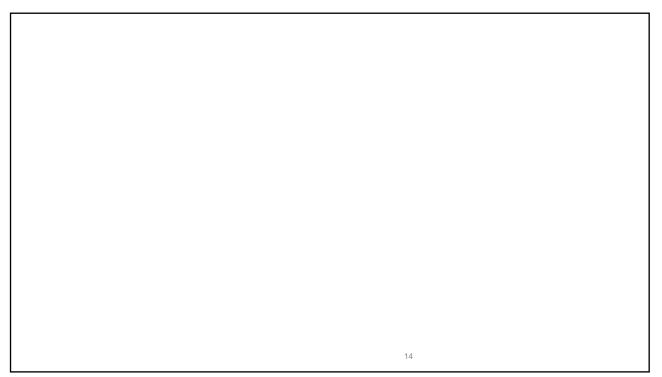




| | 5. | Dr. Tan is an Associate Professor of Urology and directs the urologic oncology fellowship. |
|-----------|----|---|
| NTER | 4. | He is the Director of Urologic Oncology for the Department of Urology and is Co-Director of Lineberger's Multidisciplinary Urologic Oncology Program at UNC. |
| PRESENTER | 3. | Raised in North Carolina, he received his medical degree from the University of Michigan, where he also completed his residency in urology. |
| OUR | | |
| | | |
| | | |

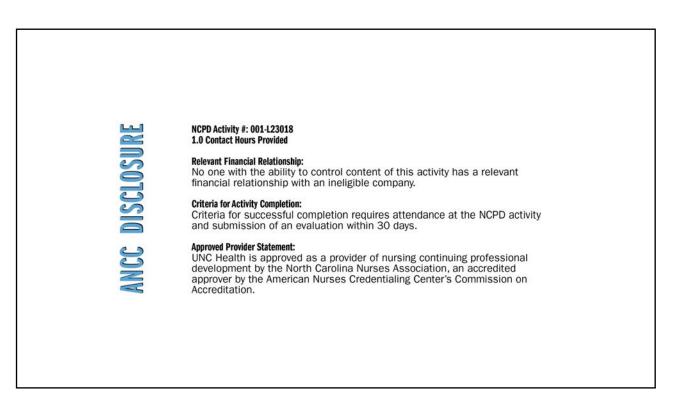


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|-----------|----|--|
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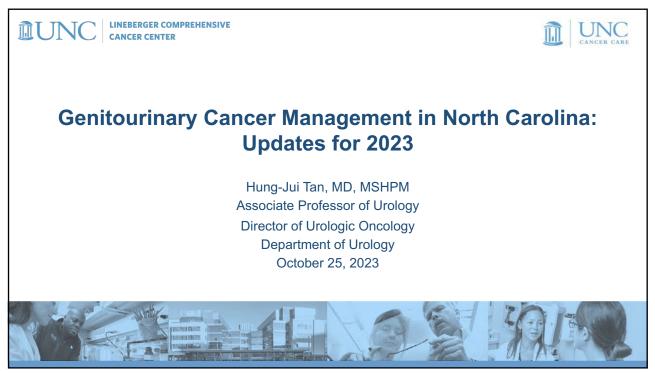


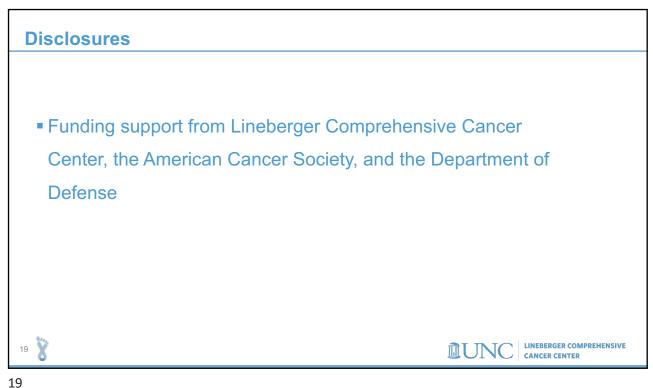
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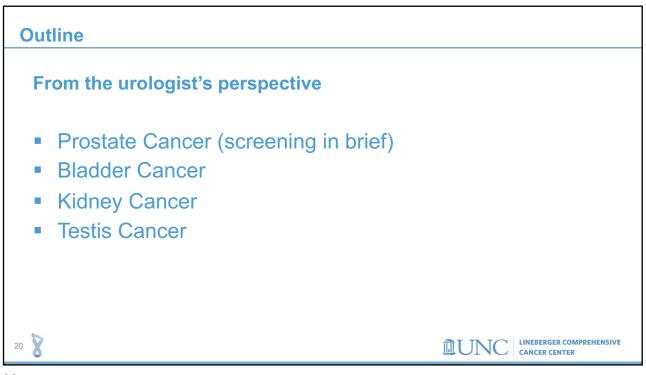
This activity has been planned and implemented under the sole supervision of the Course Director, William A. Wood, MD, MPH, in association with the UNC Office of Continuing Professional Development (CPD). The course director and CPD staff have no relevant financial relationships with ineligible companies as defined by the ACCME.
 A potential conflict of interest occurs when an individual has an opportunity to affect educational content about health-care products or services of a commercial interest with which he/she has a financial relationship. The speakers and planners of this learning activity have not disclosed any relevant financial relationships with any commercial interests pertaining to this activity.
 The presenter has no relevant financial relationships with ineligible companies as defined by the ACCME.







