## Drugs, Sex, AI, etc - threats to sport integrity

## Yannis P Pitsiladis, MMEDSci., PhD, FACSM

Following 15 years at the University of Glasgow, Scotland, where he created the largest known DNA biobank from world-class athletes and one of the largest DNA biobanks in school-aged children, Yannis was recently appointed Professor in the Department of Sport, Physical Education and Health (SPEH) at Hong Kong Baptist University, Hong Kong, where he plans to create a new center of excellence in sport and exercise science and medicine.

His current research interest is in exploring human enhancement technologies applicable to health and disease with particular focus on responsible paradigms. His most recent research is funded by the World Anti-Doping Agency (WADA) and by the International Olympic Committee (IOC), he is currently a member of the IOC Medical and Scientific Commission, a member of the Executive Committee and Chair of the Scientific Commission of the International Sports Medicine Federation (FIMS), a member of the Scientific and Education Commission of the European Federation of Sports Medicine Associations (EFSMA), a member of WADA's Health Medical Research Committee (HMRC) and is a Fellow of the American College of Sports Medicine (ACSM). He has published over 250 peer-reviewed papers, written and edited a number of books and has featured in numerous research documentaries (e.g. ESPN, BBC, NHK Japan, CNBC) and popular books (e.g. Bounce, The Sports Gene).

## "Sport is digging its own grave"





Richard William Duncan Pound "Dick Pound"

"People will say: 'I'm not going to watch something that's fixed.' If they stop watching, sponsors will question why they are sponsoring this, and broadcaster will ask why they are wasting air time. The whole thing could dry up in a nanosecond."

Home Medal Count Team USA Tracker Shop >

## 2024 Olympic boxing: Imane Khelif earns gold medal in final match amid continued gender controversy

The Algerian took down her toughest test to date -- and a familiar foe -- to walk away from Paris as champion



By Brent Brookhouse Aug 9, 2024 at 5:32 pm ET = 2 min read



**Getty Images** 



### **SPORT**

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Olympics > Schedule | Medal table | Medal results | Day-by-day quide | Video | Team GB | GB medallists | Olympic

## Lin follows Khelif by winning gold amid controversy



Lin Yu-ting beats Poland's Julia Szeremeta to win women's featherweight gold

10 August 2024

An emotional Lin Yu-ting became the second boxer in 24 hours to win women's Olympic gold despite the ongoing row over her gender eligibility.

The Taiwanese beat Polish 20-year-old Julia Szeremeta by unanimous decision to claim the featherweight title, a day after Imane Khelif became the welterweight champion.

Lin and Khelif have been allowed to compete in Paris despite being disqualified from last year's World Championships after reportedly failing gender eligibility tests.



## Petition urges IOC to "never sacrifice women's sport"

By Inside The Games

O Wednesday, 4 December 2024



A joint open letter to the International Olympic Committee, signed by over 40,000 individuals worldwide, calls for the protection of "women's sports, private spaces, and basic fairness" and raises concerns over alleged pressure to allow biological males to compete in female categories.

ADF International hand-delivered the letter to the IOC headquarters in Lausanne, Switzerland this week. It asserts, "Men and women are different. Their physical differences give men

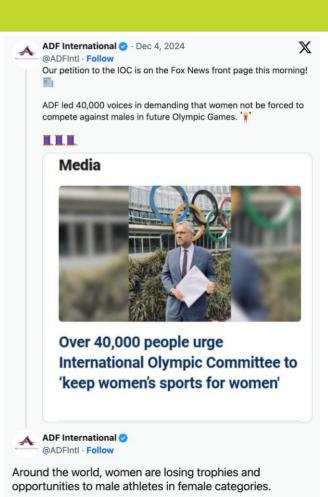
athletic advantages in sports. Scientific research continues to acknowledge this reality."

It further critiques current policies, stating, "However, governments and organisational bodies like the IOC have adopted policies that allow males who identify as female to compete in women's sports. These policies prioritise feelings over fairness, ideology over truth."

Riley Gaines, an advocate for fairness in women's sports who faced competing against males in swimming, commented on the petition. "As a college athlete, my safety and privacy in the locker room was repeatedly jeopardised because of sports bodies which put ideology before women's rights. I'm one of so many young women that have lost out on medals and opportunities – simply because I wasn't a male. What kind of message does that send?"

With the transgender debate in sports as vivid as ever, Gaines highlighted ongoing concerns raised by women athletes, stating, "Women have raised concerns repeatedly about safety, privacy and fairness in women's sports. The IOC is looked upon as a leader on sports policies. It must take heed of this petition, and take a stand for women around the world in protecting our sports – not only for this generation of athletes, but the little girls who one day dream of winning the gold."

According to a report by Reem Alsalem, the UN Special Rapporteur on Violence Against Women and Girls, women have lost over 890 medals across 29 sports to males competing in women's categories. This report was spotlighted during a recent panel event hosted by ADF International at the UN Headquarters in New York.



"The IOC...must take heed of this petition," said

@Riley\_Gaines\_

# **Key Milestones**

## 1. KEY MILESTONES



2022

IOC

Roll-out of the

Framework

5



2009
First public investigation on hyperandrogenism around Caster Semenva's case



2015
CAS ruling on
Dutee Chand
suspends any
hyperandrogenism
rule



intervention

IAAF eligibility rules

2018

The World Health
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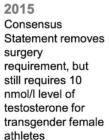
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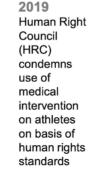














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IOC finalizes consultations and releases Framework 2024 Olympic boxing: Imane Khelif earns gold medal in final match amid continued gender controversy

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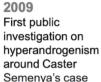


# **Key Milestones**

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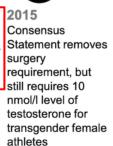
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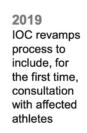


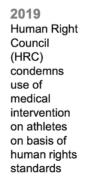




2011 IAAF introduces eligibility rules









2021 Tokyo 2020 Games with first openly transgender athletes	2022 Roll-out of the IOC Framework
IOC finalizes consultations	

and releases

5

Framework





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**TODAY IN ATHLETICS** 

IAAF NEWS AND PRESS RELEASES

REGISTER FOR PRESS RELEASES

12 APR 2011 IAAF NEWS DAEGU, KOREA

## IAAF TO INTRODUCE ELIGIBILITY RULES FOR FEMALES WITH HYPERANDROGENISM







The IAAF today became the first international sports federation to approve the adoption of new rules and regulations governing the eligibility of females with hyperandrogenism\* to compete in women's competition.

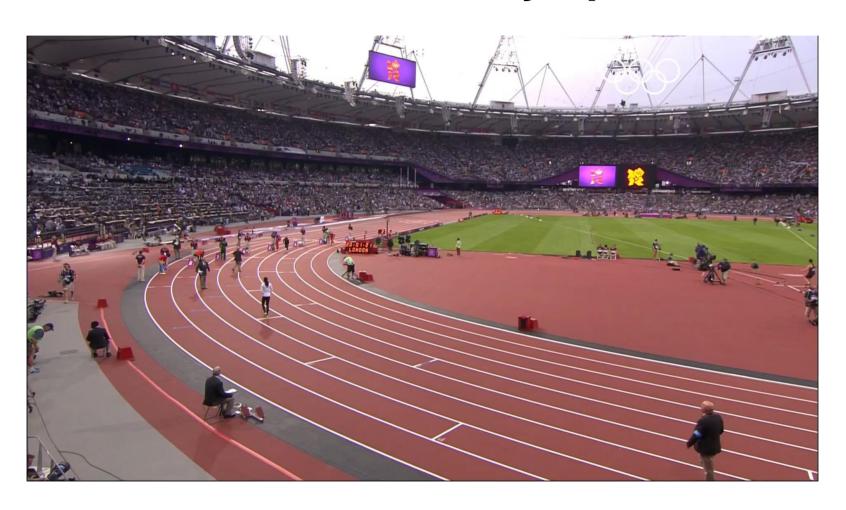
The IAAF Council's decision is the culmination of an 18 month-long review by an IAAF expert working group who have studied issues relating to the participation of female athletes with hyperandrogenism in athletics. This group has worked in close co-ordination with the IOC Medical Commission throughout this period and also participated in a series of international expert meetings held on the subject in 2010.

The new rules and regulations, which will be published and come into force for all International Competitions on 1 May 2011, provide for the following key principles:

- Competition in athletics will continue to be divided into men's and women's competition recognising that there is a difference in sporting performance between elite men and women, that is predominantly due to higher levels of androgenic hormones in men;

- A female with hyperandrogenism who is recognised as a female in law shall be eligible to compete in women's competition in athletics provided that she has androgen levels below the male range (measured by reference to testosterone levels in serum) or, if she has androgen levels within the male range she also has an androgen resistance which means that she derives no competitive advantage from such levels;
- A pool of international medical experts has been appointed by the IAAF to review cases referred to it under the regulations as an independent expert medical panel and to make recommendations to the IAAF in such cases to decide on the eligibility of female athletes with hyperandrogenism;
- A 3-level medical process under the regulations shall ensure that all potentially relevant data is made available to the expert medical panel for the purposes of evaluating an athlete's eligibility. This medical process may include, where necessary, the expert medical panel referring an athlete with potential hyperandrogenism for full examination and diagnosis in accordance with best medical practice at one of the 6 IAAF-approved specialist reference centres around the world:
- The medical process under the regulations shall be conducted in strict confidentiality and all cases shall be referred to the expert medical panel on an anonymous basis:
- A female athlete who declines, fails or refuses to comply with the eligibility determination process under the regulations shall not be eligible to compete in women's competition.

# **2012 Summer Olympics**

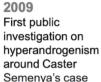


# **Key Milestones**

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2015 CAS ruling on **Dutee Chand** suspends any hyperandrogenism rule

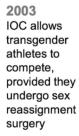


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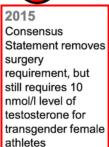
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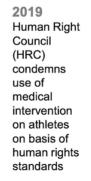




2011 IAAF introduces eligibility rules









2021 **Tokyo 2020** Games with first openly transgender athletes

**IOC** finalizes

consultations and releases

Framework

2022 Roll-out of the IOC Framework





## IOC Consensus Meeting on Sex Reassignment and Hyperandrogenism November 2015

Participants:

Prof Dr Uğur Erdener Chairman, IOC Medical & Scientific Commission

Prof Arne Ljungqvist Former Chairman, IOC Medical Commission

Dr Stéphane Bermon Monaco Institute of Sports Medicine & Surgery, IAAF Medical &

Scientific Senior Consultant

Michael Beloff, QC Barrister, Blackstone Chambers

Prof Gerard Conway Professor of Clinical Medicine, University College London

Prof Myron Genel Professor Emeritus of Pediatrics and Senior Research Scientis

Yale Child Health Research Center Yale University School of Medicine

Ms Joanna Harper Chief Medical Physicist, Radiation Oncology, Providence

Portland Medical Center

Prof Angelica Linden De

Hirschberg

Department of Woman & Child Health, Division of Obstetrics &

Gynecology, Karolinska Institutet

Prof Dr Maria Jose Martinez

Patino

Faculty of Sport Sciences, University of Vigo

Prof Martin Ritzén Professor Emeritus, Dept of Woman and Child Health

Karolinska Institutet

Dr Eric Vilain Professor of Human Genetics, Pediatrics and Urology

Director, Center for Gender-Based Biology Chief, Medical Genetics, Department of Pediatrics

Co-director, Clinical Genomic Center David Geffen School of Medicine at UCLA

Jonathan Taylor Partner, Bird & Bird

Liz Riley Barrister, Bird & Bird

Dr Robin Mitchell Vice-Chair, IOC Medical & Scientific Commission

Dr Rania Elwani Member, IOC Medical & Scientific Commission

Dr Vidya Mohamed-Ali Member, IOC Medical & Scientific Commission

Prof Yannis Pitsiladis Member, IOC Medical & Scientific Commission

Dr Richard Budgett IOC Medical & Scientific Director

Dr Lars Engebretsen IOC Head of Scientific Activities

Christian Thill IOC Senior Legal Counsel

#### 1) Transgender guidelines



In this spirit, the IOC Consensus Meeting agreed the following guidelines to be taken into account by sports organisations when determining eligibility to compete in male and female competition:

- Those who transition from female to male are eligible to compete in the male category without restriction.
- 2. Those who transition from male to female are eligible to compete in the female category under the following conditions:
  - 2.1. The athlete has declared that her gender identity is female. The declaration cannot be changed, for sporting purposes, for a minimum of four years.
  - 2.2. The athlete must demonstrate that her total testosterone level in serum has been below 10 nmol/L for at least 12 months prior to her first competition (with the requirement for any longer period to be based on a confidential case-by-case evaluation, considering whether or not 12 months is a sufficient length of time to minimize any advantage in women's competition).

