

1 **Title:**

2 Comparison of muscle activity, strength and balance, before and after a 6-month training using
3 the FIFA11+ program (part 2)

4

5 **Authors:**

6 Takeshi Oshima, M.D. Ph.D. ¹

7 Junsuke Nakase, M.D., Ph.D.¹

8 Anri Inaki, M.D., Ph.D.²

9 Takafumi Mochizuki, M.D., Ph.D.³

10 Yasushi Takata, M.D.¹

11 Kengo Shimozaki, M.D.¹

12 Seigo Kinuya, M.D., Ph.D.²

13 Hiroyuki Tsuchiya, M.D., Ph.D.¹

14

15 **Affiliations:**

16 1. Department of Orthopaedic Surgery, Graduate School of Medical Science, Kanazawa

17 University, 13-1 Takara-machi, Kanazawa 920-8641, Japan

18 2. Department of Nuclear Medicine/Biotracer Medicine, Graduate School of Medical Science

19 Kanazawa University, Kanazawa, Japan

20 3. Kanazawa Advanced Medical Center, Kanazawa, Japan

21

22 Corresponding author:

23 Junsuke Nakase, M.D., Ph.D.

24 Department of Orthopaedic Surgery, Graduate School of Medical Science, Kanazawa University

25 13-1 Takara-machi, Kanazawa 920-8641, Japan

26 TEL: +81-76-265-2374

27 FAX: +81-76-234-4261

28 E-mail: nakase1007@yahoo.co.jp

29

30 3176 words

31 **Abstract**

32 **Purpose:** Sports injuries can significantly impact an athlete's career, as well as impose a high
33 financial burden on teams. Therefore, the prevention of sports injuries is an essential aspect of
34 sports medicine. To evaluate the effects of a 6-month training period, using part 2 of the
35 FIFA11+, on the activation and strength of core and lower limb muscles, and on static and
36 dynamic balance performance.

37 **Materials and Methods:** Eight college male soccer players, 20.4 ± 0.5 years old, completed the
38 FIFA11+ program (part 2) at least 3x per week for 6 months. The following variables were
39 measured, before and after the 6-month training program: activity of more than 30 muscles
40 (with a focus on core and lower limb muscles), measured using the standardized uptake values
41 of 18F-fluorodeoxyglucose (FDG) on positron emission tomography (PET-CT); isokinetic
42 strength of the knee flexor and extensor and hip abductor muscles, measured at $60^\circ/s$; static
43 balance over a 60-s period, measured using a Gravicorder; and dynamic balance, measured
44 using the Star Excursion Balance Test.

45 **Results:** Training improved activity levels of core (obliquus externus abdominis and erector
46 spinae) and lower limb (tibialis anterior of the both legs) muscles ($p \leq 0.03$), corrected the
47 between-limb difference in activation of the semimembranosus and improved dynamic
48 balance, with a greater training effect on the non-dominant limb ($p \leq 0.02$). Training also

49 improved knee flexor force of the non-dominant lower limb ($p=0.02$).

50 **Conclusion:** Routine performance of the FIFA11+ (part 2) program can improve activation of

51 core and lower limb muscles, with a concomitant improvement in dynamic balance.

52 **Introduction**

53 Sports injuries can significantly impact an athlete's career and the financial aspect [1].

54 Therefore, the prevention of sports injuries has received increasing attention in sports

55 medicine. Generally, sports injury prevention programs include some combination of

56 plyometric, balance and agility exercises, and have been reported to be effective in decreasing

57 the incidence of injuries, regardless of sport activity level, sex and age [2,3]. 'FIFA11+' is one of

58 the most effective prevention programs, which the Fédération Internationale de Football

59 Association (FIFA) Medical and Assessment Research Center (F-MARC) has developed. The

60 FIFA11+ consists of three parts: basic running (part 1); 3 levels of difficulty of 6 exercises

61 aiming to increase strength (core and lower limbs), balance, muscle control (plyometrics), and

62 core stability (part 2); and running such as straight line running, or cutting activities (part 3).

63 Improvement and evaluation of the effectiveness of an injury prevention program

64 requires assessment of not only the change in the incidence rate of injury, but also the short-

65 and long-term effects of the training on modifying muscle activity patterns and improving,

66 strength and balance.

67 To evaluate the muscle activities, electromyography (EMG) has principally been used.

68 However, EMG can only provide information on the activation of superficial muscles, and not

69 of deep muscles of the trunk and limbs, is limited with regard to the number of muscles that
70 can be assessed simultaneously (namely those on which superficial sensors can be placed) and
71 requires equipment to be attached to the body, which is difficult during performance of sports
72 activities.

73 Previous studies have used whole-body positron emission tomography - computed
74 tomography (PET-CT) to quantify the change in muscle activity after performing the FIFA11+
75 (part 2) program, with glucose uptake in skeletal muscles being used as a proxy measure of the
76 level of muscle activity [4,5]. Unlike EMG, PET-CT provides a non-invasive observation of the
77 activity of muscles throughout the body, simultaneously, with the possibility of 3-dimensional
78 (3D) image reconstruction. As active muscle cells exhibit increased glucose uptake, the use of
79 ¹⁸F-fluorodeoxyglucose (FDG), a deoxy analog of glucose, permits the observation of glucose
80 metabolism of the skeletal muscles throughout the body. However, unlike glucose, FDG does
81 not continue along the usual glycolytic pathway but, rather, accumulates within exercising
82 muscle tissue. This metabolic trapping process forms the basis of FDG-PET. The accumulation
83 of FDG in muscle provides a parameter of glucose intake by muscles and, therefore, of the
84 intensity of muscle activity [6]. To our knowledge, FDG-PET is the only method that can
85 provide a reliable cumulative index of muscle activity for between-muscle comparison, which
86 is invaluable for the assessment of sports injuries. As such, FDG-PET would be effective to

87 measure the effects of the FIFA11+ program.

