25th ASIA-PACIFIC PROSTATE CANCER CONFERENCE



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# ANDROGEN DEPRIVATION THERAPY AND QUALITY OF LIFE

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RESEARCHER



**CLINICIAN** 





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### Disclosure Andrew Matthew, PhD., C.Psych

- None





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#### Outcomes of the AUSTRALIAN Convening (January 2025)

THE GOAL: To understand how to improve access to sexual health for more Australians with

prostate cancer.



Care coordination between treating physicians, primary care and allied health



Staged approach to workforce training



Increased dissemination of patient education and material



Improve sexual health care through primary care physicians



Patients with advanced s cancer are an overld





#### Prostate Cancer and ADT

Estimates: 45-50% of men will receive ADT during the course of their disease (MENG 2002, Gilbert 2011)

- Radiation Treatment with adjuvant androgen deprivation therapy (Herr 2023)
  - > Grade Group 5: 81.4% received ADT
  - Grade Group 4: 73.8% received ADT
  - **Grade Group 3**: 51.2% received ADT
  - **Grade Group 2**: 37.2% received ADT
  - **Grade Group 1**: 14.2% received ADT

Often time limited: 6months to 2-3 years

> ~30% of patients present with advanced disease or disease recurrence and patients will be treated with ADT (Batra 2021)



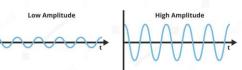


#### Prostate Cancer and ADT

- Side-effects occur within 2 months of initiating ADT (Gray 2005)
- Incidence and severity of side-effects increase with duration of therapy (Grossman 2011)
- Impact may increase with age of patient (Mohile et al., 2009)

Patients vary significantly in expression of ADT side-effect profile

Not well understood.







## ADT and Quality of Life

**Physical** 



Cognitive



Masculinity



Sexual Functioning



NOTE: Metabolic and cardiovascular consequences Emotional



01

## **PHYSICAL**









## Fatigue, Hot Flashes, and Sleep Disturbances

#### FATIGUE (Story 2011, Casey 2012)

- > 40% of men on ADT will experience fatigue
- The 2<sup>nd</sup> most distressing side-effect next to sexual dysfunction

#### **HOT FLASHES** (Frisk 2010)

- 80% of men on ADT will experience hot flashes
- > 27% most bothersome side-effect

#### **SLEEP DISTURBANCE** (Gonzalez et al., 2017)

- > 59% of men report poor sleep
- Hot flashes
- Nocturia



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## Intervention: Fatigue, Hot Flashes and Sleep

#### **FATIGUE**

> EXERCISE (Segal 2003, Yuan 2023)

#### **HOT FLASHES**

- ➤ Hormonal Agents (e.g. megestrol acetate, estrodial) (Jones 2012)
  - ➤ Could interfere with ADT, plus side-effects
- ➤ SSRI (e.g. Escitalopram) and SNRI (e.g. Venlafaxine)
- Neuroleptic agents (e.g. Gabapentin) (Loprinzi 2009)
- ➤ Acupuncture, Meditation/ Relaxation/ Deep Breathing (Jones 2012)
- Behaviour Modification layered sheets, clothing, fans

#### **SLEEP**

- **EXERCISE** (Segal 2003, Yuan 2023)
- CBTi building sleep drive (Meyers et al., 2022)



#### From the Chair

- Loss of engagement in joyful activities
- Loss of motivation
- Reduction in productivity
- Depression (Scott 2021)
- > Reduction in coping and resilience
- > Loss of meaning







### Weight Gain, Muscle Loss, and Loss in BMD

#### **WEIGHT GAIN**

➤ In the 1<sup>st</sup> year on ADT fat mass increase by 10% and lean mass decrease by 3% (Nguyen 2015)

#### LOSS in BMD

Some men will experience reductions in BMD

#### **MUSCLE LOSS**

Men on ADT will experience muscle loss and reduced muscle strength (Storer 2012)









## Intervention: Weight Gain, Muscle Loss, Loss in BMD

#### WEIGHT GAIN AND MUSCLE LOSS

- **EXERCISE** (Algoter 2021, Yuen 2023)
  - Rx: Hypertrophy training (muscle building)
  - Resistant more than aerobic
  - Pre-habilitation and Rehabilitation

#### LOSS in BMD (Poon 2018)

- Calcium, Vitamin D
- Stop smoking
- Biophosphonates and Denosumab





# U2 COGNITIVE





## Cognitive Function

- Research: a history of mixed results
  - ➤ Largest prospective study on ADT and Cognitive Function no effect (Alibhai 2010)
  - ➤ Review 47% to 69% experience impairment in at least one cognitive domain (Nelson 2008)
  - > Strongest evidence (incl. Meta-analysis) for: (Jamadar 2012; McGinty 2014)
    - Losses in verbal & spatial memory
    - Declines in visuospatial processing