# **Key Milestones**

## 1. KEY MILESTONES









2015 CAS ruling on **Dutee Chand** suspends any hyperandrogenism rule



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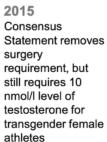
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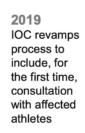






2015 IAAF Consensus introduces surgery eligibility rules

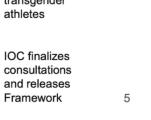






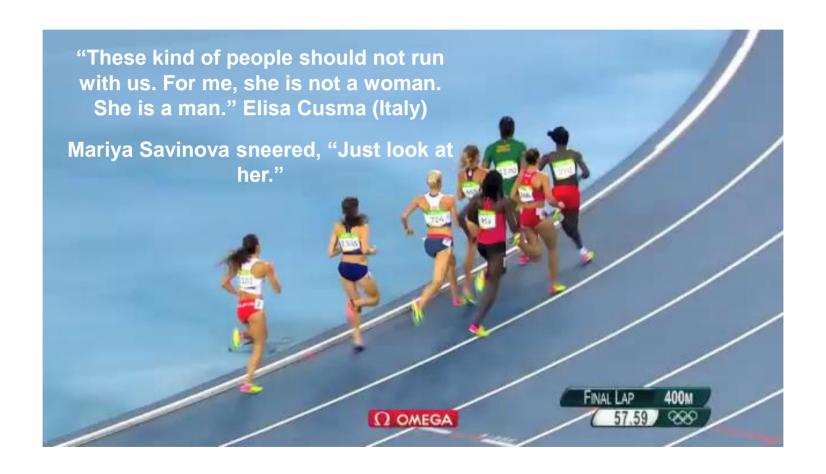
2020 HRC presents its report on discrimination on sport and gender identification.

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	IOC finalizes consultations	





## **2016 Summer Olympics**



# **Key Milestones**

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2015
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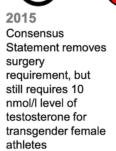


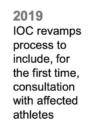


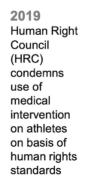




2011 IAAF introduces eligibility rules













26 APR 2018 PRESS RELEASE MONACO

## IAAF INTRODUCES NEW ELIGIBILITY **REGULATIONS FOR FEMALE CLASSIFICATION**







The IAAF this week issued new Eligibility Regulations [English | French] for Female Classification (Athlete with Differences of Sexual Development) for events from 400m to the mile, including 400m, hurdles races, 800m, 1500m, one mile races and combined events over the same distances ('Restricted Events').

The new regulations require any athlete who has a Difference of Sexual Development (DSD) that means her levels of circulating testosterone (in serum) are five (5) nmol/L or above and who is androgen-sensitive to meet the following criteria to be eligible to compete in Restricted Events in an International Competition (or set a world record in a Restricted Event at competition that is not an International Competition):

- (a) she must be recognised at law either as female or as intersex (or equivalent);
- (b) she must reduce her blood testosterone level to below five (5) nmol/L for a continuous period of at least six months (e.g., by use of hormonal contraceptives); and
- (c) thereafter she must maintain her blood testosterone level below five (5) nmol/L continuously (ie: whether she is in competition or out of competition) for so long as she wishes to remain eligible.



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ALL PAGES RELATED TO THIS ARTICLE

# **Key Milestones**

## 1. KEY MILESTONES









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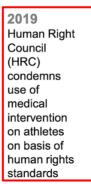






2015 Consensus Statement removes surgery requirement, but still requires 10 nmol/l level of testosterone for transgender female athletes







2021 Tokyo 2020 Games with first openly transgender athletes	2022 Roll-out of the IOC Framework
IOC finalizes consultations and releases Framework	5



#### LETTER TO THE EDITOR



#### Response to the United Nations Human Rights Council's Report on Race and Gender Discrimination in Sport: An Expression of Concern and a Call to Prioritise Research

Blair R. Hamilton 1,2,3 . Maria Jose Martinez-Patiño 4 · James Barrett 3 · Leighton Seal 3 · Ross Tucker 5 · Theodora Papadopoulou<sup>6,7,8,9</sup> · Xavier Bigard<sup>6,7,10</sup> · Alexander Kolliari-Turner<sup>1</sup> · Herbert Löllgen · Petra Zupet<sup>6</sup> · Anca Jonescu<sup>6</sup> · Andre Debruyne<sup>6,7</sup> · Nigel Jones<sup>8,11</sup> · Juergen M. Steinacker<sup>6,7,12</sup> · Karin Vonbank<sup>13</sup> · Giscard Lima 1,14 · Federica Fagnani · Chiara Fossati 14,15 · Luigi Di Luig Maurizio Casasco<sup>6,7,16</sup> · Michael Geistlinger<sup>7,17</sup> · Bernd Wolfarth<sup>7,18</sup> · Jane T. Seto<sup>19,20</sup> · Norbert Bachl<sup>6,7,21,22</sup> · Richard Twycross-Lewis 23 0 · David Niederseer 24 0 · Andrew Bosch 25 · Jeroen Swart 7,25 · Demitri Constantinou 7,26 0 · Boria Muniz-Pardos<sup>27</sup> • José Antonio Casaius<sup>27</sup> • Victoriva Badtieva<sup>7,28,29</sup> • Irina Zelenkova<sup>27,28</sup> • James L. J. Bilzon<sup>7,8,30</sup> · Michiko Dohi<sup>7,31</sup> · Christian Schneider<sup>7,32</sup> · Sigmund Loland<sup>33</sup> · Michele Verroken<sup>34,35</sup> · Pedro Manonelles Marqueta<sup>36</sup> · Francisco Arroyo<sup>7,37</sup> · André Pedrinelli<sup>7,38</sup> · Konstantinos Natsis<sup>6,7,39</sup> · Evert Verhagen<sup>40</sup> · William O. Roberts<sup>7,41</sup> · José Kawazoe Lazzoli<sup>7,42</sup> · Rogerio Friedman<sup>43</sup> · Ali Erdogan<sup>7,44</sup> Ana V. Cintron<sup>7,45</sup> · Shu-Hang Patrick Yung<sup>7,46</sup> · Dina C. Janse van Rensburg<sup>7,47</sup> · Dimakatso A. Ramagole<sup>7,47</sup> Sandra Rozenstoka<sup>6,7,48</sup> · Felix Drummond<sup>6,7,49</sup> · Nick Webborn<sup>50</sup> · Fergus M. Guppy<sup>1,2</sup> · Yannis P. Pitsiladis<sup>1,6,7,14</sup>

Accepted: 24 October 2020 © Springer Nature Switzerland AG 2020

Dear Editor,

Caster Semenya recent tion of blood testostero World Athletics [2, 3]. orders of Sex Develop testosterone concentrat least six months, and t testosterone continuou for events between 400 resolution 40/5, on rac published by the Unit (UNHRC) which calls

The UNHRC [4] and World Medical Association [5] argue that World Athletics regulation [2] denies DSD women the right to participate in certain events unless they accept "unnecessary medical intervention", and that athletes are being coerced into such treatment.

The authors agree that there is currently no direct scientific evidence that DSD women athletes with higher testosterone levels have a performance advantage in sporting events between 400 and 1500 m. This evidence will be difficult to obtain, given the low number of DSD athletes and the ethical considerations in such research. The principles

Extended author information available on the last page of the article

"Unnecessary

etween the theory and hormonal control and lective ban is questionnts at opposite ends of hlete Michael Johnson Intervention" and 400 m, as well as 100, 200, and 400 m. levels in the 1500 and lated by the policy, the

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latter is not.

Given the influence of high testosterone concentrations, there is concern that 6 months of testosterone suppression is not long enough to negate potential advantages from lifelong exposure. There is growing support for the idea that development influenced by high testosterone levels may result in retained anatomical and physiological advantages [7]. One such inherent "legacy" effect may be the phenomenon of muscle memory [8], that has been defined as the ability to rebuild muscle mass and strength after a long intervening period of inactivity and muscle mass loss [9].

The debate over sporting fairness, highlighted by the UNHRC, is particularly pertinent in DSD athletes [1, https://doi.org/10.1007/s40279-021-01451-8

Sports Medicine

#### REVIEW ARTICLE



#### Integrating Transwomen and Female Athletes with Differences of Sex Development (DSD) into Elite Competition: The FIMS 2021 Consensus Statement

Blair R. Hamilton<sup>2,3</sup> Giscard Lima<sup>1,4</sup> James Barrett<sup>3</sup> Leighton Seal<sup>3</sup> Alexander Kolliari-Turner<sup>2</sup> Guan Wang<sup>61</sup> · Antonia Karanikolou<sup>2</sup> · Xavier Bigard<sup>5,6,7</sup> · Herbert Löllgen<sup>6</sup> · Petra Zupet<sup>6</sup> · Anca Ionescu<sup>6</sup> · Andre Debruvne<sup>6,7</sup> · Nigel Jones<sup>8,9</sup> · Karin Vonbank<sup>10</sup> · Federica Fagnani<sup>4</sup> · Chiara Fossati<sup>4,11</sup> · Maurizio Casasco<sup>6,7,12</sup> · Demitri Constantinou<sup>7,13</sup> · Bernd Wolfarth<sup>7,14</sup> · David Niederseer<sup>15</sup> · Andrew Bosch 16 · Boria Muniz-Pardos 17 0 · José Antonio Casaius 17 0 · Christian Schneider 7.18 · Sigmund Loland 19 . Michele Verroken 20,21 . Pedro Manonelles Margueta 7,22 · Francisco Arrovo 7,23 . André Pedrinelli<sup>7,24</sup> · Konstantinos Natsis<sup>6,7,25,26</sup> · Evert Verhagen<sup>27</sup> · William O. Roberts<sup>7,28</sup> José Kawazoe Lazzoli<sup>7,29</sup> · Rogerio Friedman<sup>30</sup> · Ali Erdogan<sup>7,31</sup> · Ana V. Cintron<sup>7,32</sup> · Shu-Hang Patrick Yung<sup>7,33</sup> Dina C. Janse van Rensburg 34 0 · Dimakatso A. Ramagole 34 · Sandra Rozenstoka 6,7,35 · Felix Drummond 6,7,36 · Theodora Papadopoulou<sup>6,7,37</sup> · Paulette Y. O. Kumi<sup>38</sup> · Richard Twycross-Lewis<sup>39</sup> · Joanna Harper<sup>40</sup> · Vasileios Skiadas<sup>41</sup> · Jonathan Shurlock<sup>42</sup> · Kumpei Tanisawa<sup>43</sup> · Jane Seto<sup>44,45</sup> · Kathryn North<sup>44,45</sup> · Siddhartha S. Angadi<sup>46</sup> · Maria Jose Martinez-Patiño<sup>47</sup> · Mats Boriesson<sup>7,48,49</sup> · Luigi Di Luigi<sup>7,50</sup> · Michiko Dohi<sup>7,51</sup> · Jeroen Swart<sup>7,52</sup> · James Lee John Bilzon<sup>7,53</sup> · Victoriva Badtieva<sup>7,54,55</sup> · Irina Zelenkova<sup>17</sup> · Juergen M. Steinacker<sup>6,7,56</sup> Norbert Bachl<sup>6,7,57,58</sup> · Fabio Pigozzi<sup>4,6,7,11</sup> · Michael Geistlinger<sup>7,59</sup> Dimitrios G. Goulis<sup>60</sup> · Fergus Guppy<sup>2,61</sup> · Nick Webborn<sup>62</sup> · Bulent O. Yildiz<sup>63</sup> · Mike Miller<sup>64</sup> · Patrick Singleton<sup>64</sup> · Yannis P. Pitsiladis<sup>1,2,4,6,7</sup>