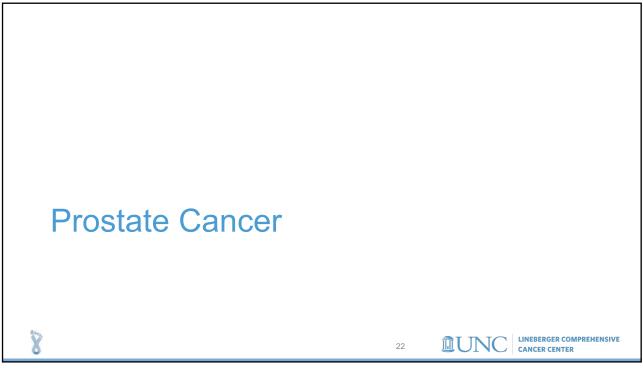




| Breast | |
|----------|----|
| | 0% |
| Prostate | |
| | 0% |
| Lung | |
| | 0% |
| Bladder | |
| | 0% |
| | |
| | |

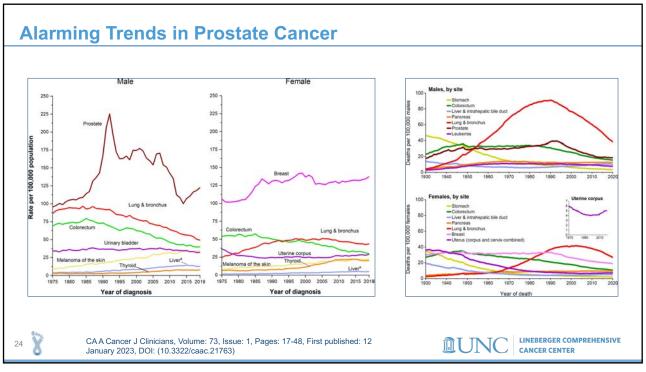
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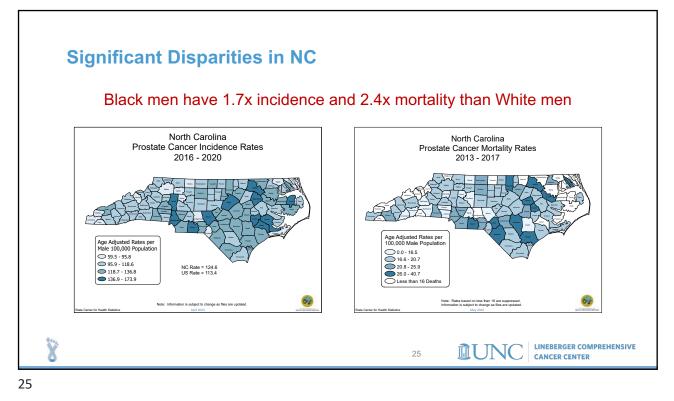
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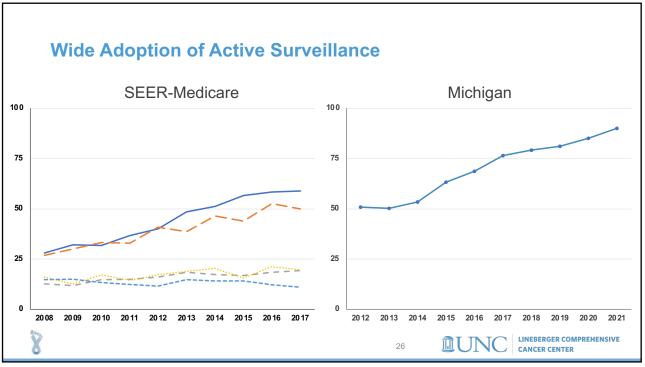


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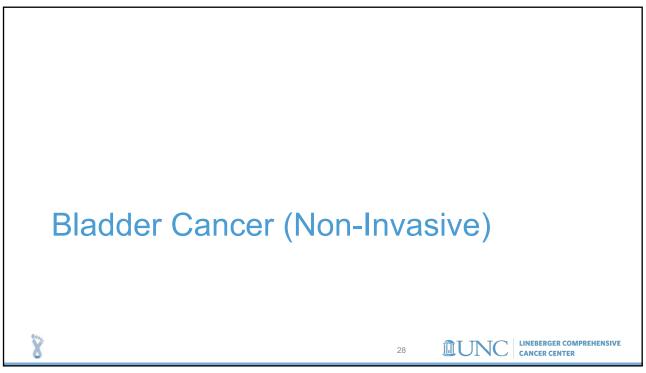




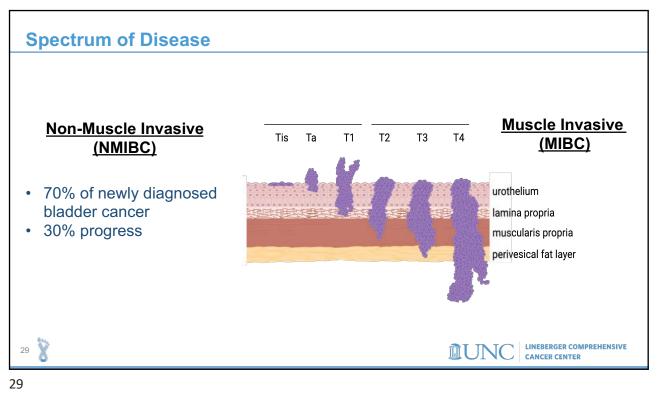


Prostate Cancer Screening Recommendations

| ACS (2023) | AUA (2023) | USPTF (2018) |
|---|---|--|
| Men should have a chance to make an informed decision about initial screening with their provider at: 50 if average risk with at least 10- year life expectancy. 45 if high risk . Black men and those with a 1° relative diagnosed with prostate cancer before age 65. 40 if even higher risk . Men with 2 or more 1° relatives with prostate cancer diagnosed at early age. If PSA <2.5, then every other year testing. Annual testing if >2.5 | Clinicians should engage in shared decision-making and may begin screening and offer a baseline PSA between ages 45-50 years. Clinicians should offer screening at age 40-45 years for people at increased risk: Black ancestry, germline mutations, strong family history of prostate cancer. Clinicians should offer regular prostate cancer screening every 2-4 years for those aged 50 to 69 years. | Grade C: An individual decision for men aged 55-69 . Small potential benefits whereas "many men will experience potential harms." Patients and clinicians should consider family history, race/ethnicity, comorbid medical conditions, patient values, and other health needs. |
| 8 | | |



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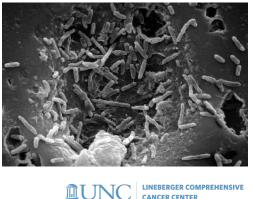
Bacillus Calmette-Guerin (BCG) - historical gold standard