88 In recent years, various benefits of the entire FIFA11+ program have been reported,
89 including a reduction in the incidence of sports injuries and improvement in the
90 neuromuscular control and strength of flexor muscles [7,8]. A review of the studies reporting
91 on the acute or chronic effects of the FIFA11+ on performance and physiological measures
92 among football players, an intervention period of 9-10 weeks yielded positive effects [9].
93 However, the effect of a long-term, routine, performance of the FIFA11+ program on the
94 metabolism of skeletal muscles remains to be clearly defined. Therefore, the aim of our study
95 was to investigate the change in muscle activity, and muscle strength and dynamic balance,
96 after performing part 2 of the FIFA11+ for 6 months. The a priori hypothesis was that the
97 FIFA11+ (part 2) program would be effective in increasing the activity of core and lower limb
98 muscles, improve muscle strength of the lower limb and improve static and dynamic balance.

99 **Materials and Methods**

100 Our study group was formed of 8 collegiate male soccer players. All participants were
101 considered to be healthy, based on their medical history and physical examination, and none
102 were taking medication at the time of the study. All participants provided informed consent,
103 and the study was approved by our Institutional Ethics Review Board.

104 Participants were asked to avoid strenuous physical activity for at least one day prior
105 to testing, and to refrain from eating and drinking for at least 6 h before testing. PET-CT images
106 were obtained as per previously described methods [4,5]. After obtaining baseline
107 (pre-training) PET-CT images, participants completed the training protocol, consisting of
108 completing part 2 of the FIFA11+ program, ≥ 3 times per week, for 6 consecutive months.
109 PET-CT images were obtained at the end of the training period, using the same protocol as at
110 baseline.

111 Regions of interest (ROI) on the images were manually segmented in 30 skeletal
112 muscles, located in 5 areas of the body: trunk, pelvis, thigh, lower leg, and the foot. (Table 1)
113 All ROIs were identified by one experienced nuclear medicine specialist, who was blinded from
114 all other results, from the plain CT images obtained concurrently. The standardized uptake
115 value (SUV) of FDG was calculated by overlapping the defined ROI and fusion PET-CT images to
116 outline the area of muscles, being careful to not include large vessels. The FDG uptake was

117 normalized to the unit volume of muscle as follows: {mean ROI count (counts per second/pixel)
118 × calibration factor (counts per second /Bq)}/{injected dose (Bq)/body weight (g)}. ROIs were
119 defined for the skeletal muscles previously described, bilaterally, and the mean SUV was
120 compared for the dominant and non-dominant side of the body (where the dominant side was
121 identified by asking participants which leg they used to kick a ball). The mean SUV for the trunk
122 was calculated as follows: $([\text{left mean SUV} \times \text{left muscle area}] + [\text{right mean SUV} \times \text{right muscle}$
123 $\text{area}])/(\text{left muscle area} + \text{right muscle area})$. FDG accumulation was compared between the
124 pre- and post-training PET-CT examinations.

125 Balance and muscle strength testing was performed by experienced physical
126 therapists. These assessments were performed one week after the PET analysis, with balance
127 tests preceding strength tests to avoid effects of muscle fatigue.

128 Static balance was measured using a Gravicorder. Postural sway, for 60 s, at a
129 sampling rate of 20-Hz sampling, under the following conditions: two-leg stance with eyes
130 open and then with eyes closed; single leg (dominant) standing with eyes open; and single leg
131 (non-dominant) standing with eyes open. All measurements were obtained in bare feet, using
132 the center of the force platform as a reference point. Two variables of balance were measured,
133 the locus length per time (LG), providing a measure of attitude control, and the environmental
134 area (AR), providing a measure of equilibrium control [10]. These two parameters have been

135 used to assess dizziness and equilibrium disorders and, more recently, to quantify balance
136 effects on anterior cruciate ligament injury [11]. All balance measurements were repeated
137 twice, with a 1-min rest between measurements; data from the second measurement, which
138 has been reported to be more accurate [12], used for analysis.

139 Dynamic postural control was evaluated using the Star Excursion Balance Test (SEBT).
140 Participants were asked to reach as far as possible along the designated line for each of the
141 following 8 directions: anterolateral, anterior, anteromedial, medial, posteromedial, posterior,
142 posterolateral, and lateral. (Fig 1) The test was performed twice, once in a clockwise direction
143 (reaching with the right leg) and once in a counterclockwise direction (reaching with the left
144 leg). The average of the length of three reaches performed in each direction was used for
145 analysis, with the distance normalized to the length of the leg (measured from the anterior
146 superior iliac spine to the distal tip of the medial malleolus). The greater the normalized length
147 of excursion, the better the dynamic balance.

148 The maximal knee flexion and extension muscle strength and the maximal isokinetic
149 hip abductor strength were tested using an isokinetic Biodex system. After 10-min warm-up,
150 knee strength measurements were obtained in the sitting position, and the hip abductor
151 strength was obtained in a sidelying position. For the strength testing, participants were asked
152 to move the knee and hip at full force, with 3 trials performed for each direction at a speed of

153 60°/s. For all strength measurements, the average value of the 3 trials was used for analysis,
154 and values were normalized to body weight (pre- and post-training) for between-subject
155 comparisons. The ratio of the strong-to-weaker leg was calculated as an index of between-limb
156 strength imbalance, converted to a percentage difference, using a previously described
157 method based on log-transformed raw data, followed by back transformation [13].

158 All statistical analyses were performed using Stata for Mac Version 15 (Stata
159 Statistical Software 2017; Stata Corp LLC, College Station, TX, USA). All data are presented as
160 mean (SD). The Shapiro-Wilk test was used to evaluate the normality of distribution. Wilcoxon
161 signed rank test was used to evaluate differences in the mean SUV and static balance, before
162 and after training, with a paired t-test used to evaluate the differences in muscle strength and
163 dynamic balance. The minimum significance level was set at $P < 0.05$. The sample size was
164 confirmed using a power analysis of 0.8, with an α value of 0.05 and effect size of 1.0.

165

166 **Fig 1:** Reaching directions on the Star Excursion Balance Test

167

168

169 Results

170 The relevant characteristics of the participants at pre-training are as follows: age;
171 20.4±0.5 years old, height; 175.4±6.2 cm, weight; 68.6±5.1 kg, 22.3±1.3 kg/m², and the leg
172 length; 89.4±3.8 cm. After training, the weight was 70.1±4.6 kg (p= 0.246) and the body mass
173 index was 22.8±0.8 kg/m² (p= 0.250), and there was no significant difference pre- and
174 post-training.