Accepted: 10 © The Author

## Abstract

Sport is hi governing be lasting a concentrati inclusion of able scient suppressio including 1 conduct lor sports to it

to be based

"Necessary Intervention for Gender Classification for Sport" s to lual eds

even the most evidence-based regulations are unlikely to eliminate all differences in performance between cisgender women with and without DSD and transwomen athletes. Any remaining advantage held by transwomen or DSD women could be considered as part of the athlete's unique makeup.

#### 1 Introduction

Since antiquity, athletic and Olympic competitions have been separated according to the traditional binary concept of male/female to promote fairness and equity, as well as being divided by criteria such as weight, age, affiliation,

Extended author information available on the last page of the article

Published online: 08 December 2020

Published online: 24 March 2021

# **Key Milestones**

## 1. KEY MILESTONES





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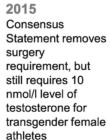




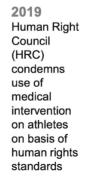




IAAF introduces eligibility rules









2021 **Tokyo 2020** its report on Games with discrimination first openly on sport and transgender athletes gender identity

> **IOC** finalizes consultations and releases Framework







## **2021 Summer Olympics**



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2015 Consensus Statement removes surgery requirement, but still requires 10 nmol/l level of testosterone for transgender female athletes

2019 IOC revamps process to include, for the first time. consultation with affected athletes

2019 Human Right Council (HRC) condemns use of medical intervention on athletes on basis of human rights standards

2020 HRC presents its report on discrimination on sport and aender identity

2021 **Tokyo 2020** Games with first openly transgender athletes

2022 Roll-out of the IOC Framework

**IOC** finalizes consultations and releases Framework



## **IOC Framework**

# 5. NEW IOC FRAMEWORK - REDEFINING THE ROLE THAT THE IOC PLAYS



2015 Consensus Statement (Current approach)

The IOC recommends a *one-size-fits-all* approach that all International Federations should apply to define eligibility.



Proposed Framework (replaces previous approach, including Consensus Statement 2015)

The IOC <u>offers guidance</u> to International Federations on how to design eligibility criteria <u>that work for their own sport/context</u>, while considering fairness, inclusion and non-discrimination.

## **IOC Framework**

## A 10-PRINCIPLE APPROACH TO DECISION-MAKING

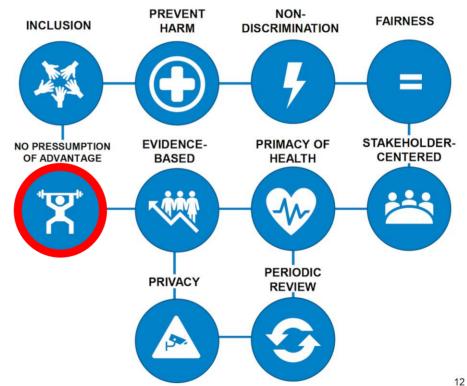




"Unnecessary Medical Intervention"









Open access Viewpoint

**BMJ Open** Sport & Exercise Medicine

Joint position statement of the **International Federation of Sports** Medicine (FIMS) and European **Federation of Sports Medicine** Associations (EFSMA) on the IOC framework on fairness, inclusion and nondiscrimination based on gender identity and sex variations

Fabio Pigozzi O, 1,2,3 Xavier Bigard, 1,4,5 Juergen Steinacker O, 1,6 Bernd Wolfarth , 1.7 Victoriya Badtieva , 1.8 Christian Schneider , 1.10 Jeroen Swart , 1.11 James Lee John Bilzon , 1.12 Demitri Constantinou , 1.13 Michiko Dohi <sup>1,14</sup> Luigi Di Luigi. <sup>1,15</sup> Chiara Fossati <sup>1,2,3</sup> Norbert Bachl. <sup>1,16,17</sup> Guoping Li, 1,18 Theodora Papadopoulou, 1,4,19 Maurizio Casasco, 1,4,20 Dina Christina (Christa) Janse van Rensburg 0, 1,21,22 Jean-François Kaux, 1,23 Sandra Rozenstoka, <sup>1,24</sup> Jose-Antonio Casajus <sup>0</sup>, <sup>1,25</sup> Irina Zelenkova, <sup>1,2</sup> Emre Ak, <sup>1,26</sup> Bulent Ulkar, <sup>1,27</sup> Francisco Arroyo <sup>1,28</sup> Anca Ionescu, <sup>4,29</sup> André Pedrinelli , 1,30 Mike Miller, 1 Patrick Singleton, 1 Malay Shroff, 1 Nick Webborn, 32,33 James Barrett, 34 Blair Hamilton , 32,35 Michael Geistlinger, 36 Gianfranco Beltrami, <sup>37,38</sup> Sergio Migliorini, <sup>39</sup> Lenka Dienstbach-Wech, <sup>40</sup> Stéphane Bermon <sup>9</sup>, <sup>41,42</sup> Yannis P Pitsiladis <sup>9</sup>, <sup>14,32,35</sup> on behalf of the International Federation of Sports Medicine (FIMS) and the European Federation of Sports Medicine Associations (EFSMA)

Steinacker J. et al. Joint position statement of the International Federation of Sports Medicine (FIMS) and European Federation of Sports Medicine Associations (EFSMA) on the IOC framework on fairness, inclusion and nondiscrimination based on gender identity and sex variations RM. Open Sport & Exercise Medicine 2022;8:e001273. doi:10.1136/ bmjsem-2021-001273

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inclusion and non-discrimination based on gender identity and sex variations. This framework is drafted mainly from a human rights perspective, with less consideration for medical/scientific issues. The framework places the onus for gender eligibility and classification entirely on the International Federations (IFs), even though most will not have the capacity to implement the framework. The position of no presumption of advantage is contrary to the 2015 IOC consensus. Implementation of the 2021 framework will be a major challenge for IFs that have already recognised the inclusion of trans and women athletes with differences of sexual development (DSD) using a scientific/medical solution. The potential consequences for sports that need to prioritise fairness or safety could be one of two extremes (1) exclusion of all transgender or DSD athletes on the grounds of advantage or (2) self-identification that essentially equates to no eligibility rules, Exclusion of all transgender or DSD athletes is contrary to the Olympic charter and unlawful

in many countries. While having no gender eligibility

rules, sport loses its meaning and near-universal support

The IOC recently published its framework on fairness,

#### Key messages

#### What is already known

- Athletes are currently excluded from the female category in sport based on transgender or differences of sexual development status.
- ► The IOC published its new framework on fairness inclusion and non-discrimination based on gende identity and sex variations in November 2021.
- ► The publication of this framework has generated debate and discussion in sport and exercise medicine

Athletes should not be under pressure to undergo medical procedures or treatment to meet eligibility criteria. However, if an athlete is fully informed and consents, ther it is their free choice to undergo carefully considered or necessary interventions for gender classification for sport to compete fairly and safely in their chosen gender. Free choice is a fundamental human right, but so is the right to fair and safe competition.

#### CONSEQUENCES

The potential consequences of these imposed constraints could be one of two unwanted extremes (1) exclusion of transgender or DSD athletes on the grounds of performance advantage or (2) self-identification that all but equates to no eligibility rules. By adopting the recent IOC framework without the possibility of necessary interventions for gender classification (the authors preferred use of the term rather than the unhelpful and emotionally charged concept of 'unnecessary medical intervention'),



# Some examples - Regs/Guidelines

- World Rugby: In 2020, WR launched their guidelines. To play women's rugby, transgender women must provide medical documentation confirming they have not experienced and are suppressing a testosterone-driven puberty. Transgender men, regardless of whether they transitioned pre- or post-puberty, may play men's rugby if they provide confirmation they have the physical ability to ensure they are not putting themselves at unacceptable risk. Transgender men who have begun a sex reassignment process that includes supplementing with testosterone may not play women's rugby.
- International Swimming Federation (FINA): In June 2022, FINA voted to bar all transgender athletes from competing in professional women's swimming, with the exception of athletes who "can establish to FINA's comfortable satisfaction that they have not experienced any part of male puberty beyond Tanner Stage 2 (of puberty) or before age 12, whichever is later". FINA also announced the creation of an "open" category for transgender swimmers to compete in.
- world Boxing Council (WBC): On Monday (15 August 2022), the international professional boxing organisation announced new guidelines on the eligibility of trans boxers in its tournaments. WBC firmly and unequivocally supports transgender rights. Neither trans women nor trans men can compete against cis athletes as it would be "unfair. A combat sport bout should occur between two equally matched competitors. At present, there is no consensus whether a bout between a transgender woman against a cisgender (biological) woman is a fair bout between two equally matched competitors. Metric such as testosterone level less than 10 nanomoles per litre (achieved by using testosterone suppression medication in the transgender woman), in isolation is inadequate to ensure fairness at the time of the bout.

- World Athletics: March 24, 2023. World Athletics has banned transgender women from competing in elite female competitions and tightened testosterone restrictions for other athletes, the governing body said on Thursday. World Athletics president Sebastian Coe said that the decision to exclude transgender women who had gone through male puberty was based "on the overarching need to protect the female category". The tighter measures around one of the most contentious and divisive issues in sport follow a similar move by World Aquatics in 2022
- UCI: July 15, 2023 UCI bans transgender women from competing in elite international races. Governing body claim their actions are to 'protect the female class. The UCI has followed other sports and banned transgender women from competing in women's category races in international competition.
- ICC: 21 November 2023. Transgender women have been barred from playing in international women's matches under new regulations from the International Cricket Council. Any player who has gone through male puberty will not be eligible for women's internationals regardless of any surgery or treatment undertaken. In September, Canada's Danielle McGahey became the first transgender cricketer to play an official international game. The new regulations will be reviewed within two years, the ICC said. Following a nine-month consultation process, the governing body said its new policy, which takes effect immediately, was based on "protection of the integrity of the women's game, safety, fairness and inclusion".

and the list will keep growing.....

