Race predictors and hemodynamic alteration after an ultra-trail marathon race

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Introduction

Ultra-trail running races have increased in popularity over the last few years.¹² This kind of race has several unique characters including a prolonged race period, specifically a distance of at least 50 km, and a high elevation route with rough terrain. Many recent studies have documented cardiac damage and pulmonary function deterioration after prolonged exercise, even in healthy runners.³⁴ Most of these studies focused mainly on flat-terrain running, in particular marathons. Standard long distance running races...
usually use easily accessible courses which are completely different from the more challenging variable terrain involved in trail running. This kind of sport requires extreme endurance despite the perceived low peak performance, which could make a significant difference when it is compared to high-strenuous, shorter duration competitions. With a recent increase in its popularity, many runners try to participate in this extreme sport, which results in a large heterogeneity of athletes competing, from elite runners to recreational individuals.1 Concerns regarding the suitability of the candidates make it difficult for the organizers to manage the race and put in place adequate safety regulations for the whole gamut of competitors. It is a standard rule for the trial course to have many checkpoints and food services along the course; however, the distance between each checkpoint is usually more than an hour. Organizers always brief the runners about the basic safety needs specific to the course, but the possibility of injuries throughout the race is an unpredictable factor. The main concern for the organizers is the safety of runners because the routes often run deep into forested area and access is difficult if any medical problems occur.

The noninvasive ultrasonic flow measurement (USCOM; Sydney, Australia) is an advanced hemodynamic assessment method with reliable accuracy when compared to other standard methods and has been validated by many previous publications.2-10 This is a noninvasive assessment which will not impair the performance of the runners before and after the race. This study was conducted to identify race predictors which could predict the race results and classify the runners allowing better safety management during the race and also to evaluate the physiologic effects of prolonged endurance physical exercise with a noninvasive flow assessment method.

Our hypothesis was that there is an association between gender, body mass index, training data, and heart rate and race performance, and we expected the hemodynamic change to be in accordance with hypovolemic change such as tachycardia, lower blood pressure, and higher stroke volume variation (SVV) after finishing the race.

Methods
This study was approved by the institutional review board and ethics committee of the Faculty of Medicine, Chiang Mai University, Chiang Mai, Thailand. The prospective observational study was conducted during the Pong Yaeng trail race on November 12, 2016. All runners were voluntarily invited to enroll onto this study without any exclusions.

The race
The 66 km-Pong Yaeng trail race is one of the most popular trail races in Thailand and is approved by the International Trail Running Association. The 66k-route takes the runners through a mixture of deciduous and pine forests along the old paths connecting the villages of the Pong Yaeng subdistrict, gaining an elevation of more than 3,400 m. The time limit is 17 hours.

Questionnaire and measurement
Informed written consent was obtained from each participant. All subjects answered the research questionnaire, including details such as the basic data (age, gender, bodyweight, height, medical history), running experience, and running records (fastest 10, 21 and 42 km records). Predicted maximal oxygen consumption values were obtained from running smartwatches which provide this function. The prerace assessment was carried out during the registration process 1 day before the race. All participants had hemodynamic parameters assessed, including blood pressure (mmHg) using a standard semi-automatic oscillometric blood pressure measurement device and oxygen saturation (SaO2; %) using a noninvasive fingertip pulse oximeter. Other hemodynamic parameters including heart rate (beats/min), stroke volume (SV; mL), SVV (%), cardiac index (CI; L/min/m2), and systemic vascular resistance (SVR; dynes/sec/cm5) were obtained using an USCOM device. The probe of the machine was applied at the suprasternal notch while the participant was in a sitting position until an adequate flow pattern was achieved. The operators were well trained physicians and familiar with this assessment. The ultrasonic flow measurement takes less than 5 minutes to complete the assessment.

All runners competed over the same course and in the same weather conditions. Water and food services were supplied adequately along the course. The time limit for completion of the course was 17 hours. The finishers were reassessed using the same methods after crossing the finish line and having a specific resting period, but the postrace assessments were done within 15 minutes of finishing the race. The same methods were used as in the prerace assessment. Finishing times were individually recorded by the race organizer. The runners who did not finish were not assessed after the race. The runners were divided into two groups: finishers (runners who crossed the finish line) and nonfinishers (runners who did not cross the finish line).

