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Physical activity and semen quality among men attending an infertility clinic

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Abstract

OBJECTIVE—To examine the association between regular physical activity and semen quality.

DESIGN—Prospective cohort study.

SETTING—Couples attending one of three IVF clinics in the greater Boston area during 1993–2003. At study entry, male participants completed a questionnaire about their general health, medical history, and physical activity. Odds ratios (OR) and 95% confidence intervals (CI) were derived using generalized estimating equations models, accounting for potential confounders and multiple samples per man.

PATIENTS—A total of 2,261 men contributing 4,565 fresh semen samples were enrolled before undergoing their first IVF cycle.

INTERVENTION—None.

MAIN OUTCOME MEASURES—Semen volume, sperm concentration, sperm motility, sperm morphology, and total motile sperm (TMS).

RESULTS—Overall, none of the semen parameters was materially associated with regular exercise. Compared with no regular exercise, bicycling ≥5 hours per week was associated with low sperm concentration (OR=1.92, 95%CI=1.03–3.56) and low TMS (OR=2.05, 95%CI=1.19–3.56). These associations did not vary appreciably by age, body mass index, or history of male factor infertility.

CONCLUSIONS—While the present study suggests no overall association between regular physical activity and semen quality, bicycling at levels of ≥ 5 hours per week was associated with lower sperm concentration and TMS.

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Keywords

physical activity; bicycling; sperm motility; sperm count; male infertility; IVF

INTRODUCTION

The American College of Sports Medicine and the American Heart Association recommend at least 30 minutes of moderate exercise on 5 days each week or at least 20 minutes of vigorous exercise on 3 days each week (1). While exercise has been associated with many health benefits, including reduced risks of obesity, diabetes, cardiovascular disease, and some cancers (2), the relation between exercise and male fertility has not been well-studied.

Male athletes, particularly long-distance runners, have been shown to have reduced testosterone levels (3–12). In addition, elevated scrotal temperature associated with certain types of exercise may cause atrophy of the testicular germinal epithelium and adversely affect spermatogenesis (4,13,14). Bicycle riding has been linked to several genitourinary problems (15,16), and two studies among competitive cyclists reported changes in semen parameters including abnormal sperm morphology (17) and reduced sperm motility during periods of racing (14).

We prospectively examined the association between regular exercise and semen quality among men attending an infertility clinic in Massachusetts during 1993–2003. The few studies that have evaluated this association have been small and confined to competitive athletes or have evaluated only a single type of exercise (4,14,17). The present study, based on a larger general male population, addresses gaps in the literature on exercise and male fertility.

MATERIALS AND METHODS

We used data from the male partner of couples undergoing infertility treatment during 1993–2003 at three IVF clinics in the greater Boston area as part of a collaborative prospective study sponsored by the NICHD. The study's primary aim was to identify predictors of IVF outcomes, including regular exercise. Couples consented to a review of their medical records and completed a self-administered questionnaire before starting IVF treatment. Men were asked to provide at least one semen sample. Sixty-five percent of couples approached agreed to participate in the study. The study was approved by the Brigham and Women's Hospital Institutional Review Board, and patients gave written informed consent before enrolment.

The baseline questionnaire included questions about medical history, lifestyle, behavioral factors, and exercise questions phrased as follows: 1) Do you regularly exercise? 2) If so, as an adult, how many years have you engaged in regular exercise? 3) How many hours per week do you exercise? 4) How many months per year do you exercise? and 5) What is your most frequent type of exercise? Participants could list more than one type of exercise.

Exercise

Our primary exposure of interest was total metabolic equivalent (MET)-hours of physical activity per week, a measure of exercise intensity. Exercise intensities were defined according to the compendium of physical activities (18). To estimate total METs per week, we summed the METs from all activity types and computed the average METs per participant across all activities. We then multiplied the average METs by the total hours of activity reported per week. We also categorized men according to their average exercise

intensity for all activities in which they were engaged, with <4 METs defined as "low," 4–6 METs "moderate," and \geq 7 METs "vigorous." The latter variable did not take into account exercise frequency. Finally, we grouped men according to frequency of their *primary type* of exercise (\leq 2, 3–4, \geq 5 hours/week).