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#### ABSTRACT

The IOC recently published its framework on fairness, inclusion and non-discrimination based on gender identity and sex variations. This framework is drafted mainly from a human rights perspective, with less consideration for medical/scientific issues. The framework places the onus for gender eligibility and classification entirely on the International Federations (IFs), even though most will not have the capacity to implement the framework. The position of no presumption of advantage is contrary to the 2015 IOC consensus. Implementation of the 2021 framework will be a major challenge for IFs that have already recognised the inclusion of trans and women athletes with differences of sexual development (DSD) using a scientific/medical solution. The potential consequences for sports that need to prioritise fairness or safety could be one of two extremes (1) exclusion of all transgender or DSD athletes on the grounds of advantage or (2) self-identification that essentially equates to no eligibility rules, Exclusion of all transgender or DSD athletes is contrary to the Olympic charter and unlawful in many countries. While having no gender eligibility rules, sport loses its meaning and near-universal support

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Athletes should not be under pressure to undergo medical procedures or treatment to meet eligibility criteria. However, if an athlete is fully informed and consents, then it is their free choice to undergo carefully considered or necessary interventions for gender classification for sport to compete fairly and safely in their chosen gender. Free choice is a fundamental human right, but so is the right to fair and safe competition.

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Home Medal Count Team USA Tracker Shop >

## 2024 Olympic boxing: Imane Khelif earns gold medal in final match amid continued gender controversy

The Algerian took down her toughest test to date -- and a familiar foe -- to walk away from Paris as champion



By Brent Brookhouse Aug 9, 2024 at 5:32 pm ET = 2 min read



**Getty Images** 



### **SPORT**

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## Lin follows Khelif by winning gold amid controversy



Lin Yu-ting beats Poland's Julia Szeremeta to win women's featherweight gold

10 August 2024

An emotional Lin Yu-ting became the second boxer in 24 hours to win women's Olympic gold despite the ongoing row over her gender eligibility.

The Taiwanese beat Polish 20-year-old Julia Szeremeta by unanimous decision to claim the featherweight title, a day after Imane Khelif became the welterweight champion.

Lin and Khelif have been allowed to compete in Paris despite being disqualified from last year's World Championships after reportedly failing gender eligibility tests.

# My thoughts on DSD/Trans

- The eligibility criteria to compete in [Olympic] sport must be based on the best available scientific/medical evidence and minimising the unavoidable discrimination needed to ensure fair or safe competition;
- Currently, hormonal control (although imperfect) is the best way to ensure as fair as possible inclusion of DSD/Trans athletes into elite sport;
- Sport specific studies are needed to quantify the impact (i.e., magnitude and time) of hormonal control on health- and performance-related phenotypes;
- Use of routine markers (i.e., free T, estradiol, prolactin, LH, FSH, SHBG, PSA) and discovery of new biomarkers are needed; and
- Political opinions must not dominate.

FUROPEAN JOURNAL OF SPORT SCIENCE https://doi.org/10.1080/17461391.2021.1938692







KEYWORDS

Transwomen; archery;

shooting: eligibility

#### Integrating transwomen athletes into elite competition: The case of elite archery and shooting

Blair R. Hamilton <sup>©</sup> a,b, Fergus M. Guppy <sup>©</sup> a, James Barrett<sup>b</sup>, Leighton Seal<sup>b</sup> and Yannis Pitsiladis <sup>©</sup> a,c,d,e,f

<sup>a</sup>Centre for Stress and Age-related Disease. University of Brighton, Brighton, UK: <sup>b</sup>The Gender Identity Clinic Tayistock and Portman NHS Foundation Trust, London, UK; 'University of Rome "Foro Italico", Rome, Italy; d'Centre for Exercise Sciences and Sports Medicine, FIMS Collaborating Centre of Sports Medicine, Rome, Italy; european Federation of Sports Medicine Associations (EFSMA), Lausanne, Switzerland; International Federation of Sports Medicine (FIMS), Lausanne, Switzerland

#### ABSTRACT

The inclusion of transwomen into elite female sport has been brought into guestion recently with World Rugby banning transwomen from the elite female competition, aiming to prioritise safety over fairness and inclusion, citing the size, force and power-producing advantages conferred to transwomen. The same question is being asked of all Olympic sports including non-contact sports such as archery and shooting. As both these Olympic sports are the polar opposite to the contact sport of rugby in terms of the need to consider the safety of athletes, the IF of both archery and shooting should consider the other elements when deciding the integration of trans individuals in their sports. Studies on non-athletic transwomen have reported muscle mass and strength loss in the range of 5-10% after 1 year of their transition, with these differences no longer apparent after 2 years. Therefore, based on the current scientific literature, it would be justified for meaningful competition and to prioritise fairness, that transwomen be permitted to compete in elite archery after 2 years of GAT. Similarly, it would be justified in terms of shooting to prioritise inclusion and allow transwomen after 1 year of GAT given that the only negligible advantage that transwomen may have is superior visuospatial coordination. The impact of this considered integration of transwomen in elite sports such as archery and shooting could be monitored and lessons learned for other sports, especially where there are no safety concerns from contact with an opponent.

Abbreviations: IF: International Sports Federation; FDS: Flexor Digitorum Superficialis; IOC: International Olympic Committee; EA: Elite Archer; NA: Non-elite Archer; INT: International Athletes; NAT: National Athletes; GAT: Gender Affirming Treatment; O2: Oxygen; cHb: Haemoglobin Concentration in Blood

#### 1. Introduction

The inclusion of transwomen into elite female sport has been brought into question in recent times, not least by the decision of World Rugby to ban transwomen from elite female competition (Rugby, 2020). As the main physical attributes of rugby are strength, speed, and power and the rules and strategies of the game encourage intense physical contact with opponents, in their recent deliberations World Rugby prioritised the safety of athletes over fairness and inclusion as reflected in the proposed decision-making triangle (Figure 1A) (Rugby, 2020). A recent review of studies conducted in non-athletic transwomen undergoing gender affirming treatment (GAT) highlights the time course changes in lean body mass, muscle cross-sectional area, and muscular strength

(i.e. 12-36 months) and haemoglobin and/or haematocrit (i.e. 3-4 months) (Harper, O'Donnell, Khorashad, McDermott, & Witcomb, 2021). Similarly, another recent review argued that lean body mass, muscle size, bone density and strength were are only trivially affected by 12 months of GAT; the period previously advocated for inclusion of transgender women in female sports categories (Hilton & Lundberg, 2021). These two recent reviews highlight the urgent need for research examining the impact of GAT in transwomen athletes to inform the decision-making process and properly consider the integration of transwomen in elite female sport (Hamilton et al., 2021). These examples also show that there is a clear need for a roadmap demonstrating how integration

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## Framework



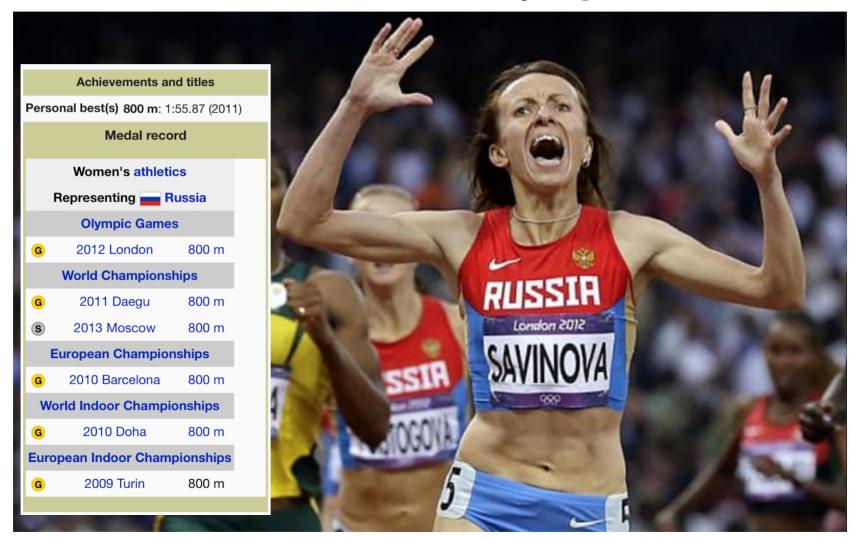
Figure 1. The declared weightings for World Rugby (A) and the derived weightings for World Archery (B) and International Shooting Sport Federation (C) using the three primary criteria for formulating the guidelines and/or rules for the fair and safe integration of transwomen athletes.



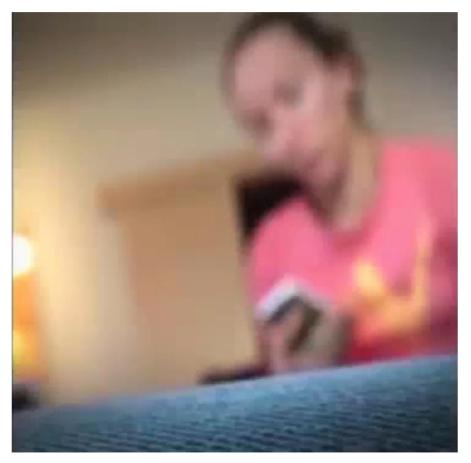
Figure 2. Proposed sliding scale tool that IF's may use to decide what weighting to prioritise in their respective sports for the inclusion/exclusion of transwoman athletes. This sliding scale may be used holistically as a sport to decide their orientation in the decision-making triangle of Figure 1, or it can be used to prioritise what to assess in an individual transwomen's eligibility case for that sport.

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## **2012 Summer Olympics**



# Russia's state-sponsored doping system



German television channel ARD, 2014

(voice dubbed)

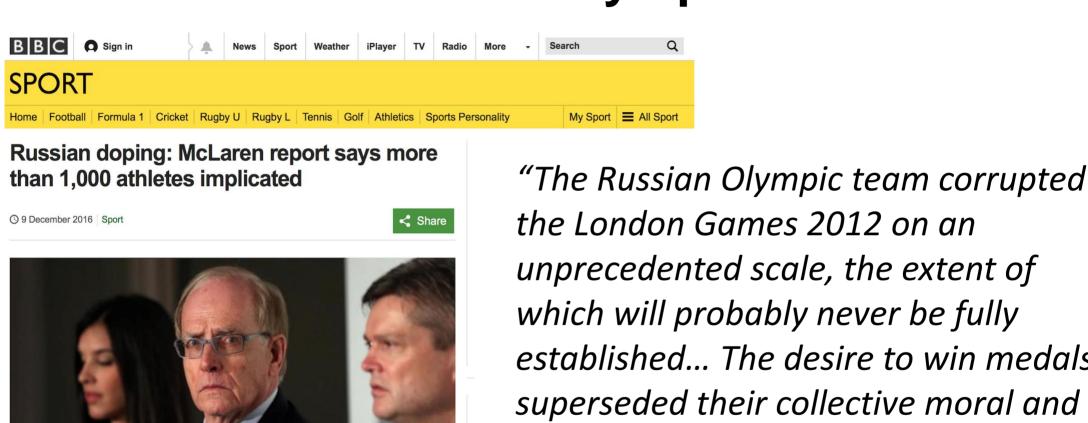
That is our system and in Russia that only works only with pharma...

My coach fortunately works with Melnikov [Russian Head Coach] and he helps to cover up the tests. They allow him to change the dates for the controls.

Oxandrolone [anabolic-androgenic steroid] is very quickly out of my body again. It takes less than 20 days. We have tested that.

My husband has very good contacts to the doping control laboratory.