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#### From the Chair

- The thought of losing cognitive capacities is extremely anxiety-provoking
  - Conceptualizing Cognitive Dysfunction is like conceptualizing an Electric Shock
  - Dr. Google can be alarmist about ADT and cognition
  - > Patients become hypersensitive to cognitive "mistakes" e.g. wordfinding
  - Can result in significant distress
- > Importance of providing a realistic and balanced expectation for patients
  - University Professor– Mathematician, 12 years on ADT



# SEXUAL FUNCTIONING









#### Loss of Sexual Desire

- > 94% of men on ADT will experience loss of sexual desire (Ng 2022, Kinnaird 2025)
- > The most distressing side-effect of ADT (Casey 2012)



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#### From the Chair

- ➤ Loss of Sexual Fantasy and Sexual Dreams
  - Men think about sex a lot
    - Results vary but it appears that men have sexual thoughts about 19 times per day (Fisher 2012)
    - Not including erotic dreams
  - ➤ Men will describe a void in their lives (loss/grief)







## Erectile Dysfunction and Orgasmic Dysfunction

➤82% suffer Erectile Dysfunction (Wibowo 2019)



- ➤ Delayed or difficulty reaching orgasm (Mazzola et al., 2016)
- ► Loss of ejaculate volume (Elliot et a., 2010)
- ➤ Anorgasmia (Gryzinski et al., 2022)

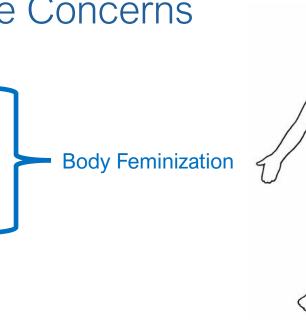


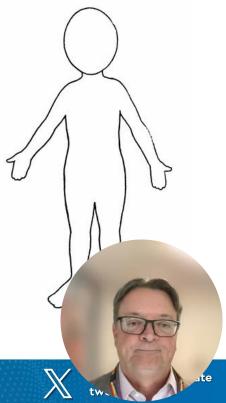


## Body Image Concerns

- Breast growth (15% men on ADT)
- loss of body hair
- genital shrinkage
- loss of muscle mass

(DiLorenzo 2005, Gentili et al., 2019)







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#### From the Chair

- > Interferes with pleasurable and healthy activity
  - Swimming at the beach (No Bondi)
  - Going to the gym
    - Showering becomes problematic



#### INTERVENTION

- Breast Growth RT, Tamoxifen (Bautista-Viz)
- > Compression clothing
- Behavioral adjustment shower at h







## Sexual Functioning – Intimacy Concerns

- 83% 93% Sexual Activity Cessation (Ng 2012)
- Distressing for Partner
- Distressing for Couple
  - Reductions in overall relationship intimacy (Walker 2015)







## Intervention: Sexual Functioning

#### BIO-PSYCHOSOCIAL APPROACH TO SEXUAL REHABILITATION (Wittman 2022)

> Integration of physical, emotional, cognitive and social factors

#### LOSS OF DESIRE

- Hardest to treat
- Acceptance & Adaptation

#### **ERECTILE DYSFUNCTION**

- > PDE5I's, ICI, VED, Surgical **Implants**
- PDE5i's less effective in context > Intimacy Coup of ADT
- Non-Penetrative Sexual Activity

#### **INTIMACY CONCERNS**

Open communication with partner (avoid assumptions)



## 04 MASCULINITY









#### "Male Traits" and ADT

Testosterone is associated with: (Archer 2006, Carre 2015, Can Honk 2010)

- Competitiveness
- Physical Strength
- Sexual Prowess
- Reactive aggression... assertiveness (Vigor)
- Stoic emotional regulation
- Decisiveness
- High energy







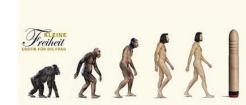
#### "Male Traits" and ADT

ADT is associated with losses in: (Ng 2006, 2012, Oliffe 2006, Donvan 2015, Harrington 2009)

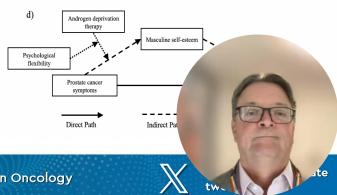
- Competitiveness
- Physical Strength
- Sexual Prowess
- Reactive aggression... assertiveness (Vigor)
- Stoic emotional regulation
- Decisiveness
- High energy



