EAA Andrology Training Centre

Centre Report





Centre identification Center for Men's Health, The Lundquist Institute at Harbor-UCLA Medical Center, Division of Endocrinology, Department of Medicine, Harbor-UCLA Medical Center, 1124 W. Carson Street, Torrance, CA 90502, USA.



**CENTRE REPORT** 

#### **History of Centre**

The Center of Men's Health (Harbor-UCLA Male Reproductive Research Center) was established in 1973 under the current director Professor Ronald Swerdloff. It is located at The Lundquist Institute at the Harbor-UCLA Medical Center. All faculty of the center are faculty of the David Geffen School of Medicine at UCLA. Harbor-UCLA Medical Center, a public hospital serving the indigent population of the South Bay of Los Angeles County, includes the hospital and clinics (both operated by the Los Angeles County Department of Health Services) along with research facilities under the auspices of The Lundquist Institute. Covered under the Center umbrella is the National Institute of Child Health and Human Development funded Contraceptive Clinical Trial Network Center (CCTN) in the Male area and National Male Reproductive Epigenomics Center. The training of Andrology in the United States is not clearly defined and generally takes place in the Division of Urology or the Division of Endocrinology. Andrology unlike Obstetrics and Gynecology is not a recognized sub-specialty and there is no formal program of training or accreditation by the American Society of Andrology or any other accreditation committee. The Center of Men's Health under the direction of Dr. Swerdloff specializes in the training of both basic and clinical andrologists. Andrology is within the Division of Endocrinology and Metabolism and an Andrology Clinic was established in 2014 with participation of urology faculty providing specialized patient care and teaching of trainees by an andrologist and a urologist. The clinical trainees include medical students, rotating medical residents, visiting scientists and endocrinology post-doctoral fellows that complete clinical training in two years after which they are eligible to be board certified in Endocrinology. The clinical trainees can stay on for extra years for research training. Basic research training includes doctoral students, and post-doctoral trainees. Research focuses development of male contraceptives, androgen replacement, development of ocular cohesive tomography for detection of occult sperm in non-obstructive azoospermia, sleep and androgen metabolism, regulation of spermatogenesis, mitochondrial derived peptides and germ cell apoptosis, cilia dyskinesia, non-hormonal male contraception, male reproductive epigenomics and epigenetic inheritance.

#### **Organization of Centre**

The Center of Men's Health (Andrology Center) is directed by Professor Ronald Swerdloff, MD (Past President of the American Society of Andrology (ASA); Recipient of the ASA Distinguished Andrology Award and Endocrine Society Laureate Distinguished Educator Award) . He is assisted by the co-director, Professor Christina Wang, MD (Past President of ASA; Past President, Past Secretary of the International Society of Andrology, ISA; Past chair, and member of the World Health Organization Task Force/Research Group on the Regulation of Male Fertility; Recipient of the Distinguished Andrology Award and Distinguished Service Award from the ASA). They led the Andrology development at our campus and have trained many andrologists from around the world. Dr. Swerdloff is the chief of Endocrinology responsible for administration of the Division and plays a leadership role shared with the director of the training program in endocrinology. He is the Director of the Center for Men's health and the Contraceptive Clinical Trials Network center (CCTN). Dr. Wang is the Medical Director of the Endocrine and Metabolic Research Laboratory (A State and Federal Certified Laboratory), Site Director of the Clinical and Translational Science Institute at The Lundquist, and Principal Investigator of the CCTN male contraceptive trials. Together Dr. Swerdloff and Wang is responsible for clinical research studies in male reproduction (usually have 5 to 10 concurrent studies) and supervised four endocrinology trainees, three research coordinators, a research scientist (Dr. Yanhe Lue) and 5 laboratory technologists and a data management consultant.

Professor Wei Yan, MD, PhD (Recipient of the ASA Young Investigators award, Past Program Chair of ASA Annual meeting and North American Testis Workshop; Past Editor-in-Chief of Biology of Reproduction and recipient of the Society for the Study of Reproduction Outstanding Research Award) is the PI of the NICHD funded Program Grant "Center of Male Reproductive Epigenomics" and several NICHD funded Investigator-Initiated grants. His laboratory has a senior scientist H. Zhang, MD, PhD; three post-doctoral fellow and a PhD student.

Professor Peter Liu, MD, PhD (Current President of the International Society of Andrology, Recipient of the ASA and ISA Young investigators award) is responsible for studies on impact of sleep on hormone and cardiometabolic health. He is also part of the male contraceptive development group.

Please see Organization Chart for other members of the center.

#### **Educational activities**

Clinical Training: The trainees include medical students, residents, fellows and visiting scholars. Didactics include weekly translational endocrinology conference, journal club, didactic series on reproduction, Harbor-UCLA- City of Hope Medical Center Joint Research Conference. Training in Andrology Clinic with Andrologists (Professors Swerdloff, Wang or Liu) and Urologists Professor Jacob Rajfer or Dr. Sriram Eleswarapu). The clinic is a teaching clinic where each patient with an andrology, sexual dysfunction or infertility problem is discussed with an endocrine and a urology fellow with all the physicians in the room. The collaboration between endocrinology and urology provides the foundation of training that enhances the curriculum with professionalism and sensitivity in the care of andrology patients and families; familiarity with contemporary andrological literature and clinical practice guidelines from professional societies; and participation in clinical or basic science research activities in any aspect of male reproductive and sexual medicine. The trainees that are interested in Andrology learn how to do semen analyses using the WHO Laboratory Manual for the Examination and Processing of Human Semen and attends the 4 modules from the Endocrine Society on Hormone Assays including Module 1: Importance of hormone measurements and assay standardization; Module 2: Hormone types and characteristics and features of an ideal assay; Module 3: Methods used to measure hormones and validation/judging the quality of an assay; and Module 4: Factors affecting the interpretation of hormone concentrations specifically for use in diagnosing endocrine disorders. Trainees are required to

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complete Laboratory hands on experience for at least a week; with weekly discussions on assays as relevant; and group discussions with Dr. Wang /Swerdloff on laboratory methods. The trainees are required to take in responsible conduct of research, protecting human participants, and training in harassment and discrimination.

**Research Training:** The trainees include high school students, undergraduate and graduate students and post-doctoral fellows. The basic research training occurs mainly in the Laboratory of Dr. Yan and Dr. Lue. The didactics include the Epigenomics Center seminars with external speakers, weekly laboratory meetings, training session for the Lundquist PhD program and the summer fellowship program. The Epigenomics Center faculty including collaborators at University of California Riverside and University of Texas San Antonio, is responsible for providing eight didactic sessions for the Summer High School Student Fellowship program and 6 teaching sessions for PhD students on reproduction and genetic and epigenetic inheritance. Supervised laboratory training is provided by Drs. Yan, Lue, and Zheng. Former selected trainees include: Shalendar Bhasin (USA), Rebecca Sokol (USA), David Handelsman (Australia), Peter Liu (formerly from Australia, now at Lundquist), Nestor Gonzalez-Cadavid (formerly from Venezuela, now at The Lundquist), Krista Erkkila (Finland), Jonas Ceponis and Indre Ceponiene (Lithuania), K.K Lee and Jason Ng (Hong Kong), Prasanth Surampudi (USA), Jaixin Xing (China) among many others. Current Trainees include: Junior faculty, Fiona Yuen, MD, MS PhD candidate; Fellows: Waleed Butt, Jerry Han, Haeseung Lee, Prativa Rajbhandari; Post-Doctoral Fellows: Kyle Wang, PhD, Hetan Wang, MD, PhD, Sheng Chen, MD, PhD; and Ph D candidate Hayden McSwiggin.