# **London 2012 Summer Olympics**



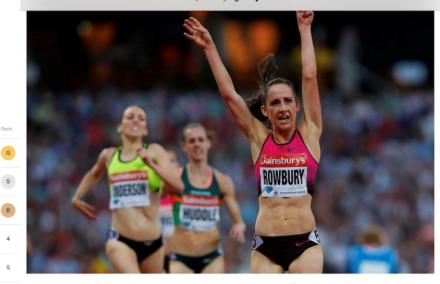
The report's author, Richard McLaren (centre), said doping took place on 'an unprecedented scale'

the London Games 2012 on an unprecedented scale, the extent of which will probably never be fully established... The desire to win medals superseded their collective moral and ethical compass and Olympic values of fair play."

# **2012 Summer Olympics**

	į	WOMEN'S 800M	<b>€</b>
7	17	RESULT - FINAL	e R
1	RUS	MARIYA SAVINOVA	1:56.19
2	RSA.	CASTER SEMENYA	1:57.23
3	RUS	EKATERINA POISTOGOVA	1:57.53
4	KEN=	PAMELA JELIMO	1:57.59
5	USA	ALYSIA JOHNSON MONTANO	1:57.93
6	RUS	ELENA ARZHAKOVA	1:59.21
7	BDI	FRANCINE NIYONSABA	1:59.63
8	KEN=	JANETH JEPKOSGEI BUSIENEI	2:00.19

#### The Washington Post



# Twelve years after finishing sixth, she's in line for Olympic bronze 1500 m

American Shannon Rowbury feels "some good closure" after another suspension in an infamous London Games race leaves her with the third-fastest time.

#### BY ADAM KILGORE

SEPTEMBER 4 AT 8:21 AM

On Tuesday morning, Shannon Rowbury was sitting with her family in the lobby of a Quito, Ecuador hotel, the requisite layover on a trip to the Galápagos Islands. The greatest American middle-distance runner of her generation, Rowbury will turn 40 later this year, and the milestone called for a bucket list vacation with her husband and two young children.

<sup>2</sup> International Association of Athletics Federations, Monaco, Monaco.

## Prevalence of Blood Doping in Samples Collected from Elite Track and Field Athletes

Pierre-Edouard Sottas, <sup>1\*</sup> Neil Robinson, <sup>1</sup> Giuseppe Fischetto, <sup>2</sup> Gabriel Dollé, <sup>2</sup> Juan Manuel Alonso, <sup>2</sup> and Martial Saugy <sup>1</sup>

	n <sup>a</sup>	Prevalence M1, % <sup>b</sup>	Prevalence M2, %
Males	4028	12 (10–15)	12 (10–15)
Country A	205	48 (35-63)	78 (54–99)
Country B	352	3 (1–11)	1 (0-2)
Country C	257	23 (15–30)	28 (17–36)
Country D	208	6 (3-19)	5 (0–17)
Country E	160	18 (11–30)	18 (7–28)
Country F	148	6 (1–25)	2 (0–22)
Country H	160	39 (20-54)	51 (21–87)

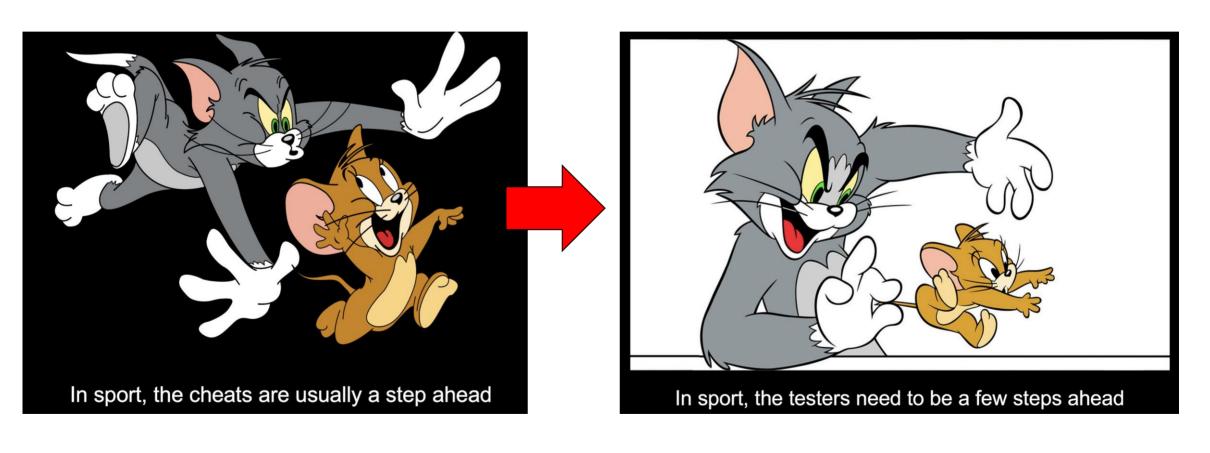
<sup>&</sup>lt;sup>a</sup> n, Number of samples from which the estimates were derived. Prevalence M1, minimal estimates without any assumptions on the doping method. Prevalence M2, estimates obtained assuming doping with rEPO microdoses.

# **Table 2.** Period prevalence estimates of abnormal blood profiles in elite track and field athletes.

Females	3261	18 (15–21)	18 (15–21)
Country A	445	46 (35–58)	50 (35–68)
Country B	130	8 (4–34)	2 (0-11)
Country C	147	12 (4-20)	14 (1–28)
Country D	103	1 (0–11)	0 (0–3)
Country E	106	11 (7–20)	8 (1–14)
Country F	110	6 (3–19)	0 (0-13)
Country H	65	36 (13-62)	36 (5–66)
All	7289	14 (12–16)	14 (12–16)
All nonendurance	1329	3 (0–8)	1 (0-3)
All endurance	4999	18 (15–22)	19 (16–22)

<sup>&</sup>lt;sup>b</sup> 95% CI estimated by bootstrapping methods, with any negative estimates rounded toward 0%.

# Paradigm Change





doi:10.1038/nature24286

Nature (2017) | doi:10.1038/nature24286

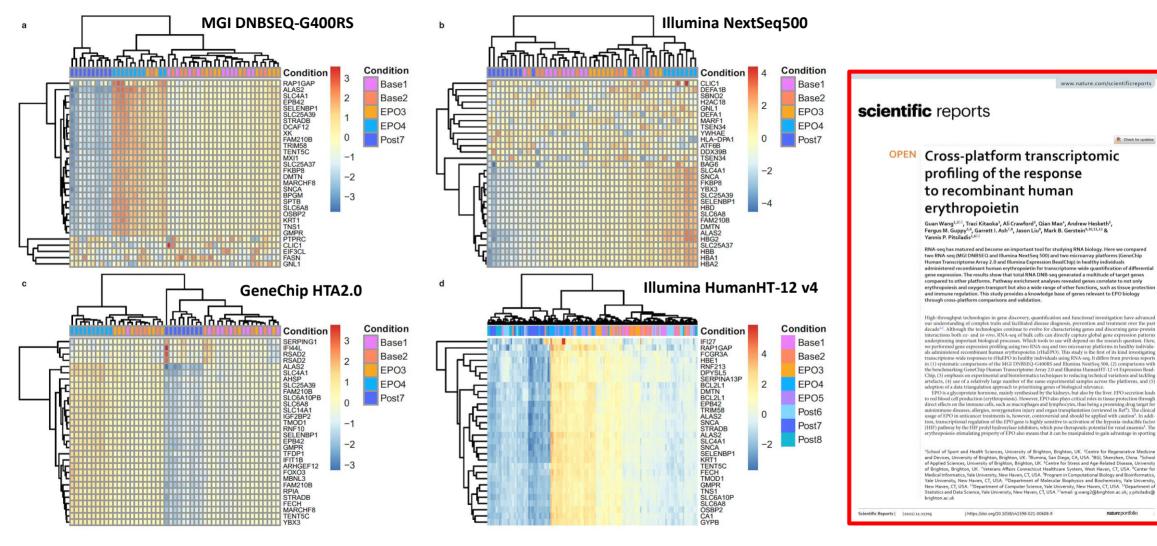
Received 13 July 2017 | Accepted 21 September 2017 | Published online 11 October 2017

# DNA sequencing at 40: past, present and future

Jay Shendure<sup>1,2</sup>, Shankar Balasubramanian<sup>3,4</sup>, George M. Church<sup>5</sup>, Walter Gilbert<sup>6</sup>, Jane Rogers<sup>7</sup>, Jeffery A. Schloss<sup>8</sup> & Robert H. Waterston<sup>1</sup>

This review commemorates the 40th anniversary of DNA sequencing, a period in which we have already witnessed multiple technological revolutions and a growth in scale from a few kilobases to the first human genome, and now to millions of human and a myriad of other genomes. DNA sequencing has been extensively and creatively repurposed, including as a 'counter' for a vast range of molecular phenomena. We predict that in the long view of history, the impact of DNA sequencing will be on a par with that of the microscope.

# Gene clustering analysis of the top 30 genes of high variance across the RNA-seq and the microarray platforms



**Figure 1.** Gene clustering analysis of the top 30 genes of high variance across the RNA-seq and the microarray platforms. (a) MGI DNBSEQ-G400RS, (b) Illumina NextSeq 500; (c) GeneChip HTA2.0; (d) Illumina HumanHT-12 v4 Expression BeadChip.

#### Effects of EPO on Blood Parameters and Running Performance in Kenyan Athletes

DIRESIBACHEW W. HAILE<sup>1,2,3</sup>, JÉRÔME DURUSSEL<sup>2</sup>, WONDYEFRAW MEKONEN<sup>1</sup>, NEFORD ONGARO<sup>3</sup> EDWIN ANJILA<sup>2</sup>, MARTIN MOOSES<sup>4</sup>, EVANGELIA DASKALAKI<sup>2</sup>, KERLI MOOSES<sup>4</sup>, JOHN D. MCCLURE<sup>2</sup>, SHAUN SUTEHALI<sup>2</sup>, and YANNIS P. PITSILADIS<sup>7,8</sup>

<sup>1</sup>Department of Physiology, College of Health Sciences, Addis Ababa University, Addis Ababa, ETHOPIA: <sup>2</sup>Institute of Cardiovascular and Medical Sciences, College of Medical, Veterinary and Idfe Sciences, University of Clargow, Glasgow, UNITED KINGDOM: <sup>2</sup>Department of Medical Physiology, School of Medicine, College of Health Sciences, Mol University, UNITEE KROLIME Digitablesis of salicity repositions, Sanon of Medicine, Custings of Irealist Scientists, 300 Customs, UNITEE KROLIME STATES, AND CUSTOMS AND CUSTO University of Rome "Foro Italien " Rome ITALY

every 2 d for 4 wk and responses or hemorlobin concentration (HGB) were h rHulipo administration. KEN had highe in SCO 6-17% vs.-10% respec

normoxic conditions. For ex to 6 wk of rHuEpo administ uptake (VO<sub>2max</sub>) by 6% to capacity at 80% of VO<sub>2max</sub> by

Address for correspondence: Yant Centre of Sports Medicine, Univ Kingdom University of Brighton Hill 7SR United Kingdom; E-mail: Y.PS Submitted for publication October 2 Accepted for publication July 2018.