The noninvasive flow measurement (USCOM) was chosen to measure the hemodynamic parameters in this study because...
of its simplicity, completely noninvasive technique, and its ability to record a wide variety of hemodynamic parameters. Previous studies have validated the accuracy of this method and have shown that it is highly accurate against other standard hemodynamic assessment techniques; this technique has also been applied in many clinical applications with critical patients. This noninvasive method is the most reasonable test for runners in both a pre- and post-race assessment setting.

**Statistical analysis**

Statistical analysis was performed using STATA version 14.0 (StataCorp, College Station, TX, USA). Categorical variables are presented as numbers and percentages, and comparison between groups was made using Fisher’s exact test. Continuous variables are presented as median and interquartile ranges (IQR) or mean and standard deviation (SD), and comparison between groups was made using a Student’s t-test or a Wilcoxon rank-sum test. Race predictors were determined by univariable and multivariable logistic regression analysis. Statistical significance was established at \( p < 0.05 \).

**Results**

289 runners were registered for this race, but 61 runners did not present at the start line, leaving 228 runners who actually competed in the race. The average age was 37 years (range 24–56 years). 163 out of the 228 (71.5%) runners were male. 130 out of the 228 (57.0%) participants answered the questionnaire before the race. 126 out of 228 (55.3%) subjects completed the prerace hemodynamic assessment, and all of this group also completed the questionnaire. 189 out of 228 (82.9%) runners crossed the finish line. The average finishing time was 920 (449–1,020) minutes. 36 out of 189 (19%) finishers completed the post-race hemodynamic assessment. A summary showing the numeric distribution of the runners is described as Figure 1.

![Flow chart demonstrates the complete data of participants.](image-url)
Univariable analysis showed that the following factors were more likely to be associated with crossing the finish line: male gender, longest distance ever run, better running records at every distance (10, 21, 42 km), and a lower diastolic blood pressure (DBP) (Tables 1 and 2).

The multivariable analysis, as shown in Table 3, indicated that lower diastolic pressure was the only significant predictor for crossing the finish line. After dividing the range of diastolic pressure into three groups, according to percentile, it was clear that the runners with a lower prerace diastolic pressure have a higher chance of finishing the race (DBP 74–84 mmHg: adjusted odd ratio 3.81; 95% CI = 1.09–13.27 and DBP <74 mmHg: adjusted odd ratio 7.74; 95% CI = 1.57–38.21). However, for only 110 subjects out of 126 runners we had complete data, including both the basic data from the questionnaire and prerace hemodynamic assessments. This therefore was the only data included in the

Table 1 Univariable analysis for race predictors from demographic data, running experience, and running records

| Variables                      | Total n (%) | Finishers (n=189) | Nonfinishers (n=39) | p-value |
|--------------------------------|-------------|--------------------|---------------------|---------|
| Age (years), mean ± SD         | 228 (100)   | 37.0±7.8           | 35.8±8.0            | 0.393   |
| Gender, n (%)                  |             |                    |                     | 0.022   |
| Female                         | 65 (28.5)   | 48 (25.4)          | 17 (43.6)           |         |
| Male                           | 163 (71.5)  | 141 (74.6)         | 22 (56.4)           |         |
| Body weight (kg), mean ± SD    | 130 (57.0)  | 64.9±10.6          | 67.3±13.1           | 0.302   |
| Height (cm), mean ± SD         | 130 (57.0)  | 169.2±7.3          | 168.6±9.5           | 0.789   |
| BMI (kg/m²), mean ± SD         | 130 (57.0)  | 22.6±2.7           | 23.5±3.3            | 0.112   |
| Experience (years)             | 127 (55.7)  | 2 (2–3)            | 2 (2–3)             | 0.289   |
| Marathon finisher, n (%)       | 127 (55.7)  | 9 (7.1)            | 7 (7.1)             | 0.964   |
| No                             | 9 (7.1)     | 30 (30.6)          | 9 (32.1)            |         |
| Yes                            | 118 (92.9)  | 91 (92.9)          | 27 (93.1)           |         |
| 50 km finisher, n (%)          | 126 (55.3)  | 30 (23.8)          | 45.6±15.6           | 0.026   |
| No                             | 39 (31.0)   | 9 (32.1)           | 143±22.6            |         |
| Yes                            | 87 (69.0)   | 9 (32.1)           | 19 (67.9)           |         |
| Longest distance (km), mean ± SD| 123 (53.9)  | 56.4±23.8          | 45.6±15.6           | 0.026   |
| Best 10 km record (min), mean ± SD| 126 (55.3)  | 55.8±9.9          | 62.5±9.6            | 0.002   |
| Best 21 km record (min), mean ± SD| 124 (54.4)  | 125±20.5          | 143±22.6            |         |
| Best 42 km record (min), mean ± SD| 116 (50.9)  | 292.8±46.2         | 326.8±36.5          | <0.001  |
| Finish time (min)              | 189 (100)   | 920 (798–977)      | 449–1,020           |         |