Semen parameters

Men were requested to observe a 2- to 5-day abstinence period before providing a semen sample. At each study visit, men collected semen samples by masturbation at the clinic and samples were analyzed within 20 minutes of collection on average (range: 15–60 minutes). Sperm concentration assessments were evaluated by hemacytometer (Improved Neubauer; Hauser Scientific Inc., Horsham, PA). Ejaculate volumes were measured using a standard 10ml laboratory pipette. Percent motile sperm was counted in a MicroCell chamber (Conception Technologies, San Diego, CA) and refers to percentage of sperm with any flagellar movement, whether twitching or progressive. Motility was analyzed using the World Health Organization (WHO) 1999 definition (19). Seminal smears were prepared at each clinical center where they were Papanicalou stained, analyzed, and stored, and where a single technician assessed sperm morphology using either the WHO (19) or Kruger method (20,21), depending on the center. Total motile sperm (TMS, units: 10⁶) was computed by multiplying sperm concentration by semen volume by percent motile sperm (22).

Exclusions

A total of 2,481 men, contributing 5,737 semen samples (IVF cycles), were enrolled. We excluded 848 samples with missing or implausible values on all semen parameters, 119 frozen samples, 96 non-ejaculate samples, and 109 samples from men with missing exercise data. After these exclusions, 2,261 men contributing 4,565 fresh semen samples remained. Excluded participants were similar to those included with respect to mean age (37.6 vs. 36.9 years), exercise intensity (16.5 vs. 16.2 METs), college education (34.5 vs. 35.2%), current smoking (11.9 vs. 10.1%), and multivitamin use (44.2 vs. 45.6%), but were more likely to have a history of male factor infertility (40.9 vs. 26.4%). Approximately 17% and 26% of couples discontinued treatment after the first and second IVF cycle, respectively, with discontinuation being more prevalent among older and parous women (23).

Data analysis

Exercise categories were based on their frequency distribution in the sample. Primary outcomes were sperm concentration, semen volume, percent motility, TMS, and percent normal morphology. We examined the odds of being classified below WHO thresholds (19) based on sperm concentration (<20 million/ml), volume (<2 ml), and motility (<50% motile sperm). For TMS, we used a cutpoint of below the 25^{th} percentile (<23 × 10^6 motile sperm). While WHO has no comparable cutoff for morphology, we defined "abnormal morphology" as normal forms <30% by standard criteria (19) and <4% by Kruger criteria (20).

We used generalized estimating equations to calculate odds ratios (ORs) and 95% confidence intervals (CIs) for the relation between exercise and semen parameters, accounting for multiple samples contributed per man. The mean (and median) number of samples provided per man was two (range: 1–10), with 1,024 (45%) providing one semen sample, 613 (27%) two samples, 355 (16%) three samples, and 269 (12%) four or more samples. Men reporting no regular exercise were the reference group in all analyses.

We controlled for known or suspected confounders, including age (years), body mass index (BMI=kg/m²), multivitamin use (yes vs. no), history of urogenital anomalies (yes vs. no), history of prostate or sexually transmitted infections (yes vs. no), history of male factor infertility (yes vs. no), current alcohol consumption (g/week), current caffeine consumption

(mg/week), smoking (current, former, never), type of underwear most commonly worn (boxers, briefs, boxers & briefs), hypertension (yes vs. no), education (≤ high school, some college or vocational, college, graduate school), and race/ethnicity (white vs. non-white). Missing data were modeled using indicator variables. We also controlled for study site and study wave (1993–1998; 1999–2003) to account for any differences in referral patterns or time period.

P-values for interaction were obtained from a likelihood ratio test comparing models with and without cross-product terms of each semen characteristic with age and BMI. All P values were based on two-tailed tests, with statistical significance indicated by P=0.05 (95% confidence interval excluding one). SAS statistical software (version 9.1) was used for all analyses (24).

RESULTS

The median age of participants was 36 years (interquartile range: 33–40 years), median BMI was 22.7 kg/m^2 (interquartile range: 20.8– 25.7 kg/m^2), and 91% were white (data not shown). About 26% reported fathering a previous pregnancy, 26% reported a history of male factor infertility, and 13% had received IVF or GIFT previously. Of the men who exercised regularly, the median number of hours exercised per week was 4.0 (interquartile range: 2.5–6.0 hours), the median METs of exercise averaged across all exercise types was 7.0 (interquartile range: 4.8–8.0), and the median total MET-hours/ week was 24.0 (interquartile range: 14.0–36.0).