#### **Research activities**

The primary purpose of the Center is to enhance collaboration within the Center members, and with members of other departments at The Lundquist as well as expanding collaborations to other institutions within the US, EAA and throughout the world. In this regard, we have been successful in recruiting Wei Yan, MD, PhD by the Endocrinology (Andrology) team of Swerdloff and Wang resulted in the successful

application of a P-50 grant and the establishment of a NIH, NICHD funded National Male Reproductive Epigenomics Center on The Lundquist Campus in 2020. The Epigenomics Center was moved to The Lundquist from University of Nevada Reno and Professor John McCarrey (University of Texas, San Antonio) and Assistant Professor Qi Chen (University of California, Riverside) were integrated across institutions. Drs. Wang and Swerdloff wrote the clinical project and the projects of Drs. Yan and McCarrey were matched with the clinical project in many aspects. Dr. Wang led the community outreach and education project for the P-50 center grant. A pilot project on Semen Microbiome in response to changes in exercise and nutrition was successfully funded as a pilot grant to Fiona Yuen MD, MS, PhD candidate at The Lundquist Institute and a UCLA Instructor in Medicine within the Epigenomic P-50 Center. Rachelle Bross PhD (Bionutritionist at The Lundquist) was integrated for the clinical nutrition portion and Dr. Harry Rossiter Research Director of the Pulmonary and Exercise Physiology led the exercise portion of the clinical project. Dr. Yan worked with Dr. Rossiter on a separate successful NIH RO1 gratnt(Investigator initiated grant funded by NIH) and with Dr. Khorram from Division of Reproductive Endocrinology on another R01 grant on leiomyoma. Dr. Yan has multiple other collaborations locally, nationally, and internationally including a collaboration with Dr. M. Skinner at Washington State University and Dr. McCarrey on transgenerational transmission of Epigenomic Modulations of Genetic Traits. His large cadre of post-doctoral fellows include many from international institutions. He has strong ties to many Chinese research Institutions growing list of collaborators in many prestigious institutions in China (See Collaboration Figure).

Drs. Swerdloff and Wang continued to collaborate with Dr. Budoff on androgens and cardiac-metabolic- reproductive projects and Drs. Budoff. Drs. Swerdloff, Wang and their senior research fellow and now junior faculty member Dr. Fiona Yuen collaborated with Dr. J Rotter on genetic assessment of changes in outcomes of The T Trial (Peter Snyder PI at U. Penn). Dr. C. Wang (PI of multiple NICHD competitive contracts through the Lundquist Contraceptive Clinical Trial Network Centers (R Swerdloff Center Director) are leading an international study on combined androgen-progestin gel for male contraception. The study sites other than The Lundquist



include the University of Washington co-lead site (S. Page, MD, PhD) and 13 other sub-sites in four continents. The Contraceptive Clinical Trials Network NICHD funding (C. Wang, R. Swerdloff) was renewed for seven additional years during the report period window. Dr. B. Nguyen (Gynecology/ Family Planning at the University of Southern California) was added as a new collaborator to The Lundquist NICHD program bringing new strength to studies on attitudes and needs for male contraception and expertise of a gynecologist in couples' studies. Dr. Wang's hormone and semen measurement laboratory (Endocrine and Metabolic Research Laboratories) continued in an increasing role as Lue, Wang and Swerdloff have developed (through funding by NICHD) new sensitive techniques for study participants home self-assessment of azoospermia and very low sperm counts (oligozoospermia). This Laboratory functions as the central laboratory of all the NICHD funded Male Contraceptive Clinical Trial Network studies and participates in many other studies on environmental effects of pre-natal hormones on the offspring. Drs. Swerdloff, Lue and Wang collaborating with Dr. Park in Bio-engineering at UC Riverside received a NICHD STIR grant and spin off a company to develop a new patented Ocular Cohesive Tomography technology as a non-invasive means of finding occult sperm in the testes of azoospermic infertile patients. Lue and his colleagues are also collaborating with the Institute of Regenerative Medicine at Wake Forest on a project on developing testis organoids for XXY mice and boys. Dr. Peter

Liu leads research on sleep and endocrinology. He participates in the contraceptive studies and collaborates with many Dr. Rotter's team from the Genomics IWI as well in investigators in other outside investigators including Dr. Veldhuis at the Mayo Clinic and Dr. Iranmanesh at the University of Virginia.

#### **Clinical activities**

The patients with andrological problems are seen at the Medical Center's Andrology, Endocrine, Pituitary, Diabetes, Pediatric Endocrinology, Urology and Reproductive Endocrinology Clinics. The Andrology clinic staffed by Andrologists and Urologists is a consultation and teaching clinic. This is where patient seen in other clinic will receive an expert opinion on androgen replacement, sexual dysfunction, and male infertility. Patients with hypogonadism are generally seen and investigated in the Endocrine clinic where approximately 5% of patients have male hypogonadism, infertility and transsexualism. Most of the patients in our Diabetic Clinic have Type 2 diabetes and about 20 % of these have male hypogonadism or erectile dysfunction. Patients with hypogonadism associated with pituitary and hypothalamic problems are investigated and treated in the Pituitary Clinic where about 25 to 35% of the patients have male hypogonadotropic hypogonadism due to pituitary tumors. The Medical Center will support the use of gonadotropins for induction of spermatogenesis with special medication request and justification for a limited number of patients. The hormone analyses for these patients are performed by the Medical Center's Clinical Chemistry Department and most of the specialized tests are sent out to Reference Laboratories. Patients with ambiguous genitalia, cryptorchidism, and micropenis, delayed or precocious puberty are referred to the Pediatric Endocrine Clinic running on the same days as the adult clinics. Harbor-UCLA Medical center has a multidisciplinary transgender medicine group where patients requiring psychological, endocrine or surgical therapy are triaged and directed to the sub-specialty clinics. In the urology clinic (Chief of Urology, Dr. Blumberg) about 7 to 10 % of the subjects present with erectile and sexual dysfunction, hypogonadism, and male infertility. In addition, Dr.

Rajfer and Eleswarapu performs testicular microdissection and sperm extraction from testis for approximately 40 to 50 patients per year with male infertility through their clinics at UCLA but not at The Lundquist Institute at Harbor-UCLA. The infertile couple is usually initially investigated in the Reproductive Endocrinology Clinic (Chief of Reproductive Endocrinology, Dr. Khorram). Because infertility treatment is not covered by insurance and the government sponsored MediCare and MediCal programs in California, patients seen in our public medical clinics have no additional resource and little access to infertility diagnosis and treatment. The Medical Center supports ovulation induction by clomiphene and a very limited ovulation induction with gonadotropins by special request. Thus the wide range of male infertility, genetics of male infertility and ART investigations and treatment are not available through the Medical Center and performed through the private clinics of Drs. Swerdloff, Rajfer and Khorram. The Center's Andrology Laboratory supports semen analyses and preparation of samples for IUI for these private clinics. IVF and ICSI and testicular extraction or epididymal aspiration of sperm are not performed on campus but in associated ART clinics (Dr. Khorram and Rajfer).

#### Name and address of Centre

The Lundquist Institute at Harbor-UCLA Medical Center, Division of Endocrinology, Department of Medicine, Harbor-UCLA Medical Center, 1124 W. Carson Street, Torrance, CA 90502, USA

Type of CentreUniversityUniversity HospitalPrivate CentreOther (please specify)Research Faculty	h Institute in an Academic Hos	spital. Faculty are UCLA
1. Director	Ronald Swerdloff MD	
Academician X	Affiliated Member	Clinical Andrologist
2a. Clinical responsible	Christina Wang, MD	
Academician X	Affiliated Member	Clinical Andrologist
2b. Clinical responsible	Peter Lue, MD, PhD	
Academician	Affiliated Member	Clinical Andrologist X
2c. Clinical responsible	Jacob Rajfer, MD at UC	LA
Academician	Affiliated Member	Clinical Andrologist X
2. Clinical responsible	Sriram Eleswarapu, MI	D, PhD at UCLA
Academician	Affiliated Member	Clinical Andrologist X
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## 3. Scientists

1)	Name Degree Specialty		Wei Yan MD, PhD Epigenetic Inheritance	
Academ	ician	Affili	ated Member	Clinical Andrologist
2)	Name Degree Specialty	-	Yanhe Lue MD Regulation of spermate	ogenesis
Academ	ician	Affili	ated Member	Clinical Andrologist
3)	Name Degree Specialty		Huili Zhang MD, PhD Male reproductive epig	genetics