175 Representative whole-body PET images, pre- and post-training, are shown in Fig 2
176 with the mean SUVs reported in Table 1. A significant pre- to post-training increase in the
177 mean SUV was identified for two core muscles, the obliquus externus abdominis (0.75±0.26
178 *versus* 1.06±0.38, respectively, p= 0.036) and erector spinae (0.67±0.16 *versus* 0.80±0.31,
179 respectively, p=0.025). The pre- to post-training significant change in the mean SUVs for the
180 muscles of the dominant and non-dominant lower limbs, from the pelvis to the foot, are
181 detected. For the dominant lower limb, the mean SUV increased for the tibialis anterior
182 (1.06±0.59 *versus* 1.53 ±0.86, respectively, p=0.017) and decreased for the triceps surae
183 (1.39±0.40 *versus* 0.88±0.19, respectively, p=0.017). A similar result was identified for the
184 non-dominant lower limb, with an increase in the tibialis anterior (1.00±0.36 *versus* 1.44±0.66,
185 respectively, p=0.025) and a decrease for the triceps surae (1.24±0.25 *versus* 0.86±0.18,
186 respectively, p=0.025). The significant side-to-side difference of SUV was detected in

187 semimembranosus in pre-training. Pre-training, the mean SUV of the semimembranosus
188 muscle was higher for the dominant than non-dominant lower limb (0.74 ± 0.14 versus
189 0.59 ± 0.10 , respectively, $p=0.012$). Of note, no significant difference in the activation of the
190 semimembranosus between the dominant and non-dominant side was observed after training.

191 The mean LG and AR values are reported in Table 2, with no significant difference
192 between pre- and post-training values. The reach distance along the 8 directions for the
193 dominant and non-dominant lower limbs are reported in the Table 3. For the dominant leg,
194 the standing reach distance increased significantly, pre- to post-training, in the anterior-lateral
195 direction (70.9 ± 9.2 cm to 74.9 ± 9.6 cm, $p=0.023$). A greater improvement in dynamic balance
196 on the non-dominant leg was observed, with an increase in the reach distance across multiple
197 directions, as follows: medial (103.6 ± 6.0 cm to 107.9 ± 7.6 cm, $p=0.002$); posterior-medial
198 (111.4 ± 6.0 cm to 115.3 ± 6.6 cm, $p=0.030$); and posterior (114.0 ± 4.5 cm to 119.1 ± 7.9 cm,
199 $p=0.022$). The pre- to post-training changes in muscle strength are reported in Table 4. For the
200 non-dominant leg, knee flexion force increased from 1.24 ± 0.15 Nm/kg to 1.39 ± 0.14 Nm/kg
201 ($p=0.023$). No effects of training on knee extensor and hip abductor strength were noted, nor
202 on the hamstring-to-quadriceps ratio or between-limb imbalance index (Table 5).

203 **Fig 2:** Representative whole-body positron emission tomography images acquired before (left)
204 and after (right) routine performance of the FIFA11+.

205 Table 1: Mean SUVs during pre- and post-training. SUV, standardized uptake value

Body area	Muscles	Pre-training SUV		Post-training SUV	
		Dominant leg	Non-dominant leg	Dominant leg	Non-dominant leg
Trunk	Rectus abdominis	0.90±0.35		1.03±0.40	
	Obliquus externus abdominis	0.75±0.26		1.06±0.38	
	Obliquus internus abdominis	0.69±0.14		0.76±0.81	
	Transversus abdominis	0.62±0.08		0.59±0.14	
	Psoas major	0.94±0.31		0.87±0.20	
	Quadratus lumborum	0.85±0.33		0.99±0.32	
	Erector spinae	0.67±0.16		0.80±0.31	
Pelvis	Gluteus maximus	1.54±0.78	1.25±0.78	1.27±0.29	1.17±0.29
	Gluteus medius	2.18±1.17	2.29±0.82	2.79±1.93	2.46±0.31
	Gluteus minimus	3.13±0.60	3.51±0.83	3.51±1.20	3.34±0.98
	Piriformis	2.95±1.70	2.37±0.88	2.13±0.74	1.99±0.65
Thigh	Quadriceps femoris	1.00±0.31	1.01±0.38	1.00±0.30	0.92±0.19

	Sartorius	0.76±0.21	0.77±0.20	0.91±0.32	0.89±0.27
	Gracilis	1.16±0.43	1.14±0.40	1.26±0.43	1.32±0.64
	Semimembranosus	0.74±0.14	0.59±0.10	0.62±0.08	0.60±0.12
	Semitendinosus	1.19±0.31	1.09±0.30	1.41±0.48	1.23±0.52
	Biceps femoris	0.59±0.08	0.63±0.10	0.61±0.10	0.62±0.13
	Adductor complex	0.81±0.18	0.82±0.18	0.83±0.14	0.76±0.15
Lower leg	Tibialis anterior	1.06±0.59	1.00±0.36	1.53±0.86	1.44±0.66
	Flexor digitorum longus	1.28±0.26	1.20±0.35	0.90±0.35	0.96±0.35
	Tibialis posterior	1.34±0.84	1.37±0.79	1.05±0.30	1.13±0.51
	Flexor hallucis longus	1.23±0.13	1.30±0.58	1.10±0.29	1.18±0.49
	Peroneus	1.50±0.89	1.32±0.36	1.13±0.61	1.21±0.52
	Triceps surae	1.39±0.40	1.24±0.25	0.88±0.19	0.86±0.18
Foot	Abductor hallucis	1.36±0.42	1.45±0.44	1.31±0.61	1.17±0.37
	Quadratus plantae	1.06±0.20	1.29±0.43	1.05±0.38	1.00±0.34
	Flexor digitorum brevis	1.43±0.42	1.61±0.58	1.68±1.49	1.49±0.67