- Intravesical immunotherapy
- Used for >40 years (First report in 1976)
- Exact mechanism unclear
- 70-80% initial response but up to 40% failure long-term
- Risks (Macloed et al., PMID: 25210559):
 - Lower urinary tract symptoms (27%-95%)
 - Fever greater than 39.5°C (2.9%)
 - Rare: granulomatous prostatitis (0.9%), pneumonitis or hepatitis (0.7%), arthralgia (0.5%), epididymitis (0.4%), severe disseminated BCG sepsis (0.4%)
- Frequent shortages associated with supply chain issues, resulting in reduced dosing and inadequate treatment

The Journal of United INTRACAVITARY BACILLUS CALMETTE-GUERIN IN THE TREATM OF SUPERFICIAL BLADDER TUMORS A. MORALES,* D. EIDINGER AND A. W. BRUCE

Morales et al., J Urol, 1976

CANCER CENTER



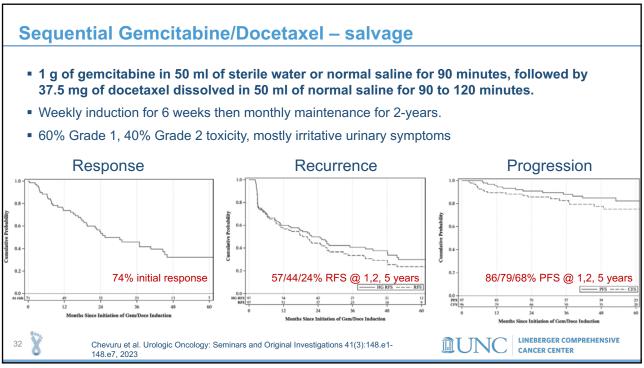
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Comparison of Sequential Intravesical Gemcitabine and Docetaxel vs Bacillus Calmette-Guérin for the Treatment of Patients With High-Risk Non–Muscle-Invasive Bladder Cancer

- Retrospective comparison of BCG (full or reduced dosing, maintenance at 3, 9, 15 months) vs. Gem/Doce (6 week induction then monthly maintenance for up to 2-years) from 2011-2021
- 67% vs. 81% RFS at 1-year >90% PFS, CFS, CSS at 2-years 0.8 9.2% vs. 2.9% discontinuation 0.6 0.4 **BRIDGE Trial (SWOG) – RCT** for BCG vs. Gem/Doce for 0.2 **HR NMIBC** now open Time since initiation of induction, mo at risk BCG 108 91 88 43 69 LINEBERGER COMPREHENSIVE X 31 氲UN(McElree et al., JAMA Netw Open. 2023;6(2):e230849. doi:10.1001/jamanetworkopen.2023.0849 **CANCER CENTER** 31

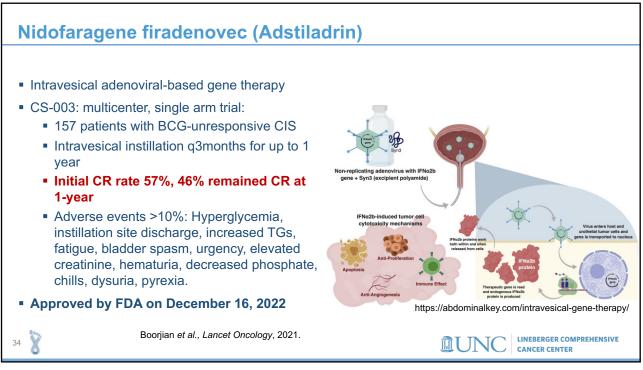


Pembrolizumab

- Anti-programmed cell death (PD)-1 antibody
- Expressed by T-cells and acts as an immune checkpoint inhibitor
- Approved for BCG-unresponsive, high-risk NMIBC based on KEYNOTE-057 (Balar et al., Lancet Oncology 2021)
 - Single arm study of 101 patients → analysis included 96 patients with high-risk CIS ineligible or unwilling to undergo cystectomy
 - Treated with 200 mg of systemic pembrolizumab every 3 weeks for up to 24 months
 - Complete response: 41% at 3 months
 - Adverse events in 2/3 of patients:
 - Most common: diarrhea, fatigue, pruritis (11 serious treatment-related)
 - Immune related adverse events in 22% (most common: hypothyroid)
- Extended follow-up (Balar et al., JCO, 2021):
 - 14% (13 of 39) with CR @ 2 years
 - 41.7% underwent cystectomy for toxicity, recurrence, progression, or persistence

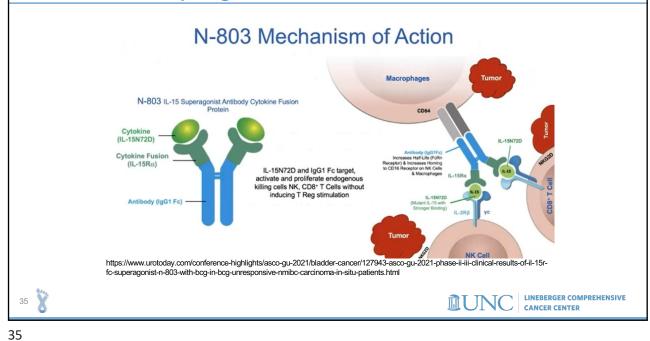
Alliance Trial Intravesical Gemcitabine + Pembrolizumab ongoing at UNC