Table 3 Multivariable analysis for race predictors (n=110)

| Variables                      | OR (95% CI) | p-value |
|--------------------------------|-------------|---------|
| Male                           | 2.02 (0.57–7.10) | 0.275   |
| Longest distance               | 1.02 (0.99–1.05) | 0.237   |
| Best 10 km record              | 1.00 (0.93–1.07) | 0.967   |
| Best 21 km record              | 0.97 (0.92–1.01) | 0.148   |
| Best 42 km record              | 1.00 (0.98–1.02) | 0.839   |
| DBP >84 mmHg                   | 1.0 Reference |         |
| DBP 75–84 mmHg                 | 3.81 (1.09–13.27) | 0.036   |
| DBP <75 mmHg                   | 7.74 (1.57–38.21) | 0.012   |

Table 2 Univariable analysis for race predictors from prerace hemodynamic parameters

| Variables                      | Total n (%) | Finishers (n=189) | Nonfinishers (n=39) | p-value |
|--------------------------------|-------------|--------------------|---------------------|---------|
| VO2 max (L/min), mean ± SD     | 43 (18.9)   | 44.4±3.9           | 41.4±1.8            | 0.055   |
| SBP (mmHg), mean ± SD          | 126 (55.3)  | 135.3±14.8         | 136.7±21.2          | 0.699   |
| DBP (mmHg), mean ± SD          | 126 (55.3)  | 78.1±9.8           | 82.6±12.7           | 0.047   |
| HR (beats/min), mean ± SD      | 126 (55.3)  | 74.9±11.9          | 78.9±14.6           | 0.134   |
| SV (mL), mean ± SD             | 125 (54.8)  | 40.2±13.4          | 35.9±9.8            | 0.120   |
| SV (%) (mL/m²), mean ± SD      | 125 (54.8)  | 50.9±35.4          | 41.0±29.9           | 0.180   |
| CI (L/min/m²), median (IQR)    | 125 (54.8)  | 2.6 (2.1–3.5)      | 2.7 (2.2–3.1)       | 0.917   |
| SVR (dyne/sec cm⁻¹), mean ± SD | 125 (54.8)  | 3,167±1,159.2      | 3,385±1,365.4       | 0.402   |
| SaO2 (%) (mL/m²), mean ± SD    | 125 (54.8)  | 97.1±1.3           | 96.7±1.2            | 0.133   |

Abbreviations: BMI, body mass index; IQR, interquartile range; SD, standard deviation.

Abbreviations: OR, odds ratio; CI, confidence interval; DBP, diastolic blood pressure; VO2 max, maximal oxygen consumption; SBP, systolic blood pressure; DBP, diastolic blood pressure; HR, heart rate; SV, stroke volume; SVV, stroke volume variation; CI, cardiac index; SVR, systemic vascular resistance; SaO2, oxygen saturation.
performance of the multivariable analysis. We did not use the multiple imputation method to correct the data, despite the data set being incomplete. This was because there was so much missing data that an error in the analysis would have been inevitable.

Out of the 33 subjects who underwent the postrace hemodynamic assessment, systolic blood pressure, heart rate, and SVV were significantly increased in the postrace status compared with prerace status, whereas SV and SaO₂ were decreased. The remaining hemodynamic parameters were not significantly changed (Table 4).