## Insert any additional staff below (if required)

MD/Biologists/	'Chemists		
1)	Name	Fiona Yuen	
	Degree	MD, MS	
	Specialty	Andrology/Endocrinology/	ogy
	Full time/part time	Full time	
Acade	mician Affil	iated Member	Clinical Andrologist X
2)	Name	Waleed Butt	
2	Degree	MBBS	
	Specialty	Endocrinology trainee	
	Full time/part time	Full time	
Acade	mician Affil	iated Member	Clinical Andrologist
3)	Name	Haeseung Lee	
,	Degree	MD	
	Specialty	Endocrinology trainee	
	Full time/part time	Full time	
Acade	mician Affil	iated Member	Clinical Andrologist
4)	Name	Jerry Han	
-	Degree	MD	
	Specialty	Endocrinology trainee	
	Full time/part time	Full time	

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Acade	mician Affiliated Member Clinical Andrologist
5)	NamePrativa RajbhandariDegreeMDSpecialtyEndocrinology traineeFull time/part timeFull time
Acade	mician Affiliated Member Clinical Andrologist
6)	NameKyle Zhuqing WangDegreePhDSpecialtyMale ReproductionFull time/part timeFull time
Acade	mician Affiliated Member Clinical Andrologist
7)	NameHetan WangDegreeMD, PhDSpecialtyMale ReproductionFull time/part timeFull time
Acade	mician Affiliated Member Clinical Andrologist
8)	NameSheng ChenDegreeMD, PhDSpecialtyMale ReproductionFull time/part timeFull time
Acade	mician Affiliated Member Clinical Andrologist
9)	NameHayden McSwigginDegreeBSSpecialtyMale ReproductionFull time / part timeFull time
Acade	mician Affiliated Member Clinical Andrologist
Insert any add	itional staff below (if required)
Specialists PhD Students	1) NameJacob Rajfer, MD, Urologist2) NameSriram Eleswarapu, MD, PhD Urologist3) NameLaura Hull MBA Data Management Consultant4) Name
ind Students	1) Name
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## Nurses/coordinators

2) Name Elizabeth Dain	
2) Name Elizabeth Ruiz	
3) Name Michael Massone	

Laboratory Technicians

1) Name	Sima Baravarian, PhD
2) Name	Maria Lajoie
3) Name	Kimberly Hernandez

## Administrative Personnel

onnel			
1) Name	Janicia ng		
2) Name			
3) Name			

## 4. Clinical Activity

A. Outpatients: Consultations per year in the last 3 years

	2019	2020	2021
Endocrinology and Diabetes	4698	5009	5440
Pituitary	793	667	674
Andrology	352	284	296

Type of andrological patients in the last years (%) in Andrology/Endocrine/Pituitary	2019	2020	2021
Infertility	7	6	10
Erectile dysfunction	27	34	32
Hypogonadotropic Hypogonadism	26	20	25
Klinefelter	23	28	23
Gynaecomastia	10	7	2
Varicocele			
Cryptorchidism			
Male sex accessory gland infections			
Testicular tumours			
Disorders of gender identity	7	5	8

B. Ultrasound (testis, penile, prostate) \*

	2019	2020	2021
Total	NA	NA	NA
Controls			

\* performed at the Department of Radiology

C. Andrological surgery procedures (Drs. Rajfer and Eleswarapu)

	2019	2020	2021
Testicular biopsies	0	0	1
Micro-Testicular extraction of sperm	12	14	13
Varicocele ligation	0	0	0
Prostate biopsies	0	0	0
BPH	70	72	72
Prostate cancer	0	0	0
Vasectomy	40	20	20
Vaso-vasostomy	0	0	0
Other			

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#### 5. A. Andrology laboratory activity

	2019	2020	2021
Semen analyses	808	714	765
Sperm preparation swim up	20	45	60
Seminal markers			

### 5. B. Andrology laboratory activity

Sperm banking donors	Yes	No	X
Sperm banking cancer patients	Yes	No	X

If yes:			
	2017	2018	2019
Number of samples			

Yes

Yes

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## **5. C.** Histopathologial evaluation of biopsies

### 5. D. Reproductive Hormones Assays

	2019	2020	2021
testosterone	2775	3880	3969
Free testosterone (Equilibrium dialysis)	0	1055	0
SHBG	983	1347	3357
FSH	1973	1471	2274
LH	1386	1636	2040
DHT	2017	2028	100
Estradiol	793	2-48	437

**5. E.** Y chromosome microdeletions according to EAA/EMQN guidelines

Yes

No	Х

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No

No

*If yes* number of tests in the past year

Participation to the EAA quality control scheme?

*If no,* specify if available in another lab of the same hospital

Blood karyotyping

*If no,* specify if available in another lab of the same hospital



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Other genetic tests (please specify) FISH sperm Pre-implantation genetic diagnosis No

### 6. Collaborations with other Clinical Units of the University/Hospital

 IVF Unit
 Yes
 X
 No

 If yes
 please specify: Children, Endocrinology, IVF, Urology, Genetics, Pathology

Urology Clinic	Yes X	No
Endocrine Clinic	Yes X	No
Genetics Lab/Unit	Yes X	No
Paediatric Unit	Yes X	No
Central Hospital Laboratory	Yes X	No
Private Centres	Yes X	No
<i>If yes</i> please specify:		
Dr. Khorram's private clinic		

### 7. Clinical teaching activity

Duration of training (years):

	Number
A: Trainees in the last five years	7
B: Trainees who passed EAA-ESAU\exam for Clinical Andrologist in the last 5 vrs	NA
C: Trainees working in the centre preparing to pass the EAA-ESAU examination	NA
D: PhD Students	2
E: Medical Students	4
F: Other students (MSc) high school	45

### 8. Formal Andrology teaching program

Yes

Years

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2

No

Months

*If yes:* specify duration (years/months):

	Hours of formal teaching per year	Professional training (weeks/months)
Medical Students	30 hours	4 weeks

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PhD Students	400 hours	
Post Graduate basic students		12 months
Trainees Clinical Post- doctoral	200 hours	Three months consult service Three half day clinics per week.
Other degrees (please specify		

#### 9. Research Activity (maximum 1 page)

#### The full list of publications (years 2018 to 2021) are presented at the end of this report.

Male Contraception Development, Androgens Replacement

Thirumalai A, Ceponis J, Amory JK, Swerdloff R, Surampudi V, Liu PY, Bremner WJ, Harvey E, Blithe DL, Lee MS, Hull L, Wang C\*, Page ST. Effects of 28 Days of Oral Dimethandrolone Undecanoate in Healthy Men: A Prototype Male Pill. J Clin Endocrinol Metab. 2019;104(2):423-32.

Wu S, Yuen F, Swerdloff RS, Pak Y, Thirumalai A, Liu PY, Amory JK, Bai F, Hull L, Blithe DL, Anawalt BD, Parman T, Kim K, Lee MS, Bremner WJ, Page ST, Wang C\*. Safety and Pharmacokinetics of Single-Dose Novel Oral Androgen 11beta- Methyl-19-Nortestosterone-17beta-Dodecylcarbonate in Men. The Journal of clinical endocrinology and metabolism. 2019;104(3):629-38

Swerdloff RS, Wang C, White WB, Kaminetsky J, Gittelman MC, Longstreth JA, Dudley RE, Danoff TM. A New Oral Testosterone Undecanoate Formulation Restores Testosterone to Normal Concentrations in Hypogonadal Men. The Journal of Clinical Endocrinology and Metabolism. 2020;105(8):2515-2531.

#### Cilia dyskinesia, epigenetic inheritance

Yuan S, Wang Z, Peng H, Ward SM, Hennig GW, Zheng H, Yan W (2021) Oviductal motile cilia are essential for oocyte pickup, but dispensable for sperm and embryo transport. Proceedings of the National Academy of Sciences USA 118(22) e2102940118.