## Metabolomic Proviling of Recombinant

#### Erythropoietin (rH Epo) in Trained Caucasian Athletes Giscard Lima, PhD, "† Jonathan Shurlock, BS

Shaun Sutehall, BSc,§ Yannis P, Pitsiladis, Mf

Objective: Recombinant human eyimropoite in 44-800 is prohibited choice for many changing distilles without by exact desiction transpround metabodimics significant or in 48-50 using an untergeted approach in bodo parameters below the significant properties of the significant properties of the significant below desicalles significant below the significant below the significant below desicalles significant below the significant which parameters of port of weekled parameters of the significant below the significant parameters of significant below the significant below the significant consideration weekled and significant discontinuous significant below the significant discontinuous discont total secun meascrates, and recommander metabolism (critical metabolism) control in the foliation included alterations of pyrimdine metabolism (critate and dihydrocrotate) and any-foe etaild carmiline), metabolic pathways that are associated with eightropolesis or eightrocyte Conclusions: Philininiary metabolic algorithms of histigo administration were identified. Fur validate these encouraging results in independent cohorts and with orthogonal techniques, such signatures derived from other "omics" analyses of ri-lulino administration (eq. transcriptomics Key Words: rHuEpo, metabolomics, mass spectrometry, anti-doping, metabolic signatures, si (Clin J Sport Med 2022;00:1-12)

#### INTRODUCTION

Recombinant human erythropoietin (rHuEpo) is a synthetic analogue of the endogenous hormone erythropoietin (EPO) ubmitted for publication May 18, 2021; accepted January 3, 2022.

From the "School of Sport and Health Sciences, University of Brighton, Brighton, Unite Ringdom; "University of Rome From Maloo," Rome, Italy; "Sport and Everoise Science

Supported by a research grant from the World Anti-Diping Agency (12009P). A. CPRessondo was supported by funds from the RATIGARISTIES from the National statute of Cleans and Markous Science, and ROTH 4-844C, ROTH. 44074. ROTH, 146151, and RPTH, 150002 from the National Heart, Lung, and Blood Institute.

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Volume 00 • Number 00 • Month 2022

increased oxygen delivery to the e Indirect methods of rHuEno detection a moderate success due to limited specificity as As such, novel strategies to detect rFfuEPO dop-unbiased and systematic fashion have become the recent research with the development of techniques al for both targeted and untargeted analyses for va endogenous and exogenous substances. "Omics" technolo gies such as metabolomics, have become a significant focus o anti-doning research with the goal to maximize the notential of these technologies to improve the longitudinal profiling of individual athlete haematological variables; the so called Athlete Biological Passport (ABP) launched by World Anti-

Athlete Biological Passport (ABP) Jaunenned by World Anti-poping Agency (WADA) in 2009. <sup>16</sup>

The use of metabolomic-based analytical techniques are rapidly growing within the fields of cancer cell metabolism and treatment, "nutrition," and disease prognosis, <sup>16,11</sup>

Bus Metabolomics allows for thousands of metaboliste to be investigated within a biological sample at any given time 1000. frame, creating a "snapshot" of the biological state of an organism, 12 with the biological (phenotypic) state inferred by profiling different biofluids. For example, in recent years, our to printed tool and are provided to the 4th or profiled to the first printed tool and are provided to the 4th or profiled to the 4th or provided t

group leveraged metabolomics approaches to determine the Countries 2022 Notice Klassy Houth, Inc. 44 patrs resona impact of high-altitude hypoxia-stimulated erythropoiesis on http://dx.doi.org/10.1007/JSM.0000000000001048

remained the most popular method of altitude training, with minor modifications on the design such as performing all but high-intensity training sessions at "high" altitude to maximize

(2500 m above sea level) while training at low altitude (<1500 m above sea level, LHTL) to attenuate the decrease in training intensity seen in previous LHTH studies. It was found that both training paradigms (LHTL and LHTH) led to an improvement in VO<sub>2</sub>max but only LHTL led to a significant increase in sea level

\$000-m time trial performance. The LHTL paradigm ha

Validation of whole-blood transcriptome signature during microdose recombinant human erythropoietin (rHuEpo) administration

real? Jonathan Shurlock<sup>3</sup> Martin Montar<sup>4</sup> Norinski Eulo<sup>5</sup> Geomie Bruinselc<sup>6</sup> Andrew Murray<sup>2</sup>, Brendan Yee<sup>8</sup>, Anne Keenan<sup>2</sup>, John D. McClure<sup>2</sup>

#### **ABT**

**BMC Genomics** 

of the adaptive model, 15 gene

Haemoglobin Mass and Running Time Trial Performance after Recombinant Human Erythropoietin Administration

Jérôme Durussel<sup>1</sup>, Evangelia Daskalaki<sup>2</sup>, Martin Anderson<sup>1</sup>, Tushar Chatterji<sup>1</sup>,

Yannis P. Pitsiladis 1

Diresibachew H. Wondimu<sup>1,3</sup>, Neal Padmanabhan<sup>1</sup>, Rajan K. Patel<sup>1</sup>, John D. McClure<sup>1</sup>

uld be detected in 14 trained subjects converse (EDR) and a 15 fold-MSR were differentially expressed iges were found 30 min after week after returning from

#### scientific reports

#### Cross-platform transcriptomic

#### mg of the response o recombinant human erythropoietin

Guan Wang<sup>1,263</sup>, Traci Kitaoka<sup>3</sup>, Ali Crawford<sup>3</sup>, Qian Mao<sup>4</sup>, Andrew Hesketh<sup>5</sup>, Fergus M. Guppy<sup>5,6</sup>, Garrett I. Ash<sup>7,3</sup>, Jason Liu<sup>9</sup>, Mark B. Gerstein<sup>5,10,13,12</sup> & Yannis P. Pitsiladis<sup>1,663</sup>

RNA-seq has matured and become an important tool for studying RNA biology. Here we compared two RNA-seq (MG DMSES Can dilliminis MexitSeq 500) and now microarray platforms (GeneChip) Human Transcriptome Array 2.0 and Illumina Expression Beed Chip) in healthy individuals administed red recombinant human septiosel for the zerostionne-wide quantification of differential gene expression. The results show that total RNA DNB-seq generated a multitude of target genes compared to other platforms. Pathway micriment analyses revealed genes correlate to not only rythropolesis and oxygen transport but also a wide range of other functions, such as tissue prond immune regulation. This study provides a knowledge base of genes relevant to EPO biology

# **⊟ wada**

- Altitude
- EPO
- Transfusions

#### Perspective

Shaun Sutehall, Fernanda Malinsky, Sven Voss, Neil Chester, Xu Xiao and Yannis Pitsiladis\*

# Practical steps to develop a transcriptomic test for blood doping

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Abstract: Blood doping remains a significant problem that threatens the integrity of sport. The current indirect method for detecting blood doping involves the longitudinal measurement of an athlete's haematological variables and identification of suspicious results that are indicative of doping (i.e., the athlete biological passport). While this has played a significant role in the protection of clean sport, improvements are needed. The development of a transcriptomic test, that can be used to detect the use of blood doping has been discussed and researched for decades and yet, an anti-doping test that can be adopted by the World Anti-Doping Agency (WADA) is yet to be established. With recent advancements in our understanding, as well as in methods of sample collection, the possibility of a transcriptomic test that can be used by WADA, is imminent. There are, however, several practical considerations that must first be made, that will be highlighted in this perspective article.

**Keywords:** anti-doping; testing; blood doping; EPO; transcriptomics

#### Introduction

In endurance sports, blood doping has the potential to significantly improve endurance performance, primarily by increasing the oxygen carrying capacity of the users' blood [1].

Baptist University, Hong Kong, China **Sven Voss**, Institute of Doping Analysis and Sport Biochemistry, Kreischa,

**Neil Chester**, Research Institute for Sport and Exercise Sciences, Liverpool John Moores University, Liverpool, UK

Xu Xiao, MGI Tech, Shenzhen, China

The use of rHuEpo to enhance athletic performance was prohibited by the IOC in 1990, however its detection to this day remains a challenge. In 1997, a "no start" rule was introduced by the UCI, with athletes providing a blood sample prior to competition [2] and if their HCT exceeded the predetermined limit, they were not permitted to race. The ABP was later introduced by the UCI in 2008 [3] and soon after adopted by WADA. Since then, it has been a critical element of anti-doping testing programmes.

The ABP measures an athlete's haematological variables longitudinally, and with the use of Bayesian statistics, generates an individual upper and lower limit [3]. If an athlete's haematological variables exceed the calculated upper or lower limit, it suggests that there is only a 1:100 chance that this is due to "natural" physiological variance [4] and can be used as evidence of doping and result in the sanctioning of an athlete [5]. It has, however, been demonstrated that it is possible to use low doses of rHuEpo and avoid detection by the ABP [6], and despite improvements to the ABP [7], questions remain about its reliability as a method of detecting doping and its efficacy as a deterrent, on a global scale [8].

A transcriptomic method of detecting rHuEpo abuse was described in 2001 [9], a year after the first direct test for rHuEpo in urine was established [10]. Recent research into transcriptomics for anti-doping has focused on identifying specific transcriptomic markers that are associated with rHuEpo abuse [11, 12], confounding variables such as altitude exposure [13] and reproducibility of transcriptomic markers in differing sample collection matrixes such as DBS [14].

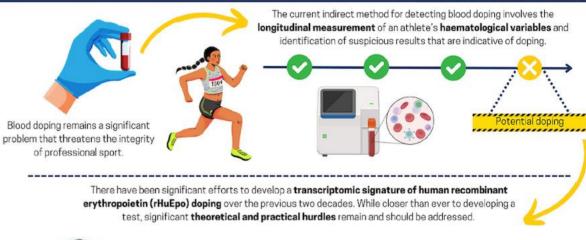
Despite significant research, a transcriptomic test, that is either standalone, or an addition to the current ABP has yet to be established. There are several theoretical and practical hurdles that must be overcome prior to the adoption of a transcriptomic test as an anti-doping tool. The summary of this article is presented in Figure 1.

#### **Identifying transcriptomic markers**

Numerous studies have characterised the transcriptomic response to blood doping in whole blood (e.g., Table 1, [15]) and DBS [16], along with conditions that may confound

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# Practical steps to develop a transcriptomic test for blood doping





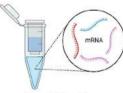




Laboratory sample processing



Blood collection and long-term storage



Identifying the transcriptomic markers

#### CONCLUSION

It is hoped that this article will assist research and anti-doping centres internationally adopt transcriptomic testing, in a WADA-compliant manner, to improve anti-doping efforts.







Figure 1: Graphical representation of this article. Figure created with BioRender.

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# My thoughts on Blood Doping

- A haematologic profile approach alone will not be able to resolve the current problem
- More sensitive methods required (i.e. gene expression)
- The notion of the "athletes passport" needs to be further investigated
- Consortium approach needed (e.g. governing bodies, scientists, statisticians, doctors, athletes, managers, lawyers)
- Serious money, not token gestures



#### Leadership

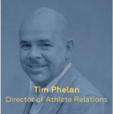






Team









## **The Enhanced Games**

The Movement





Investors



Scientific and Medical Advisory Commission













Athletes Advisory Commission













## superhumanity is on.

anced Games, the ultimate at the human body is capable



## The world's best athletes, enhanced.

When 44% of athletes already use performance enhancements, it is time to safely celebrate science.

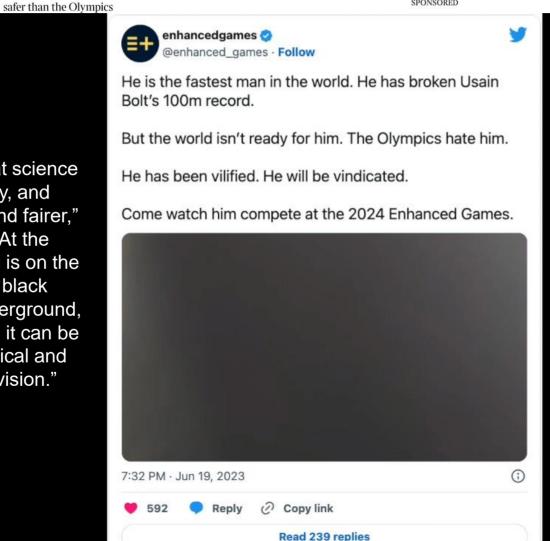
THE TIMES Today's sections V Past six days Explore V Times Radio Times-Sport ATHLETICS | MATT | AWTON Team GB athletes interested in competing in Olympics for dopers Australian plans to stage annual event free of drug testing that he claims would be

He founded Sargon, a technology infrastructure company for the pensions and superannuation industry that has more than £25 billion under management and supervision. D'Souza sold his stake in the company in 2018.