	Abductor digiti minimi	1.13±0.44	1.45±0.55	1.15±0.50	1.31±0.78
	Flexor hallucis brevis	2.12±0.65	2.24±0.57	2.27±0.83	2.12±0.90
	Interosseous	2.02±0.61	2.20±1.11	2.38±1.50	2.20±1.31

206

207 Table 2: Static balance parameter, pre- and post-training.

208

	conditions	pre-training	post-training	Confidence interval	Effect size	P value
LG (cm/s)	Two-leg, eyes opened	0.94±0.11	1.05±0.14	[-0.04, 0.27]	0.904	0.12
	Two-leg, eyes closed	1.41±0.28	1.27±0.23	[-0.39, 0.11]	0.533	0.36
	Dominant leg, eyes opened	3.93±0.61	3.96±0.67	[-0.44, 0.51]	0.052	0.78
	Non-dominant leg, eyes opened	3.88±0.68	4.03±0.31	[-0.48, 0.79]	0.195	0.40
AR (cm ²)	Two-leg, eyes opened	1.69±0.64	1.81±0.40	[-0.44, 0.69]	0.181	0.48
	Two-leg, eyes closed	2.65±0.37	2.46±0.82	[-1.01, 0.62]	0.198	0.78
	Dominant leg, eyes opened	6.84±2.77	7.24±1.56	[-2.30, 3.11]	0.124	0.78
	Non-dominant leg, eyes opened	6.91±2.66	7.45±1.99	[-2.10, 3.18]	0.177	0.67

209 Table 3: Reach distance (excursion distance/leg length×100) by balance condition and direction of reach, pre- and post-training.

210

211

Direction (cm)	Dominant leg				Non-dominant leg			
	Pre-trainin g	Post-traini ng	Confidence interval	P value	Pre-trainin g	Post-traini ng	Confidence interval	P value
anterior-lateral	70.9±9.2	74.9±9.6	[0.74, 7.2]	0.02	76.9±9.9	75.3±9.3	[-7.3, 4.0]	0.51
anterior	86.4±8.3	86.6±6.9	[-1.3, 1.7]	0.75	86.9±7.5	88.1±7.4	[-0.08, 2.5]	0.06
anterior-medial	95.5±7.3	95.0±6.6	[-3.8, 3.0]	0.77	94.8±4.7	97.3±8.0	[-0.81, 6.0]	0.11
medial	103.4±7.2	105.0±7.3	[-1.3, 4.5]	0.23	103.6±6.0	107.9±7.6	[2.2, 6.3]	<0.01
posterior-medial	112.5±6.9	113.6±7.9	[-1.5, 3.6]	0.35	111.4±6.0	115.3±6.6	[0.49, 7.3]	0.03
posterior	115.7±6.4	117.4±7.0	[-2.9, 6.4]	0.40	114.0±4.5	119.1±7.9	[1.0, 9.1]	0.02
posterior-lateral	106.4±7.3	108.9±9.0	[-5.0, 10.1]	0.45	106.1±6.6	109.9±8.9	[-2.8, 10.3]	0.22
lateral	97.7±8.7	97.1±13.3	[-8.3, 7.1]	0.86	94.8±7.8	97.5±7.8	[-2.1, 7.5]	0.22

212 Table 4: Pre- to post-training change in lower limb muscle strength.

213

214

215

	pre-training	post-training	Confidence interval	Effect size	P value
Knee flexor (%)	17.0±22.3	6.5±4.9	[-28.5, 7.52]	0.487	0.22
Knee extensor (%)	12.5±8.2	12.3±8.6	[-10.7, 10.3]	0.016	0.96
Hip abductor (%)	9.9±7.9	6.7±5.0	[-10.5, 4.0]	0.373	0.33
H/Q ratio (%)	17.5±5.8	10.6±8.4	[-16.4, 2.7]	0.596	0.14

216 Table 5: Between-limb muscle strength imbalance, pre- and post-training.

217

218

	Dominant leg					Non-dominant leg				
	pre-training	post-training	Confidence interval	Effect size	P value	pre-training	post-training	Confidence interval	Effect size	P value
Knee flexor (Nm/kg)	2.90±0.09	2.99±0.18	[-0.47, 6.44]	0.134	0.72	2.78±0.17	2.98±0.13	[-0.10, 0.50]	0.550	0.16
Knee extensor (Nm/kg)	1.31±0.20	1.41±0.21	[-0.15, 0.34]	0.338	0.37	1.24±0.15	1.39±0.14	[0.03, 0.28]	1.026	0.02
Hip abductor (Nm/kg)	2.41±0.41	2.40±0.35	[-0.41, 0.38]	0.031	0.21	2.35±0.37	2.52±0.43	[-0.05, 0.39]	0.634	0.55
H/Q ratio (%)	0.45±0.05	0.48±0.07	[-0.02, 0.08]	0.454	0.16	0.46±0.09	0.48±0.09	[-0.05, 0.09]	0.220	0.78

219

220

221 Discussion

222 Our results indicate an increase in the activation of various skeletal muscles of the
223 core and lower limbs after a 6-month training using part 2 of the FIFA 11+ program, measured
224 as an increase in the uptake of glucose: obliquus externus abdominis, erector spinae and
225 tibialis anterior. Of note was the decrease in the glucose uptake of the triceps surae, as glucose
226 uptake increased in the tibialis anterior. We also noted an improvement in the imbalance of
227 the glucose uptake in the semimembranosus, between the dominant and non-dominant, after
228 training. From a functional perspective, training produced a greater improvement in dynamic
229 balance on the non-dominant than dominant lower limb. To our knowledge, this is the first
230 study to report changes in muscle activities, associated with improvements in balance and
231 muscle strength, after long-term training using the part 2 of the FIFA 11+ program. Observed
232 changes in glucose uptake, balance and strength would, therefore, be the key mechanisms
233 explaining the association between the FIFA 11+ and a decrease in sports-related injuries.