**Discussion**

Ultra-distance trail running has several unique characters that raise numerous special concerns for the race organizer. Although the International Trail Running Association was established to control and regulate every official trail running race around the world, the health policy of this association is mainly focused on individual prerace athletes’ health status and antidoping objectives. From a review of relevant literature, no study has been conducted to determine race predictors concerning trail racing. The rationale behind the study is that if the organizers could predict the race results from the wide varieties of runners, they would be better able to arrange more effective safety management for the runners who are at greater risk of withdrawal from the race.

From the results of the univariable analysis, the only significant predictor from the basic demographic data is male gender. 86.5% (141 out of 163) of male runners finished the race, while only 73.8% (48 out of 65) of female runners were finishers. The age, weight, and body mass index could not be used to predict the race results. In this study, the questionnaire regarding running experience included the number of years the participant had been running and the race finishing history of a standard marathon race and a 50K trail race. None of the experience data achieved a level of statistical significance. The distance running records, however, showed a level of statistical significance in every field including the longest distance ever run and the best records at every distance (10 km, 21 km, and 42 km). The finishers had a record of longer runs than the nonfinishers by a distance of nearly 11 km (56.4±23.8 km in the finisher group and 45.6±15.6 km in nonfinisher group; p=0.026) and also faster times at every distance. These parameters could probably be explained by the fact that the finishers had undergone better training and had higher fitness levels than the nonfinishers. These findings was confirmed in the study by Knechtle et al,¹¹ which investigated the race performance in recreational ultra-runners in a 24-hour run. They concluded that the personal best marathon time and the longest training run, not anthropometry such as skinfold thickness or body mass index, predicted the race performance. Considering these findings, we might assume that training factors are more important than body geometry, especially in ultra-distance runners.

The prerace hemodynamic parameters, as shown in Table 2, showed DBP as the only race finish predictor from the result of the multivariable analysis. The finisher group had a lower DBP than the nonfinisher group. Blood pressure in athletes has been reviewed intensively. A meta-analytical study carried out by Berge et al¹² in 2014 reviewed 16 studies that included nonathletes as controls and showed, contradictorily, that blood pressure was lower in athletes than in controls in 9 studies and higher in athletes in 7 studies. The conclusion was that there was no significant difference in blood pressure between athletes and controls. However, most of studies in this meta-analysis were conducted among elite athletes. When comparing this study to ours, in which the participants have such a mix of training levels, this finding is hard to rationalize due to the wide variation of fitness among the athletes. Several studies in this meta-analysis reported that blood pressure in athletes who had trained >10 hours/wk was higher than in those who had trained less. On the other hand, the American Heart Association lifestyle management guidelines in 2013

| Parameters                          | N    | Prerace          | Postrace         | p-value |
|------------------------------------|------|-----------------|-----------------|---------|
| SBP (mmHg), mean ± SD              | 33   | 135.9±14.8      | 119.7±11.3      | <0.001  |
| DBP (mmHg), mean ± SD              | 33   | 76.6±10.1       | 73.1±6.8        | 0.066   |
| HR (beats/min), mean ± SD          | 33   | 72.6±10.7       | 96.4±10.4       | <0.001  |
| SV (mL), mean ± SD                 | 33   | 43.2±13.6       | 29.6±10.4       | <0.001  |
| SVV (%), median (IQR)              | 33   | 36 (25–58)      | 53 (33–78)      | <0.001  |
| CI (L/min/m²), mean ± SD           | 33   | 3.0±1.0         | 2.8±0.9         | 0.583   |
| SVR (dynes/sec cm⁻³), mean ± SD    | 33   | 3,044±1,052.8   | 2,888±871.8     | 0.431   |
| SaO₂ (%), mean ± SD                | 31   | 97.4±1.0        | 96.4±1.0        | <0.001  |

**Abbreviations:** SBP, systolic blood pressure; DBP, diastolic blood pressure; HR, heart rate; SV, stroke volume; SVV, stroke volume variation; CI, cardiac index; SVR, systemic vascular resistance; SaO₂, oxygen saturation.
still advised adults to engage in aerobic physical activity to lower blood pressure, advice being: 3–4 sessions a week, lasting on average 40 minutes per session, and involving moderate-to-vigorous intensity physical activity.13 From our point of view, the lower DBP in our data could be as a result of the exercise effects on amateur athletes which is different from the elite athlete group according to the meta-analysis data. According to the multivariable analysis, only lower DBP is a statistical significant race predictor. However, only 110 subjects in our study had completed all sections of the prerace data for the performance of this analysis.