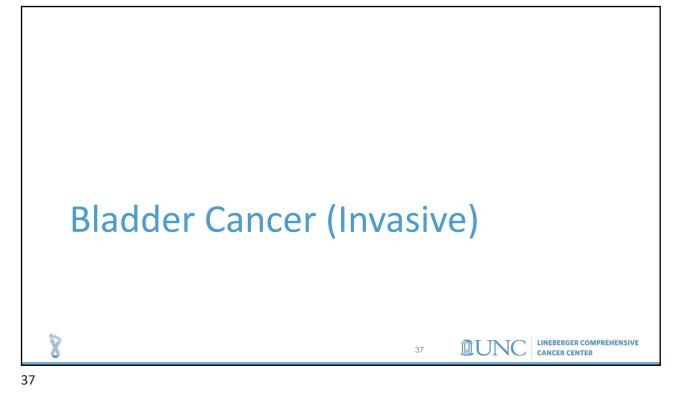


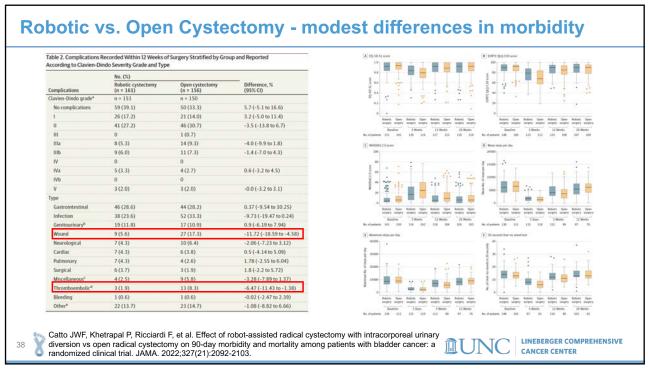
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ALT-803: IL-15 Superagonist

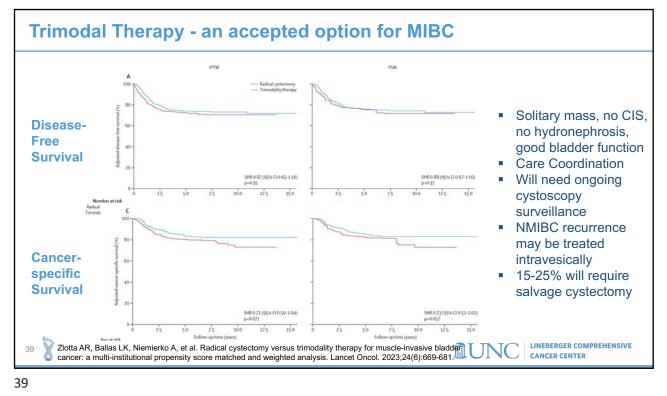


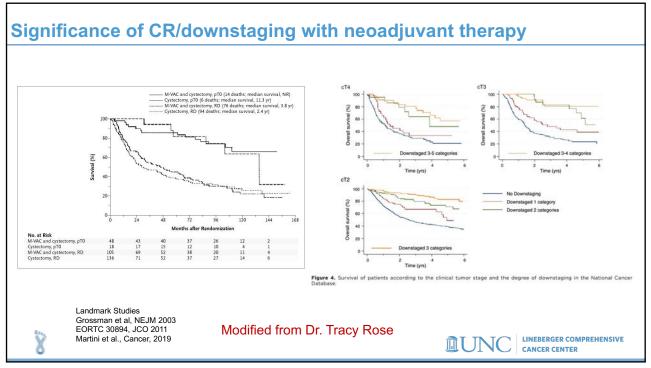
| NIMBO | | a in situ | 03 with BCG in BCG-I (CIS) patients w/wo p ? | |
|--------|--|-----------|--|--|
| Cohort | Population | Treatment | Outcomes | Develop of Cariphon Response 309 Develop of CE = 12 events 41 EIN (89% CE-U28% to 713%) |
| A | BCG- unresponsive CIS +/- Ta/T1 disease | NAI + BCG | CR: 58/82 (71%). >50% CR at 2 years. In patients with CR, probability of cystectomy-free 89.2% and DSS 100% | |
| С | BCG- unresponsive CIS +/- Ta/T1 disease | NAI alone | CR at 3 months = 2/10 patients CR at 6 months = 1/10 patients (discontinued for futility) | At Bits 31 51 52 34 34 35 36 |
| В | BCG- unresponsive HG Ta/T1 disease | NAI + BCG | DFS rate at 12-months of 55.4% Median DFS 19.3 months | 8 |





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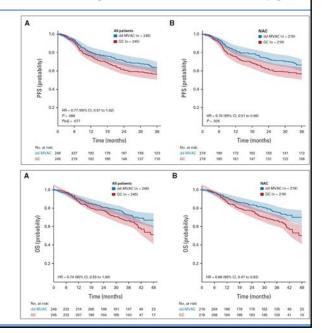


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Dose-dense MVAC vs Gem/Cisplatin for neoadjuvant chemotherapy

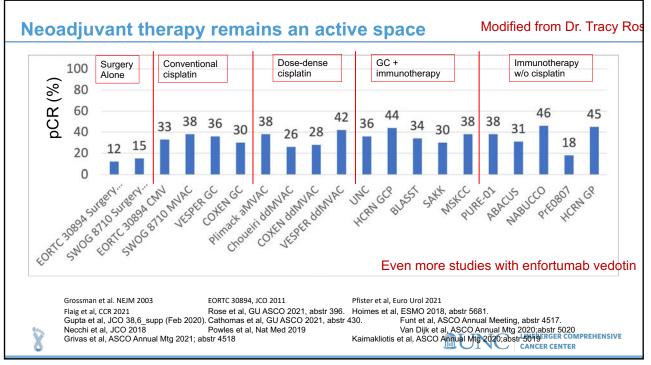


- 500 patients randomized between 6 cycles ddMVAC vs. 4 cycles GC of which 89% received as NAC
- 60% in ddMVAC received 6 cycles vs. 84% in GC received 4 cycles
- pCR 42% vs. 36%
- <ypT3 77% vs. 63%</p>
- 3-year PFS 66% vs. 56% for NAC, HR 0.70, p=0.025
- 5-year survival data in NAC: 66% vs. 57% OS, 75% vs. 60% CSS.