Baseline characteristics of the sample according to MET-hours/week of exercise are shown in Table 1. Men reporting regular exercise were more likely to be educated, lean, and use multivitamins, and were less likely to smoke, consume caffeine, or have hypertension. Positive associations were found between low TMS and male age (OR=1.04 per year, p<0.001), BMI (OR=1.03 per kg/m², p<0.001), history of urogenital anomaly (OR=1.48, p=0.02), education (\leq high school vs. graduate school; OR=1.54, p=0.03), and history male factor infertility (OR=8.93, p<0.001). Abnormal sperm morphology was associated with male factor infertility only (OR=2.77, p<0.001).

None of the semen parameters was materially associated with regular exercise, MET-hours/ week of activity, or average exercise intensity (Table 2). When we examined the relation between type of exercise and semen parameters (Table 3), bicycling was associated with low sperm concentration and low TMS. Compared with no regular exercise, ORs associated with bicycling ≤2, 3–4, and ≥5 hours per week were 1.31 (95% CI=0.57–3.02), 2.09 (95% CI=0.90–4.83), and 1.92 (95% CI=1.03–3.56), respectively, for low sperm concentration (*P*-trend=0.01) and 0.77 (95% CI=0.34–1.76), 1.44 (95% CI=0.69–3.01), and 2.05 (95% CI=1.19–3.56), respectively, for low TMS (*P*trend=0.02).

Associations of bicycling with low sperm concentration and low TMS persisted across categories of BMI and age. ORs for low sperm concentration comparing \geq 5 hours/week of bicycling with no regular exercise were 2.33 (95% CI=1.04–5.23; *P*-trend=0.01) among men with BMI <25; 2.06 (95% CI=0.64–6.60; *P*-trend=0.24) among men with BMI \geq 25 (*P*-interaction by BMI=0.92); 2.08 (95% CI=0.86–5.02; *P*-trend=0.11) among men aged <40; and 1.49 (95% CI=0.52–4.30; *P*-trend=0.10) among men aged \geq 40 (*P*-interaction by age=0.21). ORs for low TMS comparing \geq 5 hours/week of bicycling with no regular exercise were 2.15 (95% CI=1.16–3.96; *P*-trend=0.05) among men with BMI <25; 2.57 (95% CI=0.80–8.23; *P*-trend=0.12) among men with BMI \geq 25 (*P*-interaction by BMI=0.77); 1.80 (95% CI=0.80–4.10; *P*-trend=0.44) among men aged <40; and 2.16 (95% CI=0.87–5.32; *P*trend=0.01) among men aged \geq 40 (*P*-interaction by age=0.31). In addition,

associations of bicycling with low sperm concentration and TMS were apparent when we confined the analysis to: 1) men without a history of male factor infertility, or 2) the first semen sample provided by each man (N=2,261), albeit the ORs were weaker and less precise (data not shown).

DISCUSSION

In the present study, there was little evidence for an overall effect of regular exercise on sperm concentration, volume, sperm motility, TMS, or morphology. However, in the subgroup of men who reported bicycling as their primary form of exercise, bicycling at levels of ≥ 5 hours per week was associated with low sperm concentration and TMS. These findings, from a study of moderate exercisers, generally agree with previous studies that have shown deleterious effects of bicycling on semen parameters among competitive cyclists (14,17).

The few studies that have evaluated the relation between exercise and semen quality have been confined to one or two types of activity, or have examined endurance or high-intensity athletic training only, making it difficult to compare results. Our null results for jogging and running do not support the few studies that have shown significant alterations in sperm count (5), concentration (4,5,11), motility (4,5,11), and morphology (4,11) among endurance-trained runners (4), high-mileage runners (5), and high-intensity long-term treadmill runners (11). However, the runners in our study had more moderate levels of exercise, potentially limiting statistical power to detect a true effect. A previous study found that "high-mileage" runners (mean: 108 km/wk) experienced significant alterations in semen quality relative to sedentary men (5), but not runners of "moderate mileage" (mean: 54 km/wk), similar to what another study found among three groups of athletes differing in exercise intensity and frequency (12). Therefore, another explanation is that running at the levels observed in our study may not influence semen quality.