234 Glucose enters the muscle cell by facilitated diffusion via the glucose transporter-4
235 (GLUT4), with exercise stimulating an increase in the expression of GLUT4 in skeletal muscles,
236 as shown by the findings of Reichkender et al. after an 11-week program of daily moderate-
237 and high-dose aerobic exercise [14]. Similarly, an increase in GLUT4 levels in skeletal muscles is
238 a key adaptation to regular exercise training [15]. Thus, FDG accumulation in the muscle can be
239 used as a measure of the change in glucose uptake with training, as well as providing a proxy
240 measure of muscle activity [16].

241 By comparing the change in FDG accumulation of each muscle from pre- to
242 post-training, we demonstrated that routine training using part 2 of the FIFA11+ program
243 improved muscle metabolism and activation as well as the previous studies [4,5]. These
244 adaptations are important when we consider the positive effects of core and lower limb

245 strength on balance. Kaji et al. reported on the improvement in two-leg standing balance with
246 eyes closed after performing the FIFA11+ program ($p=0.005$) [17]. Granacher et al. reported on
247 the improvement in the Functional Reach test ($p<0.05$) after performing a core stability
248 training program which increased the strength of the trunk flexors ($p<0.001$), extensors
249 ($p<0.001$) and lateral flexors ($p<0.001$) [18]. In the same way, Imai et al. reported immediate
250 improvements in the posteromedial ($p<0.001$) and posterolateral directions ($p=0.002$) of the
251 SEBT after the trunk stabilization exercises [19]. Considering the effect of core stability on
252 balance, Willson et al. suggested that appropriate core strength training could reduce
253 sports-related injuries [20]. We reported similar findings, showing an increase in the mean
254 SUVs of the obliquus externus abdominis and erector spinae muscles after training, with a
255 concomitant improvement in dynamic balance, indicative of the effectiveness of the part 2
256 FIFA11+ program in improving core strength.

257 With regard to lower limb muscle activation, Day et al. reported an increase activity
258 of the tibialis anterior during active swaying (compared to static standing), which they
259 associated to the higher proprioceptive demands of balancing under more challenging sensory
260 conditions and the proprioceptive role of the tibialis anterior [21]. Similarly, Earl et al. reported
261 an increase in the general activity of lower limb muscles (vastus medialis obliquus, vastus
262 lateralis, medial hamstring, biceps femoris, and tibialis anterior) during the SEBT ($p<0.05$), with
263 the exception of the triceps surae muscles ($p=0.08$) [22]. We demonstrated comparable
264 findings post-training, supporting the effectiveness of the FIFA11+ (part 2) in improving
265 balance [8].

266 The balance of muscle strength is also an important component with regard to injury
267 prevention. For the lower limb, the hamstring-to-quadriceps ratio is an important risk factor of
268 injury [23]. Previous studies have reported on the effectiveness of the complete FIFA 11+

269 program in improving knee flexor strength and, thus, the hamstring-to-quadriceps ratio [7].
270 Between-limb strength imbalances might also be an important risk factor for lower leg injury
271 [24]. A prospective study provided evidence that a between-limb imbalance in eccentric knee
272 flexor strength increase in the risk of hamstring injuries [25]. Although we reported the
273 effectiveness of the training in eliminating the higher SUV of the semimembranosus muscles in
274 the dominant than non-dominant lower limb observed pre-training, we did not identify a
275 significant between-limb imbalance among our study participants, either pre- or post-training.
276 Therefore, we cannot stipulate if the FIFA11+ (part 2) program is effective to improve lower
277 limb muscle imbalances, although the results of our SUV analysis indicate that the program
278 would likely be of benefit in this regard.

279 Overall, our findings are consistent with previous reports on the effectiveness of
280 performing the complete FIFA 11+ program in improving balance and muscle strength, which
281 lowered the incidence of sports injuries [7–9]. The methods we used, and PET-CT in particular,
282 could be useful for evaluating the effectiveness of training programs and identifying the
283 underlying pathways.

284 We note the following limitations of our study. First, the FDG-PET method accounts
285 only for muscle glucose uptake. Other substrates, such as free fatty acids, muscle glycogen and
286 lactate, are also metabolized in active muscle cells. That being said, studies have confirmed
287 that glucose oxidation increases with exercise intensity and increases in glucose uptake, to
288 some extent, increases in proportion with glycogen utilization as exercise intensity increases. A
289 second limitation was the method we used to define the ROI. Since FDG uptake was measured
290 at an arbitrary site on the target muscle, it did not reflect the uptake of glucose for the entire
291 muscle. Lastly, taking into consideration the ethical dilemma of radiation exposure with CT
292 (even though the amount of FDG was <10% of normal PET examination), our study sample was

293 small and we did not include a control group. Our use of PET-CT was consistent with the aim of
294 our study to confirm that improvements in muscle metabolism was an important underlying
295 pathway for the previously reported effectiveness of the FIFA11+ training program. In this
296 sense, our use of PET-CT and our findings of an increase in glucose uptake post-training are
297 novel.

298

299 **Conclusions**

300 Routinely performing part 2 of the FIFA11+ program for 6 months increased glucose
301 uptake, related to muscle activity, of the obliquus externus abdominis, erector spinae and
302 tibialis anterior, while decreasing activation of the triceps surae. The training program also
303 improved knee flexor strength and dynamic balance, with no effect identified on static
304 balance. We speculate that these improvements could be beneficial in lower the risk of
305 sports-related injuries.

306

307 **Acknowledgments**

308 The authors would like to express their appreciation for the outstanding efforts
309 positive attitude of the participants. In addition, they are extremely grateful for the technical
310 assistance with measurement provided by physical therapists of our hospital.

311 No financial assistance was received for the project.

312

313 **Author Contributions**

314 Conceptualization: Takeshi Oshima, Junsuke Nakase.

315 Data curation: Takeshi Oshima, Anri Inaki.

316 Formal analysis: Takeshi Oshima, Anri Inaki, Takafumi Mochizuki.
317 Investigations: Takeshi Oshima, Yasushi Takata, Kengo Shimozaki.
318 Methodology: Takeshi Oshima, Anri Inaki, Takafumi Mochizuki, Junsuke Nakase.
319 Project administration: Seigo Kinuya, Hiroyuki Tsuchiya.
320 Resources: Takeshi Oshima, Yasushi Takata, Kengo Shimozaki, Junsuke Nakase.
321 Supervision: Seigo Kinuya, Hiroyuki Tsuchiya.
322 Validation: Takeshi Oshima, Anri Inaki, Takafumi Mochizuki.
323 Writing- original paper: Takeshi Oshima.
324 Writing- review and editing: Junsuke Nakase, Seigo Kinuya, Hiroyuki Tsuchiya.