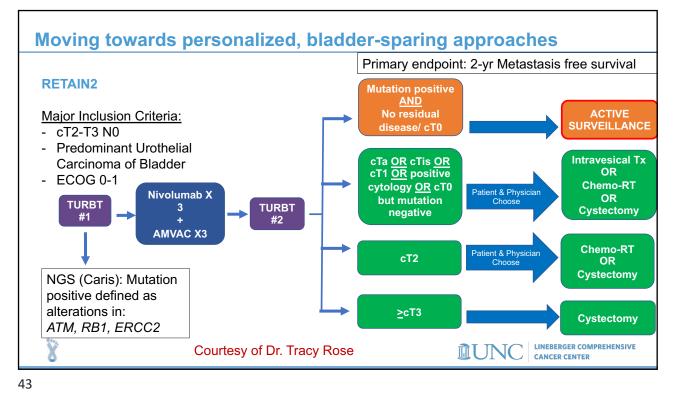


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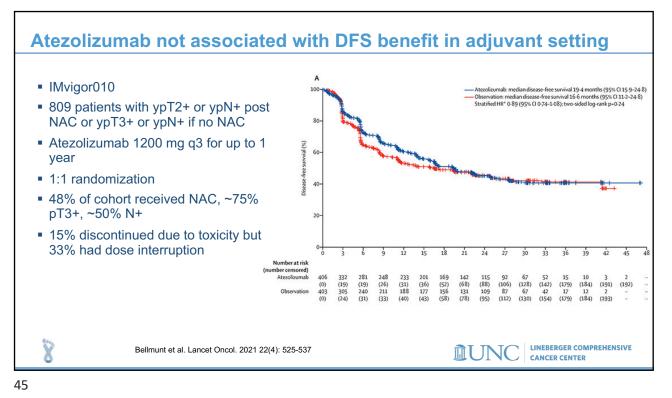


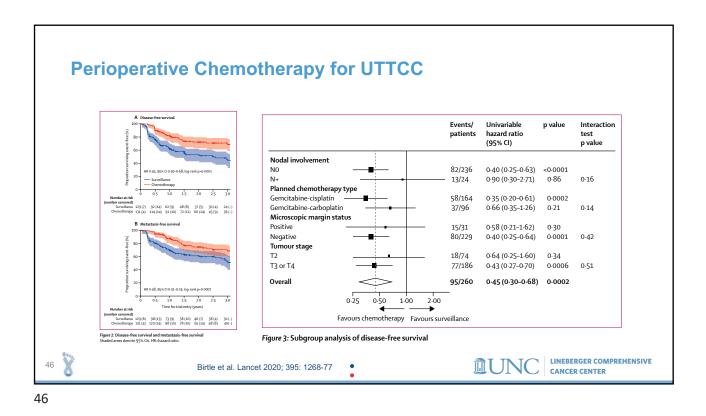
Pfister et al. VESPER JCO, 2022. Pfister at al. VESPER Eur Urol, 2021.

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Nivolumab associated with prolonged DFS in patients with MIBC tion-to-Treat Popul rvival Survival (95% CI) at 12 Mo (95% CI) CheckMate 274 74.9 (69.9–79.2) 62.8 (57.3–67.8) 60.3 (54.9–65.3) 46.6 (41.1–51.9) 709 patients with ypT2+ or ypN+ post NAC or ypT3+ or ypN+ if no NAC and a Nivolumab 240 mg IV q2 weeks for 12 15 18 21 24 27 30 33 36 39 42 45 48 51 up to 1 year 1:1 randomization with stratification 353 296 244 212 178 154 126 106 85 68 57 51 36 23 20 3 1 0 356 248 198 157 134 121 105 94 80 65 54 50 37 22 19 10 2 0 based on PDL1, NAC, nodal status ith a PD-L1 Ex ion Level of =1% Survival Survival at 6 Mo (95% CI) at 12 Mo (95% CI) 43% of cohort received NAC, ~60% 90 80 70 60 50 40 30 20 10 pT3, ~50% N+ 74.5 (66.2-81.1) 67.2 (58.4-74.5) 55.7 (46.8-63.6) 45.9 (37.1-54.2) (%) nazard ratio for disease recur 0.55 (98.72% Cl. 0.35–0.85) P<0.001 12% discontinued nivolumab due to toxicity 9 12 15 18 21 24 27 30 33 36 39 42 45 48 51 Approved by FDA on August 2021 140 113 98 91 76 68 58 50 38 31 27 24 21 12 10 1 0 0 142 90 73 59 53 49 42 37 28 22 17 16 12 7 5 3 1 0 Bajorin DF, Witjes JA, Gschwend JE, et al. Adjuvant nivolumab versus placebo in muscle-invasive urothelial LINEBERGER COMPREHENSIVE 44 carcinoma. N Engl J Med. 2021;384(22):2102-2114. **CANCER CENTER**

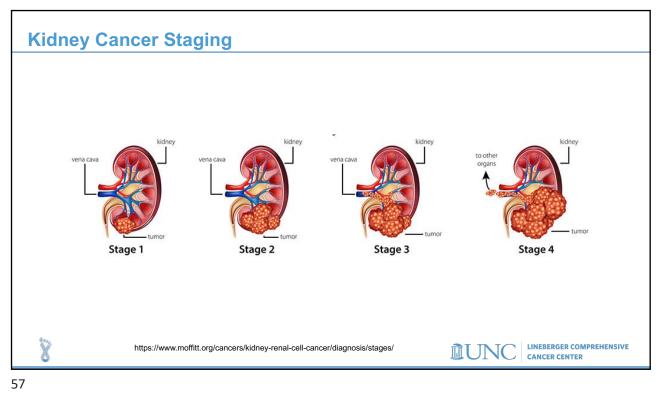


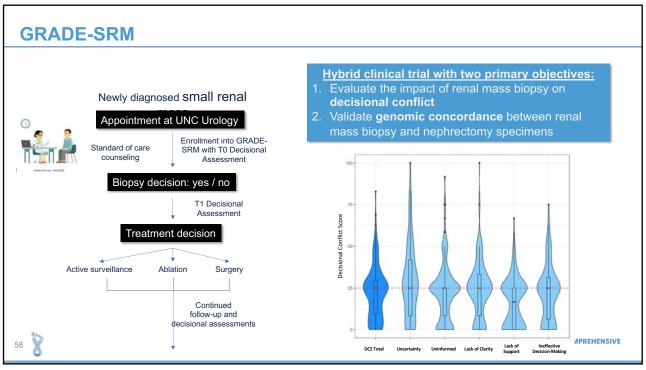




| BUNK: semantic What percent of | f clinical T1a renal masses are benign? | |
|--------------------------------|--|----|
| | | |
| 0-10% | | |
| | | 0% |
| 10-20% | | 0% |
| 20-30% | | |
| | | 0% |
| 30-40% | | |
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| | Start the presentation to see live content. For screen share software, share the entire screen. Get help at pollev.com/app | |