Study limitations include self-reported exercise data and the evaluation of activity for current exercisers only. For example, lifetime exercisers who eliminated their activity prior to completing the baseline questionnaire would have been misclassified as non-exercisers. Furthermore, in the analyses of exercise subtypes, there was no information on the actual intensity of the activity. For instance, two men who reported bicycling ≥5 hours/week could have had vastly different intensities of bicycling (e.g., leisurely cycling for transportation versus competitive cycling). Such errors in classification would generally dilute our effect estimates.

Although exclusions at baseline were similar according to several risk factors for infertility, men reporting a history of male factor infertility were more likely to be excluded. Couples whose IVF treatment included donor sperm were ineligible for enrollment, precluding analysis of their data. While such exclusions may have reduced variability in semen quality, it is unlikely that these exclusions were a significant source of bias because our findings persisted among men with and without a history of male factor infertility. Finally, our results did not differ markedly by the number of semen samples provided over follow-up, suggesting that results did not depend on success with IVF treatment.

Study strengths include the large sample size and the collection of multiple cycles of data over a 10-year period. Because men reported their exercise and covariate data before the provision of a semen sample, reporting errors are not likely to depend on the measurement of semen quality. Thus, our effect estimates comparing extreme categories of exercise are probably conservative. We controlled for a wide range of potential confounders such as age,

BMI, smoking, and history of infections. Given all outcome data were abstracted from medical records, we had nearly complete information on the study endpoints of interest.

A link between reduced sperm concentration and bicycling—the one class of exercise to show an effect—has some biologic plausibility. Bicycling has been linked to genitourinary problems (15,16), including nerve entrapment syndromes (50–91% of cyclists), erectile dysfunction (13–24%), and other less common symptoms (priapism, penile thrombosis, hematuria, torsion of spermatic cord, perineal nodular induration, and prostatitis) (16). Two studies among long-distance competitive cyclists reported changes in semen parameters including a lower percentage of sperm with normal morphology (17), a higher proportion of morphologically abnormal tapered forms (17), and reduced sperm motility (14). Moreover, small changes in total and free testosterone have been observed in male athletes (3–12), potentially affecting spermatogenesis. It remains unclear as to whether the changes associated with bicycling are due to mechanical trauma (i.e., caused by compression of scrotum on bicycle saddle), to a prolonged increase in core scrotal temperature (i.e., related to exercise itself or wearing of constrictive clothing), or some other factor (16). A report among moderate bicyclers showed no correlation between cycling duration and scrotal temperature (25).

Because men who attend infertility clinics differ from the general population of reproductive-aged men, caution should be used when interpreting our findings. If those attending infertility clinics differ with respect to exercise levels *and* semen quality, our results may not extend to a larger population of men. However, it is unlikely that the exercise levels of men in our study differed materially from those of the general population. Moreover, because associations were uniform across levels of age and BMI, our findings, if real, may extend to other groups of men with similar characteristics.

In summary, in this study of moderate exercisers, no association was observed between regular exercise and semen parameters overall, but bicycling at levels of ≥5 hours per week was associated with reduced sperm concentration and TMS. Our findings warrant confirmation in larger studies of moderate bicyclers and men from the general population.

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Table 1

Baseline characteristics among 2,261 male participants according to weekly level of exercise (IVF Study, 1993-2003)