325

326 **Conflict of interest statement**

327 This research didn't receive grants from any funding agency in the public, commercial
328 or not-for-profit sectors.

329

330 **Ethical approval**

331 This study was approved by the ethics committee of Kanazawa University (approval
332 number: 1286).

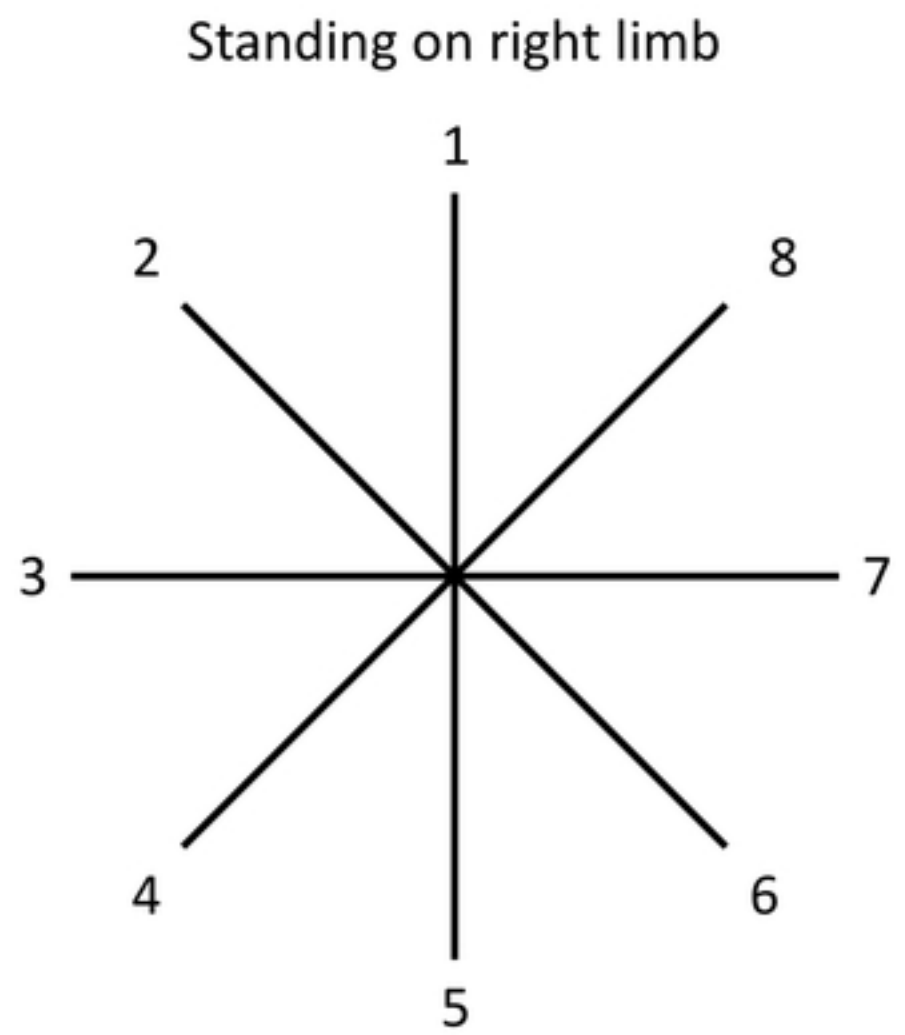
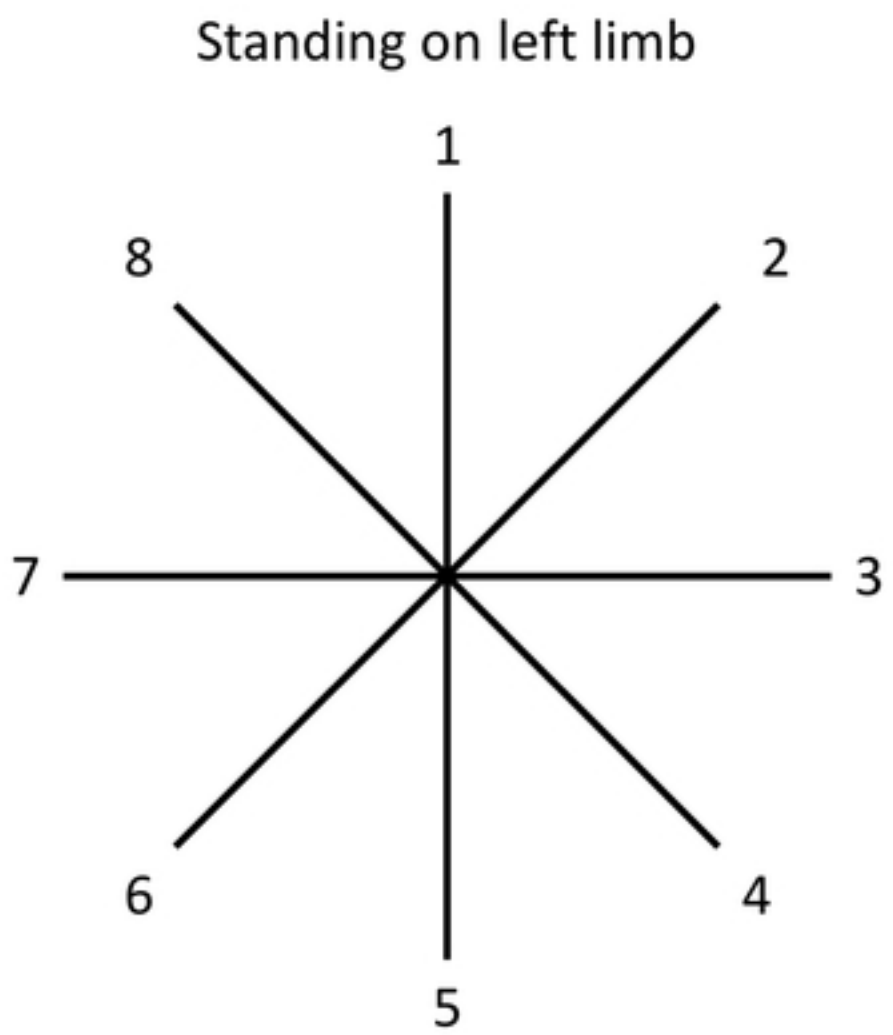
333 **References**