# Intervention: Masculinity



- Psychological Flexibility (Hoffman 2019)
- Broader perspective of masculine identity through Psychological Flexibility (Spendelow 2018, Bowie 2022)
  - ➤ Less effective in PrCa patients on ADT (Hulburt-Williams 2015)
  - Likely due to severity of symptoms



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## 05 EMOTIONAL









## **Emotional Lability**

- Exaggerated changes in emotion (Cherrier 2009, Cary 2014)
  - More irritable or angry

- Becoming Spontaneously Tearful
  - > Difficulty maintaining stoic emotional expression



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#### From the Chair

- > INABILITY TO MAINTAIN STOIC EMOTIONAL EXPRESSION
  - > e.g. Sentimental Movie
  - Public Embarrassment Politician
- Unmanly, shameful

#### INTERVENTION

- >Awareness for potential exaggerated mood
  - >Sensitivity embrace rather than view as weakness/vulnerability (mean
  - ➤ Tearfulness let intimate others know









## Depression

- Research: a history of mixed results (variable methodological rigor, small sample sizes)
  - > 11% depression in the general male population (significant symptoms) (Lee 2015)
  - > 8% to 25% depression in men with prostate cancer (Nead 2021)
  - > 39% depression in men with prostate cancer treated with ADT(Lee 2015, Watson 201



Reduced Sexual Activity/Libido

#### **TOGETHER IN DISCOVERY & CARE**



DSM V DIAGNOSTIC CRITERIA FOR DEPRESSION	ADT SIDE-EFFECT PROFILE		
Low Mood / Irritability	Depression, Emotional Lability		
Loss of Interest/Motivation	Loss of vigor/competitiveness		
Weight Gain or Loss	Weight Gain		
Poor Sleep	Hot Flushes – Sleep Disturbance		
Psychomotor Retardation			
Fatigue/Low Energy	Fatigue/Low Energy		
Diminished Concentration	Cognitive Impairment/Memory		
Indecisiveness			
Worthlessness	Losses In Masculinity		
Suicidality	Elevated In Men On ADT		

Reduced Sexual Activity/Loss of Instin



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DSM V DIAGNOSTIC CRITERIA FOR DEPRESSION		ADT SIDE-EFFECT PROFILE				
Low Mood / Irritability	Depression, Emotional La		ability			
Loss of Interest/Motivation		Loss of vigor/competitiver	ness			
Weight Gain or Loss		Weight Gain				
Poor Sleep	Under the cond	ditions of ADT's	bance			
Psychomotor Retardation	Under the conditions of ADT's physical and cognitive side effect profile men are highly susceptible to depressed mood					
Fatigue/Low Energy						
Diminished Concentration			mory			
Indecisiveness						
Worthlessness		Losses In Masculinity				
Suicidality		Elevated In Men On ADT				
Reduced Sexual Activity/Libido		Reduced Sexual Activity/L	oss of Instin			

## Intervention: Depression

- Anti—Depressant Medications
  - ➤ ADT-based depression may be resistant to anti-depressants (Sountoulides 2013)
  - Better for more severe depression (Fournier 2010)
- Psychosocial Interventions
  - Cognitive Behaviour Therapy effective for depression (Rock 2012)
- **EXERCISE** (Rock 2012, Galvao 2012, Yuan 2023)
  - Targeting physiological etiology



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## **Key Takeaways**

- Importance of an interdisciplinary team
- Invite/include partner
- Provide up-front biopsychosocial education
  - Sexual dysfunction
  - > Fatigue, Hot Flashes, Sleep
  - Body image
  - **Cognitive Function**
  - Mood
- Inquire throughout patient's journey
- Inquire about the partner and couple's journey

- 4) Highlight the importance of EXERCISE
- **Strength and physical function**
- Mental health
- Self image
- **Sexual Function (not including libido)**

Study or subgroup	Weight (%)	Standard mean differen	ce Standard m	ean difference	
	• . ,			0070 01	_
Bourke et al. (2011)	19.4	0.34 (-0.22 to 0.90)			-
Bourke et al. (2014)	38.4	0.44 (0.04 to 0.84)		_	
Segal et al. (2009)	21.6	0.21 (-0.32 to 0.74)		-	
Segal et al. (2009)	20.6	0.39 (-0.15 to 0.93)	_		_
Total (95% CI)	100.0	0.36 (0.11 to 0.61)		•	_,
Heterogeneity: $\chi^2$ =0. Test for overall effect		-1	0 –0.5 0 Favours usual care	0.5 Favours exercise	1.0





## Thank you

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Complete our session survey to enter a raffle for a FREE one-year ISSM membership!

This poll gathers insights to improve sexual health guidelines and resources in prostate cancer care through the ISSM and Movember partnership.