Chang Z, Qin W, Zheng H, Schegg K, Wang Y, Liu X, Han L, Peri S, McSwiggin H, Wang Z, Peng H, Yuan S, Wu J, Wang Y, Zhu S, Jiang Y, Nie H, Zhou Y, Tang Y\*, Yan W\* (2021) Triptonide is a reversible, nonhormonal male contraceptive agent. Nature Communications 12, 1253 DOI: 10.1038/s41467-021-21517-5

Yuan S, Liu Y, Peng H, Tang C, Hennig GW, Wang Z, Yu T, Klukovich R, Zhang Y, Zheng H, Hess RA, Xu C, Wu J, Yan W\* (2019) Motile cilia of the male reproductive system require miR-34/miR-449 for development and function to generate luminal turbulence. Proceedings of the National Academy of Sciences USA 116(9):3584-3593.

Yu T, Xie Y, Tang C, Wang Y, Yuan S, Zheng H, Yan W\* (2021) Maternal transmission of a paramutant phenotype requires intact DNMT2 functions in the male germline. Biology of Reproduction. <u>https://doi.org/10.1093/biolre/ioab086</u>

#### Male Germ Cell Apoptosis

Jia Y, Swerdloff RS, Lue Y, Dai-Ju J, Surampudi P, Cohen P, Wang C. The IL-27 component EBI-3 and its receptor subunit IL-27R alpha are essential for the cytoprotective action of humanin on male germ cells. Biol Reprod. 2021;104(3):717-30.

#### **Sleep Endocrinology**

Liu PY, Lawrence-Sidebottom D, Piotrowska K, Zhang W, Iranmanesh A, Auchus RJ, Veldhuis JD, Van Dongen HPA. Restoring testosterone and cortisol balance during sleep restriction in men improves metabolic outcomes. J Clin Endocrinol Metab. 2021. (accepted May 25, 2021).

### **10. Research Funding**

Please specify the amount of available funds in the last 3 years and their source (Government, European Union, University, Local Government, Pharmaceutical Industries, Banks, Foundations....)

"Center for Male Reproductive Epigenomics" PI: Wei Yan, Co-I Swerdloff, Wang, Bross, Rossiter NICHD, 1P50HD098593, 09/13/2019-03/31/2024.

Major Goal: To study molecular mechanism underlying intergenerational epigenetic inheritance of metabolic diseases caused by unhealthy diet or lack of physical activities.

"Epitranscriptomic regulation of spermatogenesis and male fertility" PI: Wei Yan, NICHD, R01 HD099924-A1, 09/01/2020-04/30/2025. Major Goal: To investigate how a protein called ALKBH4 controls a specific chemical modification, m6A, on mRNAs, which is known to be essential for the production of good quality of sperm, male fertility and long-term offspring health.

"Sperm tsRNAs and their RNA modifications in diet-induced epigenetic inheritance"

PI: Qi Chen, NIH/NICHD, R01HD092431, 08/09/2017-06/30/2022.

Role: Co-Investigator

Major Goal: To study how sperm-borne tsRNAs and their modifications mediate epigenetic memory.

"Epigenetic Transgenerational Diagnostics for Disease Susceptibility" PIs: Michael Skinner, Wei Yan, Larry Holder, and Paul Winchester. The Templeton Foundation, 2018-2022.

Role: co-PI

Major goal: To investigate the potential that DNA methylation, non-coding RNAs (ncRNAs) and histone modifications can be used as biomarkers of diagnostic value for disease susceptibility and disease-causing direct or ancestral exposures.

"UCLA Clinical and Translational Science Institute" PI Dubinett Role: Wang Site Director at The Lundquist Institute UL1TR001881 07/01/2016- 05/31/2022 The UCLA-CTSI an academic-clinical-community partnership designed to accelerate scientific discoveries and clinical breakthroughs to improve health in the most populous and diverse county in the United States. The mission is to create a

borderless clinical and translational research institute that brings UCLA innovations and resources to bear on the greatest health needs of Los Angeles.

"Contraceptive Development Program NICHD Contraceptive Clinical Trials Network – Male Sites"

NO1 Contract 75N94020D00007 Task Order 75N94020F00001

PI Swerdloff Role: Wang Co-PI 09/27/20 - 9/26/27

The NICHD's Contraceptive Clinical Trials network conducts Phase I, II and III clinical trials of a wide variety of new male contraceptive methods. The objective of

this contract is for CCTN contractors to develop and design new research protocols for men and to conduct male contraceptive clinical trials.

"Clinical Evaluation of Nestorone Gel and Testosterone Gel for Male Contraception."
PI Wang Co-I Swerdloff ,Liu NO1 Contract HHSN275220130024I T ask Order HHSN 2750007 9/15/2015–9/14/2023
This is a phase 2b study to assess the contraceptive efficacy of a combined transdermal Nestorone and *testosterone* gel for 12 months in couples using this method as the sole protection against pregnancy. This study started recruitment in 2018.
"Injectable DMAU for Male Contraception: Safety and Tolerability, Pharmacokinetics and Pharmacodynamics of Single IM or SC DMAU Injection

Pharmacokinetics and Pharmacodynamics of Single IM or SC DMAU Injection Dose Escalation Study in Healthy Male Volunteers " NO1 Contract HHSN275220130024I Task Order HHSN 2750002 01/13/2014–12/31/2023 PI Wang, Co-I Swerdloff, Liu This phase 1 study will administer single IM or SC injections of DMAU (a potential male contraceptive) in healthy men to assess safety tolerability, pharmacokinetics, and suppression of endogenous gonadotropins and sperm output in healthy men.

"Novel OCT Technology for Detection of Occult Sperm in the Testes in Non-Obstructed Azoospermia" PI Swerdloff Co-I: Lue ,Wang NIH/NICHD R41 HD102275 05/01/2020-4/30/2022 This project aims to develop OCT equipment to retrieve sperm in testis of men with

non-obstructive azoospermia



#### **CENTRE PHOTOS**



Ronald Swerdloff, MD Center Director



Peter Liu, MD, PhD Andrologist



Christina Wang, Center Co-Director



Yanhe Lue, MD Scientist/Investigator



Wei Yan, MD, PhD Co-Director



Sriram Eleswarapu Urology at UCLA



## FULL LIST OF PUBLICATIONS (with IF) of staff members from the last 3 years

Please use these URL links to get more recent publication of the main investigators of

our center

Swerdloff

http://www.ncbi.nlm.nih.gov/sites/myncbi/ronald.swerdloff.1/bibliography/47782158/public/?sort=date&direction=ascending

Wang

http://www.ncbi.nlm.nih.gov/sites/myncbi/christina.wang.2/bibliography/47778439/public/?sort=date&direction=descending

Yan

https://www.ncbi.nlm.nih.gov/myncbi/wei.yan.2/bibliography/public/

Liu

https://www.ncbi.nlm.nih.gov/sites/myncbi/peter.liu.1/bibliography/47766701/publ ic/

Publications are arranged in order of the appearance of the Andrology Center investigators. Duplicates are not presented under the other authors' names

### Peter Liu, M.D., Ph.D.

O'Byrne NA, Yuen F, Niaz W, Liu PY. Sleep and the Testis. Curr Opin Endocr Metab Res. 2021 Jun;18:83-93. doi: 10.1016/j.coemr.2021.03.002. Epub 2021 Mar 11. PubMed PMID: 33937581; PubMed Central PMCID: PMC8087280.

Liu PY, Lawrence-Sidebottom D, Piotrowska K, Zhang W, Iranmanesh A, Auchus RJ, Veldhuis JD, Van Dongen HPA. Clamping Cortisol and Testosterone Mitigates the Development of Insulin Resistance during Sleep Restriction in Men. J Clin Endocrinol Metab. 2021 May 27;. doi: 10.1210/clinem/dgab375. [Epub ahead of print] PubMed PMID: 34043794.

Pengo MF, Steier J, Parati G. The ANDANTE Project: A Worldwide Individual Data Meta-Analysis of the Effect of Sleep Apnea Treatment on Blood Pressure. Arch Bronconeumol. 2021 May 10;. doi: 10.1016/j.arbres.2021.05.002. [Epub ahead of print] PubMed PMID: 34088534.