- 334 1. Junge A, Lamprecht M, Stamm H, Hasler H, Bizzini M, Tschopp M, et al. Countrywide
335 Campaign to Prevent Soccer Injuries in Swiss Amateur Players. *Am J Sports Med.*
336 2011;39: 57–63. doi:10.1177/0363546510377424
- 337 2. Mandelbaum BR, Silvers HJ, Watanabe DS, Knarr JF, Thomas SD, Griffin LY, et al.
338 Effectiveness of a Neuromuscular and Proprioceptive Training Program in Preventing
339 Anterior Cruciate Ligament Injuries in Female Athletes. *Am J Sports Med.* 2005;33:
340 1003–1010. doi:10.1177/0363546504272261
- 341 3. Zebis MK, Andersen LL, Brandt M, Myklebust G, Bencke J, Lauridsen HB, et al. Effects of
342 evidence-based prevention training on neuromuscular and biomechanical risk factors
343 for ACL injury in adolescent female athletes: a randomised controlled trial. *Br J Sports*
344 *Med.* 2016;50: 552–557. doi:10.1136/bjsports-2015-094776
- 345 4. Nakase J, Inaki A, Mochizuki T, Toratani T, Kosaka M, Ohashi Y, et al. Whole Body
346 Muscle Activity during the FIFA 11+ Program Evaluated by Positron Emission
347 Tomography. Chen C-T, editor. *PLoS One.* 2013;8: e73898.
348 doi:10.1371/journal.pone.0073898
- 349 5. Takata Y, Nakase J, Inaki A, Mochizuki T, Numata H, Oshima T, et al. Changes in muscle
350 activity after performing the FIFA 11+ programme part 2 for 4 weeks. *J Sports Sci.*
351 2016;34. doi:10.1080/02640414.2016.1149606
- 352 6. Ohnuma M, Sugita T, Kokubun S, Yamaguchi K, Rikimaru H. Muscle activity during a
353 dash shown by¹⁸F-fluorodeoxyglucose positron emission tomography. *Journal of*
354 *Orthopaedic Science.* 2006. pp. 42–45. doi:10.1007/s00776-005-0972-y
- 355 7. Impellizzeri FM, Bizzini M, Dvorak J, Pellegrini B, Schena F, Junge A. Physiological and
356 performance responses to the FIFA 11+ (part 2): a randomised controlled trial on the

- 357 training effects. *J Sports Sci.* 2013;31: 1491–1502. doi:10.1080/02640414.2013.802926
- 358 8. Daneshjoo A, Mokhtar AH, Rahnema N, Yusof A. The Effects of Comprehensive
359 Warm-Up Programs on Proprioception, Static and Dynamic Balance on Male Soccer
360 Players. Lucia A, editor. *PLoS One.* 2012;7: e51568. doi:10.1371/journal.pone.0051568
- 361 9. Barengo N, Meneses-Echávez J, Ramírez-Vélez R, Cohen D, Tovar G, Bautista J. The
362 Impact of the FIFA 11+ Training Program on Injury Prevention in Football Players: A
363 Systematic Review. *Int J Environ Res Public Health.* 2014;11: 11986–12000.
364 doi:10.3390/ijerph111111986
- 365 10. Okuda K, Abe N, Katayama Y, Senda M, Kuroda T, Inoue H. Effect of vision on postural
366 sway in anterior cruciate ligament injured knees. *J Orthop Sci.* 2005;10: 277–283.
367 doi:10.1007/s00776-005-0893-9
- 368 11. Oshima T, Nakase J, Kitaoka K, Shima Y, Numata H, Takata Y, et al. Poor static balance is
369 a risk factor for non-contact anterior cruciate ligament injury. *Arch Orthop Trauma*
370 *Surg.* 2018;138: 1713–1718. doi:10.1007/s00402-018-2984-z
- 371 12. Demura S, Yamaji S, Noda M, Kitabayashi T, Nagasawa Y. Examination of Parameters
372 Evaluating the Center of Foot Pressure in Static Standing Posture from the Viewpoints
373 of Trial-to-trial Reliability and Interrelationships Among Parameters. *Equilib Res.*
374 2001;60: 44–55. doi:10.3757/jser.60.44
- 375 13. Impellizzeri FM, Bizzini M, Rampinini E, Cereda F, Maffiuletti NA. Reliability of isokinetic
376 strength imbalance ratios measured using the Cybex NORM dynamometer. *Clin Physiol*
377 *Funct Imaging.* 2008;28: 113–119. doi:10.1519/JSC.0b013e3181c06bdd
- 378 14. Reichkender MH, Auerbach P, Rosenkilde M, Christensen AN, Holm S, Petersen MB, et
379 al. Exercise training favors increased insulin-stimulated glucose uptake in skeletal
380 muscle in contrast to adipose tissue: a randomized study using FDG PET imaging. *Am J*