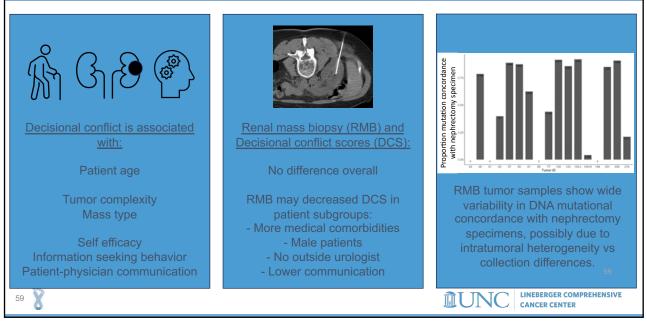
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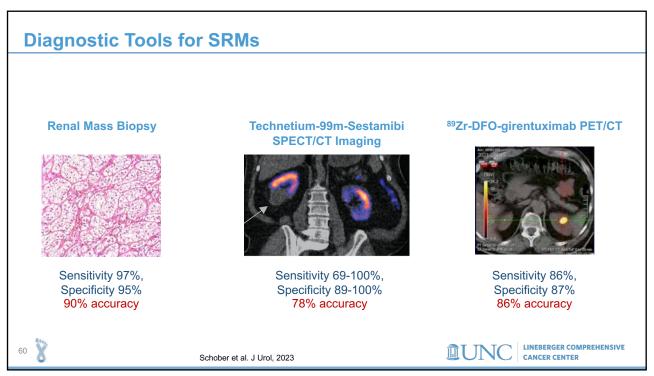


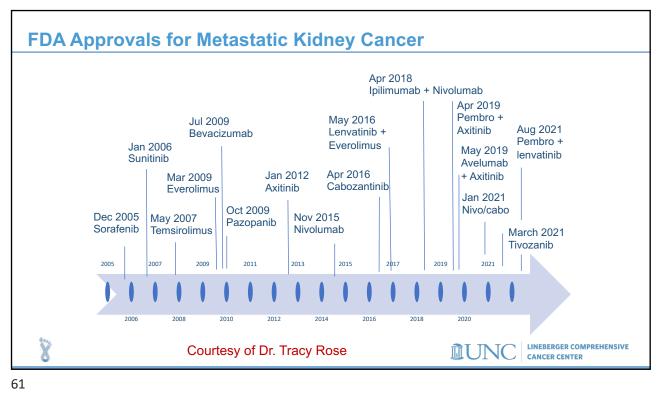


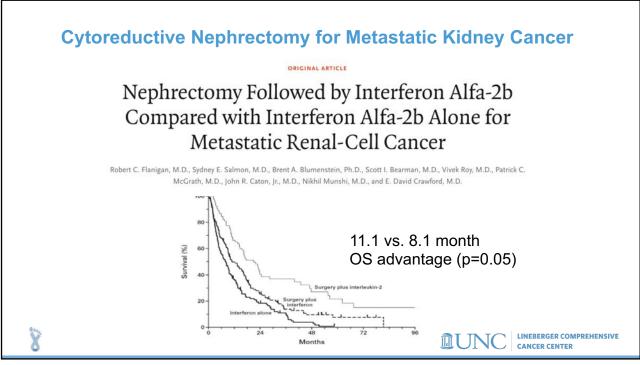
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GRADE-SRM



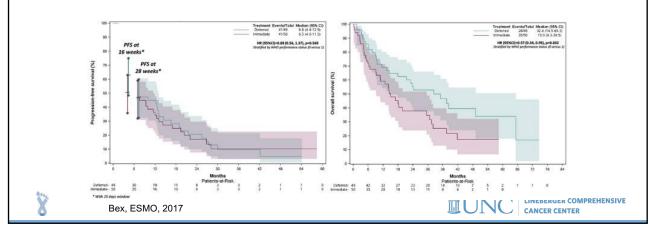


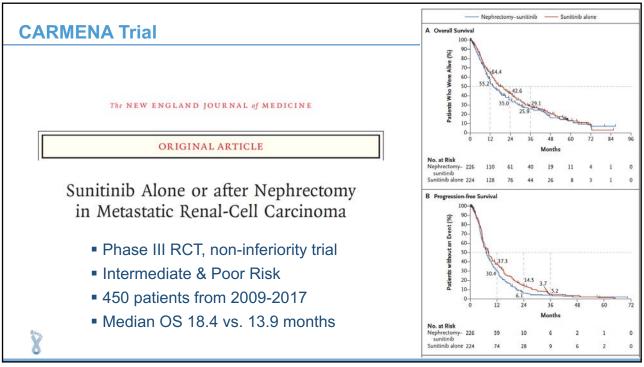




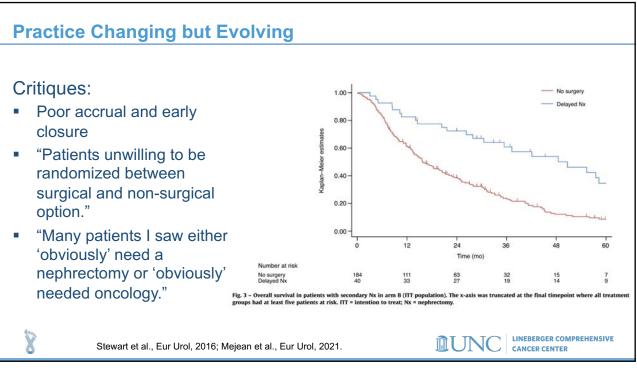
SURTIME Trial

- 99 patients from 19 institutions
- CN then sunitinib vs. 3 cycles sunitinib then CN