			Re	Regular exercise		
Characteristic ^a	No exercise (N=975)	All categories (N=1,286)	<20 MET- hrs/wk (N=503)	20–39 MET- hrs/wk (N=489)	≥40 MET- hrs/wk (N=294)	P-trend ^b
Age, years (mean)	36.8	37.1	37.3	36.9	37.2	0.84
White (%)	90.3	92.2	89.4	95.1	92.6	0.01
Body mass index, kg/m ² (mean)	25.0	23.2	23.4	22.9	23.1	<0.001
Education (%)						
Less than College	36.5	19.5	18.5	18.5	22.6	<0.001
College	36.1	35.6	34.8	35.7	36.2	0.97
Graduate school	27.4	44.9	46.6	45.8	41.2	<0.001
Multivitamin use (%)	39.1	50.7	50.0	48.8	54.0	<0.001
Current smoker (%)	16.4	5.2	5.2	4.4	6.5	<0.001
Pack-years of smoking (mean)	5.0	2.2	2.6	1.9	1.9	<0.001
Alcohol, g/wk (mean)	0.99	2.99	6.09	7.07	69.2	0.30
Caffeine, mg/wk (mean)	1,970	1,702	1,694	1,674	1,771	0.003
Diabetes treated with medication (%)	1.0	6.0	0.5	9.0	1.6	< 0.001
Hypertension (%)	10.5	5.6	5.2	7.0	2.9	<0.001
History of urogenital anomaly (%)	9.6	11.6	13.5	12.8	7.3	0.96
History of STD/prostate infection (%)	13.7	13.3	12.8	14.9	11.0	0.62
History of male factor infertility (%)	27.0	25.8	26.5	27.3	22.6	0.24
IVF or GIFT before entering study (%)	14.3	12.5	14.1	12.0	11.1	0.08
Wear briefs only (%)	56.5	49.6	52.1	50.6	43.3	<0.001
Baths preferred to showers (%)	3.3	2.0	1.8	1.2	4.2	0.41

N = number of male participants.

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a Characteristics are presented as means or percents within exercise categories and are standardized to age distribution of sample.

 $^{^{}b}$ Test of difference in baseline characteristic across physical activity categories (none, <20, 20–39, \geq 40 MET-hrs/wk).

Table 2

ORs for clinically-based cut points for semen parameters (WHO criteria) by weekly MET-hours and average intensity of exercise

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	Total N ^a	Low sperm concentration (<20×10 ⁶ /ml)	Low volume (<2 ml)	Low motility (<50%)	Low total motile concentration (<23×10 ⁶)	Abnormal morphology (<30% Standard/ <4% Kruger)
Number of samples	4,565	4,523	4,558	4,501	4,489	2,006
No regular exercise	1,951	$1.00~(\mathrm{Ref.})^b$	$1.00~(\mathrm{Ref.})^b$	$1.00~({ m Ref.})^b$	$1.00~(\mathrm{Ref.})^b$	$1.00~(\mathrm{Ref.})^b$
Regular exercise						
Any	2,614	1.22 (0.97–1.54)	0.97 (0.80–1.17)	0.97 (0.80–1.17) 1.00 (0.84–1.19)	0.93 (0.75–1.15)	1.10 (0.82-1.48)
MET-hours per week						
<20	984	1.27 (0.95–1.70)	0.92 (0.73–1.17)	1.04 (0.83–1.30)	0.89 (0.68–1.17)	1.17 (0.81–1.68)
20–39	1,030	1.26 (0.95–1.68)	0.88 (0.69–1.14)	0.90 (0.72–1.13)	0.90 (0.68–1.18)	1.02 (0.70–1.49)
≥40	009	1.07 (0.76–1.51)	1.19 (0.89–1.59)	1.10 (0.86–1.41)	1.05 (0.77–1.44)	1.15 (0.75–1.77)
P -trend c		0.30	0.63	0.95	0.89	0.64
Average intensity (METs)						
Low (<4)	402	1.32 (0.88–1.97)	1.03 (0.74–1.45)	1.03 (0.74–1.45) 1.16 (0.86–1.58)	0.94 (0.64–1.38)	1.46 (0.94–2.26)
Moderate (4–6)	299	1.27 (0.92–1.77)	0.85 (0.63-1.16)	1.07 (0.82–1.40)	0.85 (0.62–1.17)	1.57 (0.96–2.57)
Vigorous (≥7)	1,613	1.17 (0.90–1.52)	1.00 (0.81–1.24)	0.92 (0.76–1.12)	0.96 (0.75–1.22)	0.91 (0.65–1.27)
P -trend $^{\mathcal{C}}$		0.18	0.81	0.37	0.48	0.67

Values are odds ratios (95% confidence intervals) adjusted for age, body mass index, study wave, study site, multivitamin use, history of urogenital anomaly, history of STI or prostate infection, male factor infertility, alcohol, caffeine, smoking, wearing of boxers or briefs, hypertension, education, and race/ethnicity. Each column is restricted to number of samples shown.