- 381 Physiol Metab. 2013;305: E496–E506. doi:10.1152/ajpendo.00128.2013
- 382 15. Richter EA, Hargreaves M. Exercise, GLUT4, and Skeletal Muscle Glucose Uptake.
383 Physiol Rev. 2013;93: 993–1017. doi:10.1152/physrev.00038.2012
- 384 16. Bojsen-Møller J, Kalliokoski KK, Seppänen M, Kjaer M, Magnusson SP. Low-intensity
385 tensile loading increases intratendinous glucose uptake in the Achilles tendon
386 [Internet]. Journal of Applied Physiology. 2006. pp. 196–201.
387 doi:10.1152/jappphysiol.00004.2006
- 388 17. Kaji A, Sasagawa S, Kubo T, Kanehisa H. Transient Effect of Core Stability Exercises on
389 Postural Sway During Quiet Standing. J Strength Cond Res. 2010;24: 382–388.
390 doi:10.1519/JSC.0b013e3181c06bdd
- 391 18. Granacher U, Lacroix A, Muehlbauer T, Roettger K, Gollhofer A. Effects of Core
392 Instability Strength Training on Trunk Muscle Strength, Spinal Mobility, Dynamic
393 Balance and Functional Mobility in Older Adults. Gerontology. 2013;59: 105–113.
394 doi:10.1159/000343152
- 395 19. Imai A, Kaneoka K, Okubo Y, Shiraki H. Comparison of the immediate effect of different
396 types of trunk exercise on the star excursion balance test in male adolescent. The
397 International Journal of Sports Physical Therapy. 2014. pp. 428–435.
- 398 20. Willson JD, Dougherty CP, Ireland ML, Davis IM. Core Stability and Its Relationship to
399 Lower Extremity Function and Injury. J Am Acad Orthop Surg. 2005;13: 316–325.
400 doi:10.5435/00124635-200509000-00005
- 401 21. Day JT, Lichtwark GA, Cresswell AG. Tibialis anterior muscle fascicle dynamics
402 adequately represent postural sway during standing balance. J Appl Physiol. 2013;115:
403 1742–1750. doi:10.1152/jappphysiol.00517.2013
- 404 22. Earl JE, Hertel J. Lower-Extremity Muscle Activation during the Star Excursion Balance

- 405 Tests. *J Sport Rehabil.* 2001;10: 93–104. doi:10.1123/jsr.10.2.93
- 406 23. Croisier J-L, Ganteaume S, Binet J, Genty M, Ferret J-M. Strength Imbalances and
407 Prevention of Hamstring Injury in Professional Soccer Players. *Am J Sports Med.*
408 2008;36: 1469–1475. doi:10.1177/0363546508316764
- 409 24. Yeung SS, Suen AMY, Yeung EW. A prospective cohort study of hamstring injuries in
410 competitive sprinters: preseason muscle imbalance as a possible risk factor. *Br J Sports*
411 *Med.* 2009;43: 589–594. doi:10.1136/bjism.2008.056283
- 412 25. Bourne MN, Opar DA, Williams MD, Shield AJ. Eccentric Knee Flexor Strength and Risk
413 of Hamstring Injuries in Rugby Union. *Am J Sports Med.* 2015;43: 2663–2670.
414 doi:10.1177/0363546515599633
- 415



- | | |
|------------------|-------------------|
| 1. Anterior | 5. Posterior |
| 2. Anteromedial | 6. Posterolateral |
| 3. Medial | 7. Lateral |
| 4. Posteromedial | 8. Anterolateral |

Figure 1

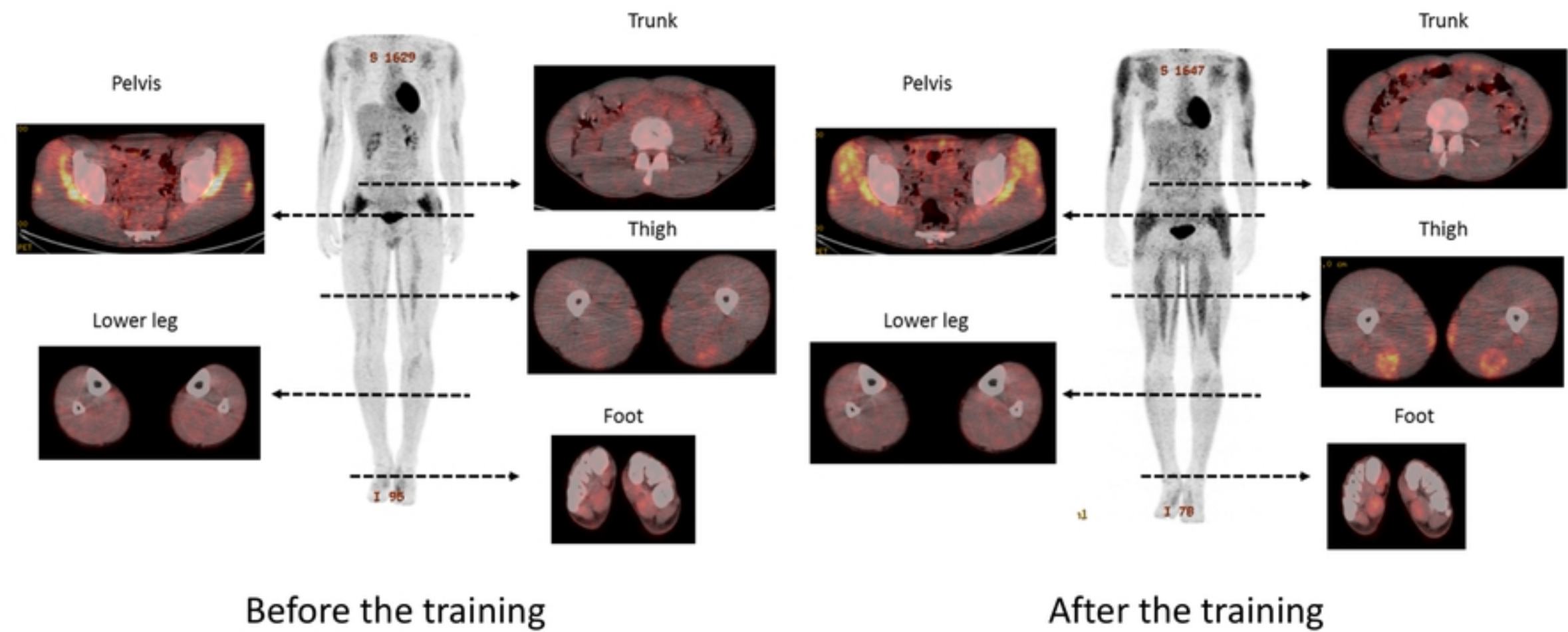


Figure 2