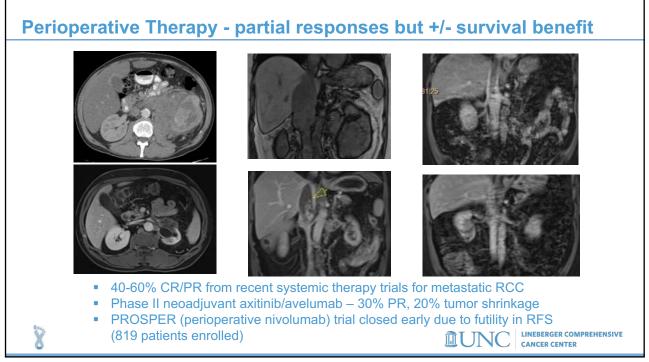




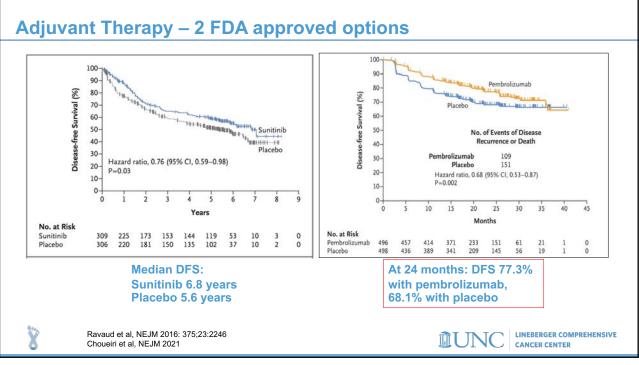
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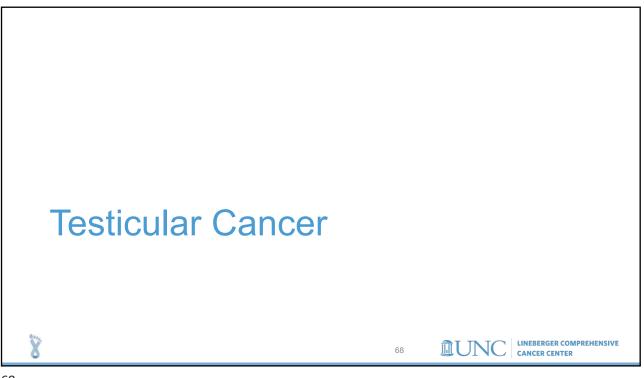






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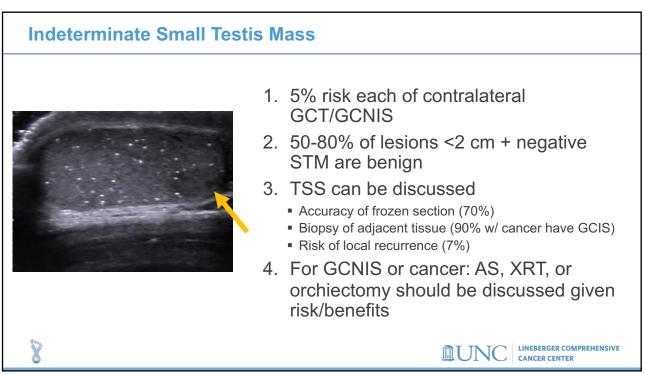
| Active surveillance | |
|---------------------|----|
| | 0% |
| Chemotherapy | |
| | 0% |
| Radiation Therapy | |
| | 0% |
| RPLND | 0% |
| | 0% |
| | |
| | |

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10/25/20

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Testis Cancer Guidelines noma and non-semine ma: ESMO-EURACAN Clinical Practic Highlights from NCCN/ESMO/EAU Suideline for diagnosis, treatment and follow-up 1. Stage I Risk factors Seminoma: retes testes, >3 cm – 15-30% recurrence NSGCT: LVI/embryonal – 30-50% recurrence **Testicular Cancer** 2. Minor elevation in STM \neq chemo Version 2 2022 — January 4, 2022 3. PC-late relapses (>2 years) likely yolk **EAU Guidelines on** sac or teratoma **Testicular** 4. PET only for seminoma, PC >3cm Cancer 8 LINERERGER COMPREHENSIVE **UNC** CANCER CENTER 72



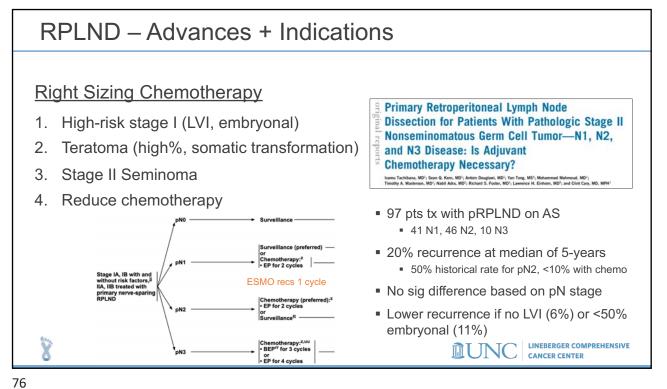
RPLND for Seminoma

Primary RPLND for Seminoma

| | SEMS (2021 GU ASCO) | PRIMETEST (2022 GU ASCO) |
|--------------|---|--|
| Population | - 55 patients with Stage II A/B Seminoma - Max LN 3 cm - 14 progression on AS - 15 sites in North America | - 33 patients with Stage II A/B Seminoma - 9 de novo, 19 progression on AS, 5 progression after carboplatin - Single site in Germany |
| Intervention | Primary RPLND (open, modified) | Primary RPLND (robot and open) |
| Comparison | Single Arm | Single Arm |
| Outcomes | - 10 recurrences (18%) at 2-years - 2 (4%) with major complications - No retrograde ejaculation | - 10 (31%) recurrences at median 2-years - 3 in-field recurrences - 3 (10%) with major complications - No difference between robotic and open |