He clearly considers himself something of a visionary but he also describes himself as an "agitator", and in this case the International Olympic Committee (IOC) is very much in his sights.

SPONSORED

"We believe that science makes humanity, and sports, better and fairer," D'Souza said. "At the moment doping is on the dark web. It's a black market, it's underground, unsafe. With us it can be done under clinical and scientific supervision."



A handful of former Olympic stars have nevertheless joined Enhanced's athletes advisory commission. e.g., swimming gold medallist Ronald Schoeman. Right now, the plan is to have five sports categories athletics, swimming, weightlifting, gymnastics and combat sports — with the event held annually.

D'Souza hopes to attract athletes with the comfort of test-free competition and the chance, unlike the majority of Olympic athletes, to make some serious cash. "We will offer them equity".

## **Ignore History at Your Peril**

Editorial

#### We want healthy athletes and fair competition

Yannis P Pitsiladis <sup>o</sup> , <sup>1,2</sup> Fabio Pigozzi, <sup>1,3</sup> Michael Geistlinger, <sup>1,4</sup> José Kawazoe Lazzoli, <sup>1,5</sup> Maher Zahar, <sup>1,6</sup> Norbert Bachl, <sup>1,7</sup> Pedro Manonelles Margueta, 1,8 William O Roberts (1),19 Klaus-Michael Braumann. 1,10 Eduardo H De Rose Demitri Constantinou o, 1,11 Anca Ionescu, 1,12 Christian Schneider. 1,13 Patrick Shu-Hang Yung, 1,14 Xavier Bigard 0, 1,15 International Federation of Sports Medicine (FIMS)

#### THE ENHANCED GAMES

The Enhanced Games<sup>1</sup> is a planned international multisport event masterminded by Aron D'Souza, Australian businessman, where athletes will not be subject to any form of testing by the World Anti-Doping Agency (WADA) for prohibited list of substances and methods. Initially planned to be held in Australia in 2024. after the Paris 2024 Olympics, the event is now destined to take place in 2025. Reactions from the sporting world have been extremely negative, highlighting the dangers of encouraging performanceenhancing drugs (box 1).

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15 Injon Cycliste Internationale Ainle Switzerland

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Notably, some countries, such as Austria, France and Italy, have criminalised the use of a World Anti-Doping Code (WADC) Prohibited Substance while many more countries have also enacted sports-specific legislation that criminalises other Anti-Doping Rule Violations beyond the use of a WADC Prohibited Substance including (1) the possession of WADC Prohibited Substances or Methods; (2) the supply or distribution of WADC Prohibited Substances or Methods (trafficking); (3) the administration or prescription of WADC Prohibited Substances or Methods to athletes and (4) failing to cooperate with antidoping investigations being conducted by an International or National Anti-Doping

#### Organization or other public authority." THE INTERNATIONAL FEDERATION OF SPORTS MEDICINE POSITION

Despite much opposition, the Enhanced Games in some forms are likely to happen given the financial backing.3 The International Federation of Sports Medicine (FIMS), an international organisation comprising national sports medicine associations that span all five continents, with an executive committee that includes representative members from IOC, WADA, Association of Summer Olympic International Federations and Association of International Olympic Winter Sports Federations, and whose mission is to assist athletes in achieving optimal performance by high-quality medical care and training,4 is adamantly opposed to the Enhanced Games.

The President of the Enhanced Games holds the opinion that 'Drug testing is about fairness, not safety.' He states as follows: 'At the Olympics, athletes are drug tested after each competition. Clearly, the goal of the Olympics' drug testing regime is to ensure fairness through the threat of punishment for those who use enhancements. There is little

Box 1 Reactions from the sporting world highlighting the dangers of encouraging performance-enhancing

'There's only one message and that is if anybody is moronic enough to feel that they want to take part in that, and they are from the traditional, philosophical end of our sport, they'll get banned and they'll get banned for a long time.'12

'Joining with WADA other governments and the International Olympic Committee to express deep concerns over both the planned 'Enhanced Games' without antidoping requirements and the Russian organised 'Friendship Games' alternative to the Olympics'

'The idea of the Enhanced Games does not merit any comment. If you want to destroy any concept of fair play and fair competition in sport, this would be a good

way to do it '14 Members of the Executive Committee (ExCo) of the World Anti-Doping Agency (WADA) also affirmed their opposition to the so-called 'Enhanced Games', an event that would encourage the use of performance-enhancing drugs by athletes. The ExCo was united in condemning the 'Enhanced Games' as a dangerous and irresponsible concept. The health and well-being of athletes is WADA's number-one priority. Clearly, this event would jeopardise both by promoting the use of potentially harmful substances and methods. As we have seen throughout history, performance-enhancing drugs have taken a terrible physical and mental toll on many athletes. Some have died. The ExCo stressed WADA's position on this subject. warning athletes and support personnel, who wish to participate in clean sport, that if they were to take part, they would risk committing Anti-Doping Rule Violations under the Code. To be clear WADA will encourage anti-doping organisations worldwide to test involved athletes before. during and after this event, in order to protect the integrity of legitimate sport

Lord Coe, President of World Athletics12 Dr. Rahul Gunta Director of the White House Office of National Drug Control Policy<sup>13</sup>: Statement from the International Olympic Committee (IOC)<sup>14</sup>; Statement from

concern for safety, as the Olympics have no mandated health-testing programme. Olympians, 44% of whom have used enhancements in the past year, are able to compete regardless of any undiscovered health risks. This is why

athletes competing in the Enhanced Games will receive free medical screening prior to competition, which will ensure that they are tion with the President of the Enhanced healthy and able to compete. The screening will involve an echocardiogram, blood tests and genomic sequencing, in order to build up a full picture of the athlete's health, and allow athletes to make an informed decision about whether they are safe to compete. The Enhanced Games aim to be the safest sporting event in the world. At the Enhanced Games. we prioritise athlete safety. In partnership with top scientists and clinicians, we are

developing a full-system medical precompe-

tition screening protocol—to protect athletes

competing in the Enhanced Games.

FIMS strongly distances itself from this kind of interpretation of competitive sport. In citing the difference between fairness and safety, the president of the Enhanced Games seems to move the discussion onto a philosophical level. The priority of FIMS must be to safeguard the three cardinal principles that form the basis of medical care of athletes without exception: scientific analysis, study of the physical condition of the individual and protection of health.6 The claim that antidoping tests do not protect the safety of elite athletes has no scientific basis and therefore, this claim can vehemently be dismissed. Notably, for a substance or method to be added to the WADA Prohibited List, it must meet at least two of three criteria of Article 4.3 for Including Substances and Methods on the Prohibited List,7 that includes an actual or potential health risk to athletes (in addition to having the potential to enhance or enhances sport performance and violates the spirit of sport). Even if there is a reduced health concern to elite athletes participating in the Enhanced Games due to the declared medical controls-although this remains to be determined. Our concern as FIMS also focuses on the health hazards of drug use in young aspiring athletes. especially those who will never make it into the Enhanced Games. The grave concern of FIMS is that young individuals will be exploited in the quest for fame and fortune and the allure of the

#### MEDICAL SCREENING INSUFFICIENT TO GUARANTEE THE SAFETY OF 'ENHANCED' ATHLETES

Enhanced Games.

The procedures and protocols under development by the proponents of the Enhanced Games are reported to safeguard athlete health while upholding the

principle of personal choice in the use of enhancements (personal communica-Games). It is their intention to conduct numerous evaluations of athletes participating in the Enhanced Games to inform a multitiered, risk-based approach to informed consent, athlete's eligibility to compete and health insurer coverage. In scenarios where athletes do not meet the specified health criteria for participation, athletes (it would appear), will have the option to appeal the decision or seek continued involvement. This process under development is expected to involve an established system of external arbitration allowing an impartial third party to review the case and consider any appeals or arguments put forth by the athlete regarding their health status and eligibility. The Enhanced Games also intend to apply a similar system of external arbitration to athletes wishing to contest any sporting and/or financial sanctions that may be imposed in the event of alleged use of prohibited enhancements-what enhancements will be permitted and prohibited remain to be declared. The Enhanced Games policy under development will consider all relevant factors in determining whether an athlete can be held responsible and what penalties should be imposed. Considerations will include the athlete's intent, fault, negligence or knowledge of use, as well as a realistic assessment of

#### TESTING FOR FAIRNESS AND HEALTH

sporting advantage gained.

FIMS welcomes medical screening capable of discovering possible health risks for competitors, Indeed, antidoping is not just to guarantee fairness. The fundamental rationale of the WADC is that 'antidoping programmes seek to protect the health of Athletes and to provide the opportunity for athletes to pursue human excellence without the Use of Prohibited Substances and Probibited Methods,3 Furthermore, the statement above by the President of the Enhanced Games disregards the current distribution of competencies in Olympic sports. The medical screening of athletes to ensure that they are healthy and fit to compete is organised by the Olympic International Federations and not by the IOC itself. Each federation administers their own sport and is responsible for the health of their athletes when competing internationally, including at the Olympic Games. This typically entails them overseeing, or obliging, their members to be responsible for the medical screening of athletes to minimise the risk to their health. As a

major tenet of antidoping is the health of athletes, antidoping is an essential part of the all-encompassing medical care for healthy athletes. An athlete, who participates in doping, above all else endangers their own health. FIMS, therefore, holds that medical screening as such is not sufficient to guarantee the safety of athletes. For example, many of the side effects of banned substances will only be observed after several years of use and will not be detected by a single blood test. There are also many examples where screening is ineffective in detecting the long-term side effects of banned substances, such as psychiatric disorders induced by anabolicandrogenic steroids,8 the effects of growth hormone on reproductive function, the acceptable level of left ventricular hypertrophy or the risk of henatocellular carcinoma. (eg,9). Clearly, the type of medical assessment suggested in the context of Enhanced Games will not be sufficient to detect the side effects of many banned

#### ACCESS TO RESOURCES

Another consideration relates to access to financial and medical resources. Of concern to FIMS is how aspiring athletes, attracted to the offerings of the Enhanced Games but without significant financial resources, and athletes from developing countries, will access such specialist medical care and 'protect' their health. It is well publicised that enhancements are being prescribed under medical supervision to interested celebrities, influencers and the very rich in exclusive clinics, where a single 1-hour consultation can cost a few thousand dollars. A further unfortunate effect of the Enhanced Games is a likely further widening of the gap between developing and wealthier countries in sporting performance.