Liu PY, Irwin MR, Krueger JM, Gaddameedhi S, Van Dongen HPA. Night shift schedule alters endogenous regulation of circulating cytokines. Neurobiol Sleep Circadian Rhythms. 2021 May;10:100063. doi: 10.1016/j.nbscr.2021.100063. eCollection 2021 May. PubMed PMID: 33748539; PubMed Central PMCID: PMC7970107.

Mok Y, Melehan KL, Phillips CL, Yee BJ, Miller C, Grunstein RR, Bartlett D, Liu PY, Wong KK, Hoyos CM. Does CPAP treat depressive symptoms in individuals with OSA? An

analysis of two 12-week randomized sham CPAP-controlled trials. Sleep Med. 2020 Sep;73:11-14. doi: 10.1016/j.sleep.2020.04.021. Epub 2020 May 4. PubMed PMID: 32769027.

Liu PY, Takahashi PY, Yang RJ, Iranmanesh A, Veldhuis JD. Age and time-of-day differences in the hypothalamo-pituitary-testicular, and adrenal, response to total overnight sleep deprivation. Sleep. 2020 Jul 13;43(7). doi: 10.1093/sleep/zsaa008. PubMed PMID: 31993665; PubMed Central PMCID: PMC7355405.

Roelfsema F, Liu PY, Yang R, Takahashi P, Veldhuis JD. Interleukin-2 drives cortisol secretion in an age-, dose-, and body composition-dependent way. Endocr Connect. 2020 Jul;9(7):637-648. doi: 10.1530/EC-20-0211. PubMed PMID: 32520721; PubMed Central PMCID: PMC7424344.

Seimon RV, Wild-Taylor AL, McClintock S, Harper C, Gibson AA, Johnson NA, Fernando HA, Markovic TP, Center JR, Franklin J, Liu PY, Grieve SM, Lagopoulos J, Caterson ID, Byrne NM, Sainsbury A. 3-Year effect of weight loss via severe versus moderate energy restriction on body composition among postmenopausal women with obesity - the TEMPO Diet Trial. Heliyon. 2020 Jun;6(6):e04007. doi: 10.1016/j.heliyon.2020.e04007. eCollection 2020 Jun. PubMed PMID: 32613096; PubMed Central PMCID: PMC7322133.

Roelfsema F, Liu PY, Takahashi PY, Yang RJ, Veldhuis JD. Dynamic Interactions Between LH and Testosterone in Healthy Community-Dwelling Men: Impact of Age and Body Composition. J Clin Endocrinol Metab. 2020 Mar 1;105(3). doi: 10.1210/clinem/dgz246. PubMed PMID: 31790144; PubMed Central PMCID: PMC7025815.

Seimon RV, Wild-Taylor AL, Keating SE, McClintock S, Harper C, Gibson AA, Johnson NA, Fernando HA, Markovic TP, Center JR, Franklin J, Liu PY, Grieve SM, Lagopoulos J, Caterson ID, Byrne NM, Sainsbury A. Effect of Weight Loss via Severe vs Moderate Energy Restriction on Lean Mass and Body Composition Among Postmenopausal Women With Obesity: The TEMPO Diet Randomized Clinical Trial. JAMA Netw Open. 2019 Oct 2;2(10):e1913733. doi: 10.1001/jamanetworkopen.2019.13733. PubMed PMID: 31664441; PubMed Central PMCID: PMC6824325.

Hoyos CM, Murugan SM, Melehan KL, Yee BJ, Phillips CL, Killick R, Cayanan EA, Wong KK, Liu PY, Grunstein RR, Marshall NS. Dose-dependent effects of continuous positive airway pressure for sleep apnea on weight or metabolic function: Individual patient-level clinical trial meta-analysis. J Sleep Res. 2019 Oct;28(5):e12788. doi: 10.1111/jsr.12788. Epub 2018 Nov 19. PubMed PMID: 30450787.

Liu PY. A Clinical Perspective of Sleep and Andrological Health: Assessment, Treatment Considerations, and Future Research. J Clin Endocrinol Metab. 2019 Oct 1;104(10):4398-4417. doi: 10.1210/jc.2019-00683. Review. PubMed PMID: 31042277; PubMed Central PMCID: PMC6735730.

Theorell-Haglöw J, Hoyos CM, Phillips CL, Yee BJ, Melehan KL, Liu PY, Cistulli PA, Grunstein RR. Associations Between Obstructive Sleep Apnea and Measures of Arterial

#### Book I

Stiffness. J Clin Sleep Med. 2019 Feb 15;15(2):201-206. doi: 10.5664/jcsm.7616. PubMed PMID: 30736873; PubMed Central PMCID: PMC6374088.

Roelfsema F, Yang RJ, Liu PY, Takahashi PY, Veldhuis JD. Feedback on LH in Testosterone-Clamped Men Depends on the Mode of Testosterone Administration and Body Composition. J Endocr Soc. 2019 Jan 1;3(1):235-249. doi: 10.1210/js.2018-00317. eCollection 2019 Jan 1. PubMed PMID: 30623162; PubMed Central PMCID: PMC6320245.

Zhang W, Piotrowska K, Chavoshan B, Wallace J, Liu PY. Sleep Duration Is Associated With Testis Size in Healthy Young Men. J Clin Sleep Med. 2018 Oct 15;14(10):1757-1764. doi: 10.5664/jcsm.7390. PubMed PMID: 30353813; PubMed Central PMCID: PMC6175801.

Theorell-Haglöw J, Hoyos CM, Phillips CL, Yee BJ, Herrmann M, Brennan-Speranza TC, Grunstein RR, Liu PY. Changes of vitamin D levels and bone turnover markers after CPAP therapy: a randomized sham-controlled trial. J Sleep Res. 2018 Aug;27(4):e12606. doi: 10.1111/jsr.12606. Epub 2017 Sep 25. PubMed PMID: 28944524.

### Yan He Lue, M.D.

Yen K, Mehta HH, Kim SJ, Lue Y, Hoang J, Guerrero N, Port J, Bi Q, Navarrete G, Brandhorst S, Lewis KN, Wan J, Swerdloff R, Mattison JA, Buffenstein R, Breton CV, Wang C, Longo V, Atzmon G, Wallace D, Barzilai N, Cohen P. The mitochondrial derived peptide humanin is a regulator of lifespan and healthspan. Aging (Albany NY). 2020 Jun 23;12(12):11185-11199. doi: 10.18632/aging.103534. Epub 2020 Jun 23. PubMed PMID: 32575074; PubMed Central PMCID: PMC7343442.

Deebel NA, Galdon G, Zarandi NP, Stogner-Underwood K, Howards S, Lovato J, Kogan S, Atala A, Lue Y, Sadri-Ardekani H. Age-related presence of spermatogonia in patients with Klinefelter syndrome: a systematic review and meta-analysis. Hum Reprod Update. 2020 Jan 1;26(1):58-72. doi: 10.1093/humupd/dmz038. PubMed PMID: 31822886.

Jia Y, Lue Y, Swerdloff RS, Lasky JL, Panosyan EH, Dai-Ju J, Wang C. The humanin analogue (HNG) prevents temozolomide-induced male germ cell apoptosis and other adverse effects in severe combined immuno-deficiency (SCID) mice bearing human medulloblastoma. Exp Mol Pathol. 2019 Aug;109:42-50. doi: 10.1016/j.yexmp.2019.104261. Epub 2019 May 11. PubMed PMID: 31085184; PubMed Central PMCID: PMC6680022.

Lue Y, Gao C, Swerdloff R, Hoang J, Avetisyan R, Jia Y, Rao M, Ren S, Atienza V, Yu J, Zhang Y, Chen M, Song Y, Wang Y, Wang C. Humanin analog enhances the protective effect of dexrazoxane against doxorubicin-induced cardiotoxicity. Am J Physiol Heart Circ Physiol. 2018 Sep 1;315(3):H634-H643. doi: 10.1152/ajpheart.00155.2018. Epub 2018 May 18. PubMed PMID: 29775411; PubMed Central PMCID: PMC6734085.