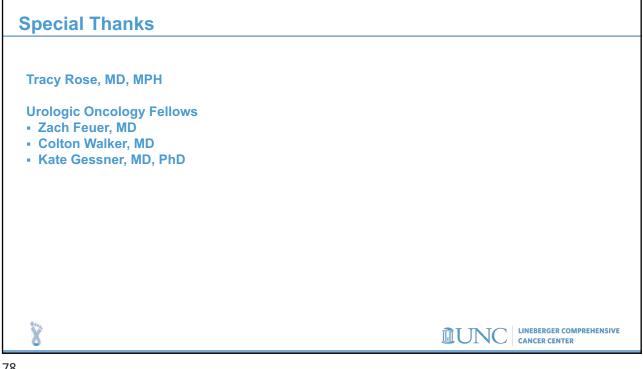
| | | Adverse Security 1 October of Associated with | |
|--|---|---|--|
| Primary Robot-assisted Retroperitoneal | | Adverse Surgical Outcomes Associated with Robotic Retroperitoneal Lymph Node Dissection | |
| for Men with Nonseminomatous Germ Cell Tumor: Experience from a Multi-institutional Cohort | | Among Patients with Testicular Cancer | |
| lacob Taylor ^{6,*} , Ezequiel Becher ⁶ , James S. Wysock ⁶ , And lacob Jipp ⁵ , Peter Langenstroer ⁶ , Scott Johnson ⁶ , Marc A Brian R. Lane ⁶ , William C. Huang ⁶ WU Langene Health, New York, NY, USA ⁴ University of California Les Angeles. Les Angeles. University of Nerth Carolina at Chapel Hill, Chapt Hill, NC, USA ⁴ Sperrum Health, Cand | . Bjurlin ^d , Hung-Jui Tan ^d , CA. USA: [«] Medical College of Wisconsin, Milwaukee, WI, USA: | Adam C. Calaway ^{®, *} , Lawrence H. Einhorn [®] , Timothy A. Masterson [®] , Richard S. Foster [®] , Clint Cary [®] *Department of Unelogy, Indiana University School of Medicine, Indianapolit, IN, USK [®] Department of Oncology, Indiana University School of Med Indianapolit, IN, USK | |
| Operative Time (median) | 288 minutes | | |
| EBL (median) | 100 cc | | |
| LOS (median) | 1 day | | |
| Readmissions | 6% | C D | |
| Major Complications | 4% | | |
| Recurrences (median 15 months) | 8% | | |
| Lymph Node Count (median) | 32 | | |

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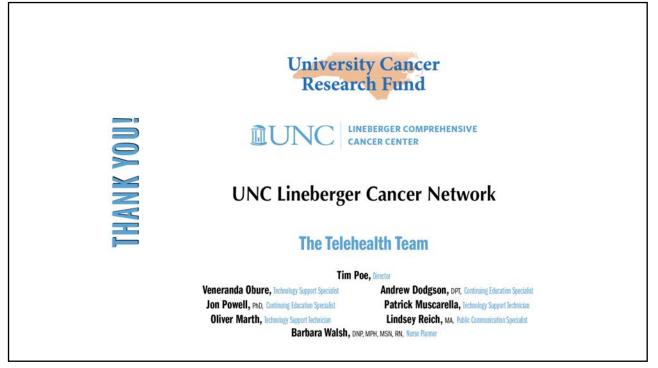


Monitoring Advances TRISST: Modality + Intensity **Imaging Modality and Frequency in** OF Surveillance of Stage I Seminoma Testicular CT vs. MRI: 2.6% vs. 0.6% Cancer: Results From a Randomized, Phase III, Noninferiority Trial (TRISST) san K. Joffe, MBBS, MD'; Fay H. Cafferty, PhD⁺; Laura Murphy, PhD⁺; Gordon J.S. Rustin, MBBS, MS-; MD⁺; Schalb, BS-, MBBS⁺; Rhian Gabe, PhD⁺; Saliry P. Stenning, MS⁺; Elizabeth James, MS⁺; Ojoa Noor, MS⁺; Simona Wade, BS sca Schiwrone, PhD⁺; Savan Bwitt, MRCP⁺; Eliaine Dumoodie, MBChB, MRCP, MD⁺; Marcia Hall, MBBS PhD⁺*; Sharma, MBBS, MICD⁺; Joney Braynobe, BSc, BM, PhD⁺; Jonathan Sharmash, MBCR, MD⁺; John Lopev, M⁺B NGrding, BS-; RMBS⁺, MRCP⁺; Ivo Hennig, MD, PhD⁺; Jeff White, MBChB, DM⁺; Savah Rudman, BSCHons), PhD, MBBS Jonefing, BS-; RMBCP⁺; Too Hennig, MD, PhD⁺; Jeff White, MBChB, DM⁺; Savah Rudman, BSCHons), PhD, MBBS Jones, MBChB, MD⁺; Michael Secki, PhD⁺; Graham MacDonald, MBChB, MRCP⁺; Ramachandrar Venkitaraman, MIC 7 vs 3 scans: 0.3% vs. 2.8% Thiaganjan Seeninasan, MBBS, MCPEKainiy^{11,12}, ustnam MacDonald, MBChB, MRCP¹¹, Danish Mashar, PhD¹¹, Nictosia Goyle, MBBCh, BAO, PhD¹¹, Martin Highley, MD¹¹, Tom Geldart, MBBS, BSc¹¹, Robert Laing, MBBS, MRCP¹¹, Richard S, Kaplan, MD¹¹, and Robert A. Huddart, PhD, MA(Oxon), MBBS, MRCP¹¹, on behalf of the TRISST Trail Management Group and Investigators MD¹⁸; Michael Seckl, PhD¹⁹; Graham n, MBBS, MRCP(Edin)^{21, 22}; Satish Kur V DFS similar across all 4 669 stage I seminoma pts in 35 UK centers groups (85-89%) Non-inferior RCT: 7 vs 3 scans; CT vs. MRI >80% compliance X LINEBERGER COMPREHENSIVE <u><u></u>∭UN(</u> **CANCER CENTER**

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Presented 10/25/20