a Maximum N of samples in exercise category; number may be lower depending on data available for given semen characteristic.

 b Reference group for within-column comparison.

crests for trend conducted by modeling the ordinal categorical version of each exercise variable and calculating p-value from the Wald test statistic, with men reporting no exercise comprising the lowest exposure category. Page 9

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Table 3

Odds ratios for clinically-based cut points for semen parameters (WHO criteria) by primary type of exercise

	Total N a	Low sperm concentration (<20×10 ⁶ /ml)	Low volume (<2 ml)	Low motility (<50%)	Low total motile concentration (<23×10 ⁶)	Abnormal morphology (<30% Standard/ <4% Kruger)
No regular exercise	1,951	$1.00~({ m Ref.})^b$	$1.00~(\mathrm{Ref.})^b$	$1.00~(\mathrm{Ref.})^b$	$1.00~({ m Ref.})^b$	$1.00~(\mathrm{Ref.})^b$
Regular exercise (primary type):						
Running/jogging						
Any	744	1.05 (0.77–1.42)	0.87 (0.65–1.15)	0.77 (0.60–1.01)	0.76 (0.56–1.04)	1.07 (0.70–1.62)
Hours per week						
≤2	223	0.93 (0.57-1.51)	0.75 (0.48–1.18)	0.60 (0.39-0.92)	0.60 (0.37–0.97)	0.69 (0.33–1.43)
3-4	265	1.03 (0.64–1.65)	0.62 (0.39–1.00)	0.81 (0.55-1.20)	0.69 (0.43–1.10)	1.13 (0.58–2.18)
25	256	1.18 (0.76–1.83)	1.26 (0.86–1.86)	0.91 (0.64–1.29)	1.03 (0.66–1.61)	1.45 (0.83–2.52)
P - $trend^{\mathcal{C}}$		0.54	96.0	0.27	0.43	0.28
Bicycling						
Any	231	1.73 (1.09–2.73)	0.90 (0.65–1.15)	1.07 (0.73–1.56)	1.37 (0.91–2.07)	0.67 (0.32–1.42)
Hours per week						
≤2	81	1.31 (0.57–3.02)	0.82 (0.45–1.47)	0.86 (0.45–1.66)	0.77 (0.34–1.76)	0.84 (0.31–2.29)
3-4	57	2.09 (0.90-4.83)	0.79 (0.34–1.83)	1.30 (0.66–2.59)	1.44 (0.69–3.01)	0.93 (0.22–3.94)
>5	93	1.92 (1.03–3.56)	1.05 (0.55–2.03)	1.10 (0.63–1.93)	2.05 (1.19–3.56)	0.38 (0.09-1.55)
P -trend $^{\mathcal{C}}$		0.01	0.84	09.0	0.02	0.21
Weightlifting						
Any	309	1.11 (0.74–1.67)	0.81 (0.57-1.15)	1.44 (1.03–2.00)	0.97 (0.68–1.37)	1.78 (1.04–3.05)
Hours per week						
≤2	51	1.27 (0.52–3.11)	0.92 (0.38–2.27)	2.05 (0.98–4.28)	1.22 (0.55–2.69)	8.36 (3.54–19.8)
3-4	94	1.23 (0.63–2.41)	0.60 (0.33-1.07)	1.66 (0.96–2.88)	0.73 (0.41–1.32)	0.95 (0.32–2.78)
≥5	164	1.01 (0.59–1.74)	0.91 (0.58–1.44)	1.18 (0.76–1.84)	1.03 (0.65–1.61)	1.49 (0.75–2.93)
P -trend c		0.76	0.34	0.12	0.85	0.15

Values are odds ratios (95% confidence intervals) adjusted for age, body mass index, study wave, study site, multivitamin use, history of urogenital anomaly, history of STI or prostate infection, male factor infertility, alcohol, caffeine, smoking, wearing of boxers or briefs, hypertension, education, and race/ethnicity.

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 $^{\it q}$ Maximum N in exercise category; number may be lower depending on data available for given semen characteristic.

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 $^{\it b}$ Reference group for within-column comparison.

Crests for trend conducted by modeling the ordinal categorical version of each exercise variable and calculating a p-value from the Wald test statistic, with men reporting no exercise comprising the lowest exposure category.