#### ANTI-DOPING MUST EVOLVE

Despite what one may think about the Enhanced Games, its inevitability could ironically help detect those who dope by better understanding the methods used to dope. Currently, due to the secrecy of the drug culture in sport, nobody knows what the dopers are up to until it is too late. As a result, preventative measures are not as effective as they could be and the exact adverse health implications for athletes are yet to be determined. There is also the reputational damage to sport to consider. Sports fans do not know what events are clean, how much doping is going on or how efficient International Federations, the IOC, WADA and other antidoping

organisations are at detecting those who dope. And, of course, any detection and punishment is often after the fact.10 Clean athletes are cheated out of a place on the podium and fans are left in the dark-wondering, cynical or just losing interest. It remains to be determined if the Enhanced Games can bring doping into a light where it can be properly studied. This may involve an uncomfortable evolution of antidoping science so ethical research could be conducted during doping to allow us to understand the life cycle of different substances and/or methods and their effects on elite performance, the body and health, FIMS and antidoping organisations armed with this information could then come up with smarter more directed ways to detect drug use in sport and better inform athletes regarding potential adverse health effects of drug use. With this rigorously researched and peer-reviewed knowledge, antidoping organisations could shorten the periodisation window thus making masking of drug use more difficult and determine the best combination of detection processes to use to make testing more effective.

It would appear that athletes participating in the Enhanced Games must participate in longitudinal studies in order externally neer reviewed to compete for the riches being offered, then FIMS and antidoping organisations could gather precise data for the first time from a large cohort of athletes who have admitted to doping to determine the long-term effects of various types of drug use. This information could be used to demonstrate the true health consequences of drug use or to dispel myths and inform decisions about the prohibited list. Ironically, the inevitable negative health impact of performance-enhancing drugs could act to discourage athletes from participating in the Enhanced Games and the increased knowledge on doping practices and the development of new antidoping tests could also strengthen the trust of athletes in drug free Olympic sport, However, all these possibilities and options cannot overrule the commitment of all sports and medical people involved to comply with the WADA. FIMS must insist that the Enhanced Games respect the ethical principles of sport medicine. Like never before

in the history of modern day sport, there is an urgent need to protect the ideals of the Olympics and Olympism. FIMS remains thoroughly dedicated to the Olympic values and committed to the Olympic Movement Medical Code. 11

Collaborators International Federation of Sports Medicine (FIMS): The International Federation of Sports Medicine (Fódération Internationale de Mádacina du Sport or EIMS) is an international organisation comprising national sports medicing associations that span all five continents, with an Executive that includes representative members from IOC, WADA, Association of Summer Olympi International Federations (ASOIF) and Association of International Olympic Winter Sports Federations (AIOWF) and fundamental nurnose to assist athletes in achieving optimal performance by high-quality medical care and training (www.fims.org)

Contributors YPP wrote the first draft of the manuscript, and all authors critically revised subsequent versions until all authors could approve the final manuscript. YPP is the quarantor

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Competing interests YPP, FP and MZ are members of the IOC Medical and Scientific Commission, All authors are members of the Executive Committee of the International Federation of Sports Medicine (FIMS)

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**Editorial** 

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#### Drugs, sex, and enhancement: Threats to sports integrity at the **Olympics**

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The Olympic Games represents a majestic sporting showcase on a uniquely tage (Figure 1). global scale. The Paris 2024 Olympics could mark the start of a new era for the Olympic Movement. This viewpoint focuses on three existential threats to modern Olympic Games: doping, gender issues, and enhancement, while cutting-edge research and science need to be at the heart of a concerted reaction to restore integrity of competition and protect the concept of "true" sport. Performance enhancement in the context of sport denotes or relates to a drug or other substance or method used with the prospect of improving an athlete's performance: specifically, substances and methods that enhance any number of components of physical and mental performance, such as agility, speed, strength, power, endurance, and competition preparation and execution.

#### ANTI-DOPING CHALLENGES

The World Anti-Doping Agency (WADA) was established in 1999 to harmonize doping control and provide educational strategies to avoid doping. However, some 20 years later, the prevalence of doping in sport has not substantially abated. While the true prevalence of doping in sport is impossible to accurately determine, it is substantially higher than the adverse analytical findings reported. For example, the figures for adverse analytical findings have varied between 1.1% (2008) and 1.6% (2016), while a recent large systematic review of 175 studies published between 1975 and 2019 reported doping prevalence rates in competitive sport ranging from 0% to 73% for doping behavior, with most falling under 5%. It is important, therefore, that new methods are devel-

of choice by those wishing to cheat and gain a significant performance advan-

Despite a number of positive developments in anti-doping, the last major development aimed at providing a step change in blood doping detection was the Athlete Biological Passport (ABP) and implemented in 2009. While urine testing is the main method of detecting recombinant human erythropoietin (rHuEpo), its relatively short window of detection (less than 24 h in some cases) renders this method ineffective for blood doping detection. The hematological module of the ABP uses Bayesian inference techniques and longitudinal measurements of blood parameters. Despite its appeal, the ABP has important limitations due to the simplicity of metrics and its impact by confounding factors such as hydration status and the natural hematological response to altitude training that complicate interpretation of ABP profiles. The ABP is also unable to effectively detect administration of micro-doses of rHuEpo, given the relatively small fluctuations in hematological parameters.2 Therefore, there is an urgent need for a paradigm shift away from "in sport, the cheats are usually a step ahead" to "in sport, the testers need to be a few steps ahead."

A potential game changer that has, until now, not been used substantially in the anti-doping field is "omics" technologies; genome, transcriptome, proteome. and metabolome. In a series of closely interconnected studies conducted in our laboratory and funded by WADA and the International Olympic Committee (IOC) using "omics" from microarrays to RNA sequencing technology, we successfully identified, replicated, and validated the blood "molecular signature" of rHuEpo administration,3 indicating that "omics" technologies can improve current oped with greater sensitivity and specificity to detect blood doping—a method anti-doping methods such as the ABP. The results we have generated to date,

## Thank you

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replicated by others, confirm that it is only a matter of when rather than if "omics" ment after only short-term use; for example, psychiatric disorders induced by methods will revolutionize anti-doping.

#### GENDER ISSUES: INTEGRATION OF INTERSEX AND TRANSGENDER ATHLETES INTO ELITE SPORT Sport is historically designated by binary categorization of male and female

that conflicts with the science of sex and modern society. This frequently overlooked fact, appreciated since antiquity, has come to the fore more recently with attempts to include transgender athletes and athletes with differences in sex development (DSD) into elite sports.4 The issue is that the performanceenhancing effect of testosterone, which is widely accepted in the literature. and the exposure to higher concentrations of testosterone in trans women and DSD women athletes is expected to provide a competitive performance advantage. This expectation has led to an ever-increasing number of international federations (IFs) banning participation of trans female athletes in elite sports and the same expected for DSD athletes. While these policy rules and their backing assumptions to primarily ban these athletes from competing may, in time, be proven correct, until data from longitudinal transgender and DSD athlete case comparison studies that control for variations in hormonal exposure and involve numerous indices of performance become available, there is just as much circumstantial evidence to support the policy to ban these athletes than there is to oppose it. The International Federation of Sports Medicine (FIMS) published its position with the European Federation of Sports Medicine Associations on the IOC framework on fairness, inclusion, and non-discrimination based on gender identity and sex variations, highlighting the need for innovative longitudinal research studies with specific athlete populations to generate physiological and sport performance knowledge and more reliable biomarkers for a fairer classification of athletes. The statement highlighted the fact that serum testosterone can be a valuable marker to regulate the inclusion of transgender athletes and DSD athletes into the elite female category but has limitations due to the multifactorial nature of sports performance. This led the authors of the joint position to conclude that longitudinal and sport-specific gender affirmation hormone therapy studies are needed to enhance our knowledge of impact of testosterone on performance and identify biomarkers of testosterone sensitivity/responsiveness, including molecular tools to determine the functional status of androgen receptors. Subsequently, the IOC published its position after consulting with many stakeholders, including FIMS. In general, it re-affirmed the position of FIMS but, also, importantly, assigned to individual IFs the responsibility to establish their own rules to ensure equitable and fair competition. How this complex situation resolves will depend on the outcomes of research studies using appropriate experimental designs and state-of-the-art technologies that remain to be commissioned and how individual IFs prioritize fairness, safety, and inclusion in sport. There is also the need to consider the ever-changing legal landscape, where more and more countries are supporting the right to self-identify. Policy makers must also consider the Olympic Charter, which clearly stipulates that "The enjoyment of the rights and freedoms set forth in this Olympic Charter shall be secured without discrimination of any kind, such as race, colour, sex, sexual orientation, language, religion, political or other opinion, national or social origin, property, birth or other status."

#### ENHANCED GAMES

The Enhanced Games is a planned annual international event involving five sports (track and field, swimming, gymnastics, weightlifting, and combat sports). masterminded by Aron D'Souza, an Australian businessman, where athletes will not be tested by WADA. The premise of the Enhanced Games is to safeguard athlete health while upholding the principle of personal choice to use any enhancements. Despite much opposition, the inaugural event is destined to happen in 2025. The primary concern, especially of the sports medicine community,5 is the health hazards of drug use in young aspiring athletes, who could be exploited in the guest for fame and fortune and the allure of the Enhanced Games. Although the Enhanced Games plan to conduct medical screening capable of discovering possible health risks for competitors, it is obvious that medical screening is insufficient to detect the side effects of many banned substances and methods. Many health side effects of banned substances and methods can be expected only after several years of use and not even with a most comprehensive medical assess-

anabolic-androgenic steroids or the risk of hepatocellular carcinoma.

Despite what one may think about the Enhanced Games, ironically, its launch may help detect those who cheat by better understanding the methods used to done. Due to the secrecy of the drug culture in sport, there is limited knowledge about sophisticated doping practices. A particular difficulty in anti-doping science is the inaccessibility to samples from doped athletes to allow new testing to be developed, improved, and perfected. With this in mind, the Enhanced Games could bring doping into focus, where it can be properly studied, but this would involve an uncomfortable evolution of anti-doping science so ethical research could be conducted during doping associated with the Enhanced Games to allow anti-doping science to better understand the life cycle of different substances and/or methods and their effects on elite performance and health. This partnership of convenience between the Enhanced Games and anti-doping organizations could result in smarter, more effective ways to detect drug use in sport while at the same time informing athletes about the true adverse health effects of drug use.

#### PROTECTING THE CONCEPT OF "TRUE" SPORT

In this viewpoint, we highlight three major threats to the future of the Olympics, but we also highlight solutions that are firmly based on science, experimentation, innovation, and application of cutting-edge technologies. Common to the doping and gender issues we address, it is the real or perceived unfairness that underpins both of these existential threats for Olympic sport, but also limited consensus on what constitutes "fairness." Part of the confusion is due to the use of terms to set the rules of sport, such as "the spirit of sport," that have no universal meaning. For example, WADA states that anti-doping policies have been developed to protect certain values in sport. The World Anti-Doping Code 2021 states that "The spirit of sport is the celebration of human spirit, body and mind. It is the essence of Olympism and is reflected in values we find in and through sport, including: Health; Ethics; fair play and honesty; Athletes' rights as set forth in the Code; Excellence in performance; Character and Education; Fun and joy, Teamwork; Dedication and commitment; Respect for rules and laws; Respect for self and other Participants; Courage; Community and solidarity. The spirit of sport is expressed in how we play true. Doping is fundamentally contrary to the spirit of sport." But such descriptions can only serve their purpose if these terms mean the same thing to different societies and people, which clearly is not the case. For example, the Enhanced Games use the same term, "spirit of sport," very differently. The only way forward is for each sport to decide what constitutes the "true" essence of their sport and prioritize its main constituents, such as inclusion, safety, and fairness. Such a decision-making process to achieve meaningful competition is already being explored in the integration of trans women into elite competition. Once there is consensus, it is essential that all stakeholders who participate in that sport must agree to play by the rules, and these rules need to be enforceable so that everyone has a chance to succeed, albeit not an equal chance. This decision-making approach and a concerted effort involving cutting-edge research and scientific data are urgently needed and represent the best response to the three major threats to integrity of competition we will witness in Paris in 2024

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