#### Ronald Swerdloff, MD/ Christina Wang, MD

Thirumalai A, Ceponis J, Amory JK, Swerdloff R, Surampudi V, Liu PY, Bremner WJ, Harvey E, Blithe DL, Lee MS, Hull L, Wang C, Page ST. Effects of 28 Days of Oral Dimethandrolone Undecanoate in Healthy Men: A Prototype Male Pill. J Clin Endocrinol

Metab. 2019 Feb 1;104(2):423-432. doi: 10.1210/jc.2018-01452. PubMed PMID: 30252061; PubMed Central PMCID: PMC6306388.

Nguyen BT, Yuen F, Farrant M, Thirumalai A, Fernando F, Amory JK, Swerdloff RS, Anawalt BD, Blithe DL, Long JE, Liu PY, Page ST, Wang C. Acceptability of the oral hormonal male contraceptive prototype, 11β-methyl-19-nortestosterone dodecylcarbonate (11β-MNTDC), in a 28-day placebo-controlled trial. Contraception. 2021 Jun 18;. doi: 10.1016/j.contraception.2021.06.009. [Epub ahead of print] PubMed PMID: 34153318.

Yuen F, Thirumalai A, Fernando FA, Swerdloff RS, Liu PY, Pak Y, Hull L, Bross R, Blithe DL, Long JE, Page ST, Wang C. Comparison of metabolic effects of the progestational androgens dimethandrolone undecanoate and 11β-MNTDC in healthy men. Andrology. 2021 Apr 28;. doi: 10.1111/andr.13025. [Epub ahead of print] PubMed PMID: 33908182.

Jia Y, Swerdloff RS, Lue Y, Dai-Ju J, Surampudi P, Cohen P, Wang C. The IL-27 component EBI-3 and its receptor subunit IL-27Rα are essential for the cytoprotective action of humanin on male germ cells<sup>†</sup>. Biol Reprod. 2021 Mar 11;104(3):717-730. doi: 10.1093/biolre/ioaa225. PubMed PMID: 33330922.

Garcia JM, Biller BMK, Korbonits M, Popovic V, Luger A, Strasburger CJ, Chanson P, Swerdloff R, Wang C, Fleming RR, Cohen F, Ammer N, Mueller G, Kelepouris N, Strobl F, Ostrow V, Yuen KCJ. Sensitivity and specificity of the macimorelin test for diagnosis of AGHD. Endocr Connect. 2021 Jan;10(1):76-83. doi: 10.1530/EC-20-0491. PubMed PMID: 33320108; PubMed Central PMCID: PMC7923131.

Thirumalai A, Yuen F, Amory JK, Hoofnagle AN, Swerdloff RS, Liu PY, Long JE, Blithe DL, Wang C, Page ST. Dimethandrolone Undecanoate, a Novel, Nonaromatizable Androgen, Increases P1NP in Healthy Men Over 28 Days. J Clin Endocrinol Metab. 2021 Jan 1;106(1):e171-e181. doi: 10.1210/clinem/dgaa761. PubMed PMID: 33090208; PubMed Central PMCID: PMC7765650.

Swerdloff R, Wang C. Reflections on the T Trials. Andrology. 2020 Nov;8(6):1512-1518. doi: 10.1111/andr.12901. Epub 2020 Oct 17. Review. PubMed PMID: 32902162.

Swerdloff RS, Wang C, White WB, Kaminetsky J, Gittelman MC, Longstreth JA, Dudley RE, Danoff TM. A New Oral Testosterone Undecanoate Formulation Restores Testosterone to Normal Concentrations in Hypogonadal Men. J Clin Endocrinol Metab. 2020 Aug 1;105(8). doi: 10.1210/clinem/dgaa238. PubMed PMID: 32382745; PubMed Central PMCID: PMC7282712.

Shaikh K, Ellenberg SS, Nakanishi R, Snyder PJ, Lee J, Wenger NK, Lewis CE, Swerdloff RS, Preston P, Hamal S, Stephens-Sheilds A, Bhasin S, Cherukuri L, Cauley JA, Crandall JP, Cunningham GR, Ensrud KE, Matsumoto AM, Molich ME, Alla VM, Birudaraju D, Nezarat N, Rai K, Almeida S, Roy SK, Sheikh M, Trad G, Budoff MJ. Biomarkers and Noncalcified Coronary Artery Plaque Progression in Older Men Treated With Testosterone. J Clin Endocrinol Metab. 2020 Jul 1;105(7). doi: 10.1210/clinem/dgz242. PubMed PMID: 31784747; PubMed Central PMCID: PMC7209773.

Nguyen BT, Farrant MT, Anawalt BD, Yuen F, Thirumalai A, Amory JK, Swerdloff RS, Bremner WJ, Liu PY, Blithe DL, Page ST, Wang C. Acceptability of oral dimethandrolone undecanoate in a 28-day placebo-controlled trial of a hormonal male contraceptive prototype. Contraception. 2020 Jul;102(1):52-57. doi: 10.1016/j.contraception.2020.04.006. Epub 2020 Apr 13. PubMed PMID: 32298717; PubMed Central PMCID: PMC7287214.

Yuen F, Nguyen BT, Swerdloff RS, Wang C. Continuing the search for a hormonal male contraceptive. Best Pract Res Clin Obstet Gynaecol. 2020 Jul;66:83-94. doi: 10.1016/j.bpobgyn.2020.02.003. Epub 2020 Feb 19. Review. PubMed PMID: 32197832; PubMed Central PMCID: PMC7375909.

Swerdloff RS, Dudley RE. A new oral testosterone undecanoate therapy comes of age for the treatment of hypogonadal men. Ther Adv Urol. 2020 Jan-Dec;12:1756287220937232. doi: 10.1177/1756287220937232. eCollection 2020 Jan-Dec. PubMed PMID: 32655691; PubMed Central PMCID: PMC7328356.

Yuen F, Thirumalai A, Pham C, Swerdloff RS, Anawalt BD, Liu PY, Amory JK, Bremner WJ, Dart C, Wu H, Hull L, Blithe DL, Long J, Wang C, Page ST. Daily Oral Administration of the Novel Androgen 11β-MNTDC Markedly Suppresses Serum Gonadotropins in Healthy Men. J Clin Endocrinol Metab. 2020 Mar 1;105(3). doi: 10.1210/clinem/dgaa032. PubMed PMID: 31976519; PubMed Central PMCID: PMC7049261.

Cunningham GR, Ellenberg SS, Bhasin S, Matsumoto AM, Parsons JK, Preston P, Cauley JA, Gill TM, Swerdloff RS, Wang C, Ensrud KE, Lewis CE, Pahor M, Crandall JP, Molitch ME, Cifelli D, Basaria S, Diem SJ, Stephens-Shields AJ, Hou X, Snyder PJ. Prostate-Specific Antigen Levels During Testosterone Treatment of Hypogonadal Older Men: Data from a Controlled Trial. J Clin Endocrinol Metab. 2019 Dec 1;104(12):6238-6246. doi: 10.1210/jc.2019-00806. PubMed PMID: 31504596; PubMed Central PMCID: PMC6823728.

Anawalt BD, Roth MY, Ceponis J, Surampudi V, Amory JK, Swerdloff RS, Liu PY, Dart C, Bremner WJ, Sitruk-Ware R, Kumar N, Blithe DL, Page ST, Wang C. Combined nestorone-testosterone gel suppresses serum gonadotropins to concentrations associated with effective hormonal contraception in men. Andrology. 2019 Nov;7(6):878-887. doi: 10.1111/andr.12603. Epub 2019 Apr 10. PubMed PMID: 30969032; PubMed Central PMCID: PMC6768743.

Yuen F, Wu S, Thirumalai A, Swerdloff RS, Page ST, Liu PY, Dart C, Wu H, Blithe DL, Sitruk-Ware R, Long J, Bai F, Hull L, Bremner WJ, Anawalt BD, Wang C. Preventing secondary exposure to women from men applying a novel nestorone/testosterone contraceptive gel. Andrology. 2019 Mar;7(2):235-243. doi: 10.1111/andr.12577. Epub 2018 Dec 16. PubMed PMID: 30556332; PubMed Central PMCID: PMC6422752.