The effects of ultra-endurance physical exercise on hemodynamic and pulmonary function have been described in many previous studies.5,14–16 From the hemodynamic results in our study, systolic blood pressure, heart rate, and SVV were significantly increased, while stroke volume was significantly reduced but SVR and CI did not change. These findings could be interpreted as the results of hypovolemia, which has a physiologic response by lowering systolic pressure, initiating tachycardia, and lowering SV. However, these standard parameters are poor markers of hypovolemia.17 SVV is one of the dynamic markers which has been shown as being superior to static markers in predicting fluid responsiveness.18 In this study, the SVV was statistically significantly higher after the race. This may confirm the effects of hypovolemia that occurred in the runners even though the athletes were allowed to hydrate at will during the race. However, there are numerous limitations of SVV in clinical usage such as the need for mechanical ventilation in order to achieve complete controlled ventilation without spontaneous breathing and the control of tidal volumes (>8 mL/kg) without dysrhythmia. The CI did not change significantly; this is a result of the normal physiologic compensation of the body by driving the heart rate to compensate for the lower SV.

From our data, the runners had a lower SaO₂ after finishing the race. The worsening pulmonary function after a prolonged endurance race has been described by many studies. Seedhouse et al16 has studied 8 triathletes competing in the 3-day World Ultraman Championships, and the results from the study showed that a reduction in peak expiratory flow, forced vital capacity, and forced expiratory volume in 1 second from baseline at the end of each race day. Rogers et al13 also found that lower SaO₂, reduction of forced vital capacity (up to 22%), and forced expiratory volume in 1 second decrease (up to 25%) after finishing an Ultra-endurance multisport competition. The explanations for these findings were elusive. One of the possible causes is exercise-induced bronchospasm. The cause of this phenomenon would be from a cooled and dried airway caused by hyperventilation. It can occur in healthy individuals with no underlying pulmonary disorder and has a greater prevalence in elite athletes and those undergoing continuous high-intensity exercise.19 The most likely etiology of exercise-induced pulmonary dysfunction is exercise-induced noncardiogenic pulmonary edema from hyponatremia which will be spontaneously resolved when the initiated electrolyte disorder is corrected.19 This finding was also found by Zavorsky et al in their study focusing on marathon-induced lung edema, and they found that mild pulmonary edema occurs in at least 17% of subjects and it is not possible to predict the occurrence or severity of the edema. However, this physiologic change is usually mild in severity and does not affect the runner’s performance.

This study is one of the very few studies that is focused on the ultra-trail running race using both historical data and basic demographic data along with the sophisticated biological data collected by noninvasive methodology. However, there are many limitations to this study. 130 out of 228 (57.0%) volunteers answered the questionnaire; 120 out of 228 (52.6%) participants had the hemodynamic flow assessment 1 day before the race; and only 39 out of 189 (20.6%) finishers had the postrace assessment after crossing the finish line. This incomplete data collection may interfere with the statistical results of this study, and so the results may not be extrapolated to represent the whole of this population of runners. We cannot assess the hemodynamic change in the nonfinisher group due to the nature of the experimental design which did not include the monitoring of runners who withdrew from the race at any checkpoint along the course. We did believe that the USCOM would be the most appropriate noninvasive hemodynamic assessment tool for this situation, but there were several limitations to this device as its main clinical application is for fully sedated, critically ill patients under controlled ventilation condition, not for healthy spontaneously ventilated individuals.

The clinical implications of this study could be used by the race organizers who could use the significant univariable factor from the gender and the training factors to classify the runners into groups by predicted race finishing time. This would enable the targeting of the safety arrangements and the overall management of the trail race. The diastolic pressure is the sole significant factor from the multivariable analysis, but classification of the runners using this factor would be difficult with a single mean. However, it will be a field of great interest for the future research conducted specifically around this factor. Future research would be conducted with a slightly different study design and with an incentivized runner recruitment to avoid the large number of dropout participants. For improved physiologic assessment and
monitoring, invasive assessments such as blood chemistry tests should be used and a specific volunteer control group must be assigned.