Garcia JM, Biller BMK, Korbonits M, Popovic V, Luger A, Strasburger CJ, Chanson P, Medic-Stojanoska M, Schopohl J, Zakrzewska A, Pekic S, Bolanowski M, Swerdloff R, Wang C, Blevins T, Marcelli M, Ammer N, Sachse R, Yuen KCJ. Macimorelin as a Diagnostic Test for Adult GH Deficiency. J Clin Endocrinol Metab. 2018 Aug 1;103(8):3083-3093. doi: 10.1210/jc.2018-00665. PubMed PMID: 29860473.

### **Christina Wang**

Wang C, Yuen F, Swerdloff R. Extrahypothalamic ER Alpha Are Required for Testosterone Effects on Physical Activity and Fat Mass in Mice. Endocrinology. 2021 Jun 1;162(6). doi: 10.1210/endocr/bqab075. PubMed PMID: 33842977.

Wright MJ, Monti MM, Lutkenhoff ES, Hardy DJ, Litvin PY, Kelly DF, Guskiewicz K, Cantu RC, Vespa PM, Hovda DA, Lopez WD, Wang C, Swerdloff R, Fuster JM. Memory in repeat sports-related concussive injury and single-impact traumatic brain injury. Brain Inj. 2020 Oct 14;34(12):1666-1673. doi: 10.1080/02699052.2020.1825806. Epub 2020 Sep 29. PubMed PMID: 32990043.

Day DB, Collett BR, Barrett ES, Bush NR, Swan SH, Wang C, Sathyanarayana S. Prenatal sex hormones and behavioral outcomes in children. Psychoneuroendocrinology. 2020 Mar;113:104547. doi: 10.1016/j.psyneuen.2019.104547. Epub 2019 Dec 24. PubMed PMID: 31901731; PubMed Central PMCID: PMC7759302.

Redmon JB, Drobnis EZ, Sparks A, Wang C, Swan SH. Semen and reproductive hormone parameters in fertile men with and without varicocele. Andrologia. 2019 Nov;51(10):e13407. doi: 10.1111/and.13407. Epub 2019 Aug 25. PubMed PMID: 31448444; PubMed Central PMCID: PMC6819219.

Stephens-Shields AJ, Wang C, Preston P, Snyder PJ, Swerdloff RS. Clinically Meaningful Change in Sexual Desire in the Psychosexual Daily Questionnaire in Older Men from the TTrials. J Sex Med. 2019 Jul;16(7):951-953. doi: 10.1016/j.jsxm.2019.04.003. Epub 2019 May 14. PubMed PMID: 31101538; PubMed Central PMCID: PMC7359814.

Kaminetsky JC, McCullough A, Hwang K, Jaffe JS, Wang C, Swerdloff RS. A 52-Week Study of Dose Adjusted Subcutaneous Testosterone Enanthate in Oil Self-Administered via Disposable Auto-Injector. J Urol. 2019 Mar;201(3):587-594. doi: 10.1016/j.juro.2018.09.057. PubMed PMID: 30296416.

Barrett ES, Mbowe O, Thurston SW, Butts S, Wang C, Nguyen R, Bush N, Redmon JB, Sheshu S, Swan SH, Sathyanarayana S. Predictors of Steroid Hormone Concentrations in Early Pregnancy: Results from a Multi-Center Cohort. Matern Child Health J. 2019 Mar;23(3):397-407. doi: 10.1007/s10995-018-02705-0. PubMed PMID: 30659461; PubMed Central PMCID: PMC6397082.

Clark RV, Wald JA, Swerdloff RS, Wang C, Wu FCW, Bowers LD, Matsumoto AM. Large divergence in testosterone concentrations between men and women: Frame of reference for elite athletes in sex-specific competition in sports, a narrative review. Clin Endocrinol (Oxf). 2019 Jan;90(1):15-22. doi: 10.1111/cen.13840. Epub 2018 Sep 27. Review. PubMed PMID: 30136295.

Ceponis J, Swerdloff R, Leung A, Hull L, Bai F, Longstreth J, Dudley R, Danoff T, Wang C. Accurate measurement of androgen after androgen esters: problems created by ex vivo esterase effects and LC-MS/MS interference. Andrology. 2019 Jan;7(1):42-52. doi:

#### Book I

10.1111/andr.12554. Epub 2018 Oct 21. PubMed PMID: 30345711; PubMed Central PMCID: PMC6519384.

Bhasin S, Ellenberg SS, Storer TW, Basaria S, Pahor M, Stephens-Shields AJ, Cauley JA, Ensrud KE, Farrar JT, Cella D, Matsumoto AM, Cunningham GR, Swerdloff RS, Wang C, Lewis CE, Molitch ME, Barrett-Connor E, Crandall JP, Hou X, Preston P, Cifelli D, Snyder PJ, Gill TM. Effect of testosterone replacement on measures of mobility in older men with mobility limitation and low testosterone concentrations: secondary analyses of the Testosterone Trials. Lancet Diabetes Endocrinol. 2018 Nov;6(11):879-890. doi: 10.1016/S2213-8587(18)30171-2. PubMed PMID: 30366567; PubMed Central PMCID: PMC6816466.

Dyar KA, Lutter D, Artati A, Ceglia NJ, Liu Y, Armenta D, Jastroch M, Schneider S, de Mateo S, Cervantes M, Abbondante S, Tognini P, Orozco-Solis R, Kinouchi K, Wang C, Swerdloff R, Nadeef S, Masri S, Magistretti P, Orlando V, Borrelli E, Uhlenhaut NH, Baldi P, Adamski J, Tschöp MH, Eckel-Mahan K, Sassone-Corsi P. Atlas of Circadian Metabolism Reveals System-wide Coordination and Communication between Clocks. Cell. 2018 Sep 6;174(6):1571-1585.e11. doi: 10.1016/j.cell.2018.08.042. PubMed PMID: 30193114; PubMed Central PMCID: PMC6501776.

Jones L, Wells K, Lin HJ, Wang C, Alo AK, Williams P, Jones F, Dickson PI, Han S, Pardo D, Norris K, Jones A, Wright A, Young K, Rotter JI. Community Partnership in Precision Medicine: Themes from a Community Engagement Conference. Ethn Dis. 2018;28(Suppl 2):503-510. doi: 10.18865/ed.28.S2.503. eCollection 2018. PubMed PMID: 30202204; PubMed Central PMCID: PMC6128349.

Wang C, Stephens-Shields AJ, DeRogatis LR, Cunningham GR, Swerdloff RS, Preston P, Cella D, Snyder PJ, Gill TM, Bhasin S, Matsumoto AM, Rosen RC. Validity and Clinically Meaningful Changes in the Psychosexual Daily Questionnaire and Derogatis Interview for Sexual Function Assessment: Results From the Testosterone Trials. J Sex Med. 2018 Jul;15(7):997-1009. doi: 10.1016/j.jsxm.2018.05.008. PubMed PMID: 29960633; PubMed Central PMCID: PMC6435333.

#### Wei Yan, MD, PhD

Suzuki S, McCarrey JR, Hermann BP. Differential RA responsiveness among subsets of mouse late progenitor spermatogonia. Reproduction. 2021 May 5;161(6):645-655. doi: 10.1530/REP-21-0031. PubMed PMID: 33835049; PubMed Central PMCID: PMC8105290.

Suzuki S, McCarrey JR, Hermann BP. An mTORC1-dependent switch orchestrates the transition between mouse spermatogonial stem cells and clones of progenitor spermatogonia. Cell Rep. 2021 Feb 16;34(7):108752. doi: 10.1016/j.celrep.2021.108752. PubMed PMID: 33596419; PubMed Central PMCID: PMC7980622.

Wang Y, Yamauchi Y, Wang Z, Zheng H, Yanagimachi R, Ward MA, Yan W. Both Cauda and Caput Epididymal Sperm Are Capable of Supporting Full-Term Development in FVB and CD-1 Mice. Dev Cell. 2020 Dec 21;55(6):675-676. doi: 10.1016/j.devcel.2020.11.022. PubMed PMID: 33352141; PubMed Central PMCID: PMC7799177.

Zhang X, Trebak F, Souza LAC, Shi J, Zhou T, Kehoe PG, Chen Q, Feng Earley Y. Small RNA modifications in Alzheimer's disease. Neurobiol Dis. 2020 Nov;145:105058. doi: 10.1016/j.nbd.2020.105058. Epub 2020 Aug 21. PubMed PMID: 32835860; PubMed Central PMCID: PMC7572745.

Cheng K, Chen IC, Cheng CE, Mutoji K, Hale BJ, Hermann BP, Geyer CB, Oatley JM, McCarrey JR. Unique Epigenetic Programming Distinguishes Regenerative Spermatogonial Stem Cells in the Developing Mouse Testis. iScience. 2020 Oct 23;23(10):101596. doi: 10.1016/j.isci.2020.101596. eCollection 2020 Oct 23. PubMed PMID: 33083754; PubMed Central PMCID: PMC7552105.

Bae B, Gruner HN, Lynch M, Feng T, So K, Oliver D, Mastick GS, Yan W, Pieraut S, Miura P. Elimination of Calm1 long 3'-UTR mRNA isoform by CRISPR-Cas9 gene editing impairs dorsal root ganglion development and hippocampal neuron activation in mice. RNA. 2020 Oct;26(10):1414-1430. doi: 10.1261/rna.076430.120. Epub 2020 Jun 10. PubMed PMID: 32522888; PubMed Central PMCID: PMC7491327.

Lewon M, Wang Y, Peters C, Peterson M, Zheng H, Wang Z, Hayes L, Yan W. Assessment of operant learning and memory in mice born through ICSI. Hum Reprod. 2020 Sep 1;35(9):2058-2071. doi: 10.1093/humrep/deaa167. PubMed PMID: 32766772; PubMed Central PMCID: PMC7485617.

Matzuk MM, Georg GI, Yan W. A special issue on contraceptive development: past, present, and future. Biol Reprod. 2020 Aug 4;103(2):145-146. doi: 10.1093/biolre/ioaa120. PubMed PMID: 32701150; PubMed Central PMCID: PMC7401340.

Wang Z, Wang Y, Wang S, Gorzalski AJ, McSwiggin H, Yu T, Castaneda-Garcia K, Prince B, Wang H, Zheng H, Yan W. Efficient genome editing by CRISPR-Mb3Cas12a in mice. J Cell Sci. 2020 May 11;133(9). doi: 10.1242/jcs.240705. PubMed PMID: 32393674; PubMed Central PMCID: PMC7240296.

Tang C, Xie Y, Yu T, Liu N, Wang Z, Woolsey RJ, Tang Y, Zhang X, Qin W, Zhang Y, Song G, Zheng W, Wang J, Chen W, Wei X, Xie Z, Klukovich R, Zheng H, Quilici DR, Yan W. m6A-dependent biogenesis of circular RNAs in male germ cells. Cell Res. 2020 Mar;30(3):211-228. doi: 10.1038/s41422-020-0279-8. Epub 2020 Feb 11. PubMed PMID: 32047269; PubMed Central PMCID: PMC7054367.

Wang Z, Xie Y, Wang Y, Morris D, Wang S, Oliver D, Yuan S, Zayac K, Bloomquist S, Zheng H, Yan W. X-linked miR-506 family miRNAs promote FMRP expression in mouse spermatogonia. EMBO Rep. 2020 Jan 7;21(1):e49024. doi: 10.15252/embr.201949024. Epub 2019 Dec 6. PubMed PMID: 31808593; PubMed Central PMCID: PMC6944911.

Wang HT, Tong X, Zhang ZX, Sun YY, Yan W, Xu ZM, Fu WN. MYCT1 represses apoptosis of laryngeal cancerous cells through the MAX/miR-181a/NPM1 pathway. FEBS J. 2019 Oct;286(19):3892-3908. doi: 10.1111/febs.14942. Epub 2019 Jun 13. PubMed PMID: 31152622. Yan W, Clarke H. New horizons in reproductive biology: a special issue. Biol Reprod. 2019 Sep 1;101(3):513. doi: 10.1093/biolre/ioz138. PubMed PMID: 31350845.

Sadler-Riggleman I, Klukovich R, Nilsson E, Beck D, Xie Y, Yan W, Skinner MK. Epigenetic transgenerational inheritance of testis pathology and Sertoli cell epimutations: generational origins of male infertility. Environ Epigenet. 2019 Jul;5(3):dvz013. doi: 10.1093/eep/dvz013. eCollection 2019 Jul. PubMed PMID: 31528361; PubMed Central PMCID: PMC6736068.

Wang Z, Yan W. Inflammation induced by faulty replication during embryonic development causes skewed sex ratio. Biol Reprod. 2019 Aug 1;101(2):259-261. doi: 10.1093/biolre/ioz104. PubMed PMID: 31189185; PubMed Central PMCID: PMC6736220.

Wang Z, McSwiggin H, Newkirk SJ, Wang Y, Oliver D, Tang C, Lee S, Wang S, Yuan S, Zheng H, Ye P, An W, Yan W. Insertion of a chimeric retrotransposon sequence in mouse Axin1 locus causes metastable kinky tail phenotype. Mob DNA. 2019;10:17. doi: 10.1186/s13100-019-0162-7. eCollection 2019. PubMed PMID: 31073336; PubMed Central PMCID: PMC6500023.

Yuan S, Liu Y, Peng H, Tang C, Hennig GW, Wang Z, Wang L, Yu T, Klukovich R, Zhang Y, Zheng H, Xu C, Wu J, Hess RA, Yan W. Motile cilia of the male reproductive system require miR-34/miR-449 for development and function to generate luminal turbulence. Proc Natl Acad Sci U S A. 2019 Feb 26;116(9):3584-3593. doi: 10.1073/pnas.1817018116. Epub 2019 Jan 18. PubMed PMID: 30659149; PubMed Central PMCID: PMC6397547.

Klukovich R, Nilsson E, Sadler-Riggleman I, Beck D, Xie Y, Yan W, Skinner MK. Environmental Toxicant Induced Epigenetic Transgenerational Inheritance of Prostate Pathology and Stromal-Epithelial Cell Epigenome and Transcriptome Alterations: Ancestral Origins of Prostate Disease. Sci Rep. 2019 Feb 18;9(1):2209. doi: 10.1038/s41598-019-38741-1. PubMed PMID: 30778168; PubMed Central PMCID: PMC6379561.

Nilsson E, Klukovich R, Sadler-Riggleman I, Beck D, Xie Y, Yan W, Skinner MK. Environmental toxicant induced epigenetic transgenerational inheritance of ovarian pathology and granulosa cell epigenome and transcriptome alterations: ancestral origins of polycystic ovarian syndrome and primary ovarian insufiency. Epigenetics. 2018;13(8):875-895. doi: 10.1080/15592294.2018.1521223. Epub 2018 Oct 2. PubMed PMID: 30207508; PubMed Central PMCID: PMC6224216.

Wang Z, Lee S, Oliver D, Yuan S, Tang C, Wang Y, Zheng H, Yan W. Prps111, a testis-specific gene, is dispensable for mouse spermatogenesis. Mol Reprod Dev. 2018
Oct;85(10):802-804. doi: 10.1002/mrd.23053. Epub 2018 Sep 7. PubMed PMID: 30113107;
PubMed Central PMCID: PMC6202250.

Grozdanov PN, Li J, Yu P, Yan W, MacDonald CC. Cstf2t Regulates expression of histones and histone-like proteins in male germ cells. Andrology. 2018 Jul;6(4):605-615. doi: 10.1111/andr.12488. Epub 2018 Apr 19. PubMed PMID: 29673127; PubMed Central PMCID: PMC6105451.