The race predictors from the univariable analysis included male gender, longest distance ever run, faster running records over specific distances (10, 21, 42 km), and lower diastolic pressure. However, from the multivariable analysis of the 40 runners included, the sole significant race predictor is lower diastolic pressure. From the hemodynamic data of 39 out of 189 (20.6%) finishers, runners seem to have a hypovolemic physiologic response including lower systolic blood pressure, faster heart rate, lower SV, and higher SVV with exercise-induced pulmonary injury which presented as lower oxygen saturation. The data from this study would help extreme sport organizers to prepare more safely for their races so that more appropriate medical service regulation for the competitors could be provided.

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Author contributions
Study concept and design (N Taksaudom, N Tongsiri); obtaining funding; acquisition of the data (N Taksaudom, N Tongsiri, AP, CL, AC); analysis of the data (N Taksaudom, AT); drafting of the manuscript (N Taksaudom); critical revision of the manuscript (N Taksaudom, AT); approval of final manuscript (N Taksaudom, N Tongsiri, AT). All authors contributed toward data analysis, drafting and critically revising the paper and agree to be accountable for all aspects of the work.

Disclosure
The USCOM device was loaned free of charge by the Saint Medical Group Co. Ltd. Natee Tongsiri is the chief organizer of the Pong Yaeng 66 km-ultra-trail race. The authors report no other conflicts of interest in this work.

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References
1. A Special Report on Trail Running; 2010. Available from: http://www.outdoorfoundation.org/pdf/research/trailrunning.pdf.
2. Outdoor Recreation Participation; 2014. Available from: http://www.outdoorfoundation.org/pdf/ResearchParticipation2014.pdf.
3. Middleton N, George K, Whyte G, Gaze D, Collinson P, Shave R. Cardiac troponin T release is stimulated by endurance exercise in healthy humans. J Am Coll Cardiol. 2008;52(22):1813–1814.
4. Shave RE, Whyte GP, George K, Gaze DC, Collinson PO. Prolonged exercise should be considered alongside typical symptoms of acute myocardial infarction when evaluating increases in cardiac troponin T. Heart. 2005;91(9):1219–1220.
5. Rogers IR, Speedy D, Hillman D, Noftsinger B, Inglis S. Respiratory function changes in a wilderness multisport endurance competition: a prospective case study. Wilderness Environ Med. 2002;13(2):135–139.
6. Zavorsky GS, Milne EN, Lavorini F, et al. Small changes in lung function in runners with marathon-induced interstitial lung edema. Physiol Rep. 2014;2(6):e12056.
7. Beltramo F, Menteer J, Razavi A, et al. Validation of an ultrasound cardiac output monitor as a bedside tool for pediatric patients. Pediatr Cardiol. 2016;37(1):177–183.
8. Chong SW, Peyton PJ. A meta-analysis of the accuracy and precision of the ultrasonic cardiac output monitor (USCOM). Anaesthesia. 2012;67(11):1266–1271.
9. Phillips RA, Smith BE, Madigan VM, et al. Assessment of the clinical utility of an ultrasonic monitor of cardiac output (the USCOM) and agreement with thermodilution measurement. Crit Care Resusc. 2010;12(3):209–213.
10. Thom O, Taylor DM, Wolfe RE, et al. Comparison of a supra-ternal cardiac output monitor (USCOM) with the pulmonary artery catheter. Br J Anaesth. 2009;103(6):800–804.
11. Knechtle B, Knechtpe Rosemann T, Lepers R. Personal best marathon time and longest training run, not anthropometry, predict performance in recreational 24-hour ultrarunners. J Strength Cond Res. 2011;25(8):2212–2218.
12. Berge HM, Isern CB, Berge E. Blood pressure and hypertension in athletes: a systematic review. Br J Sports Med. 2015;49(11):716–723.
13. Ecker RH, Jakicic JM, Ard JD, et al. 2013 AHA/ACC guideline on lifestyle-style management to reduce cardiovascular risk: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines. Circulation. 2014;129(25 Suppl2):S76–S99.
14. Jouffroy R, Caille V, Perrot S, Vicelillard-Baron A, Dubourg O, Mansencal N. Changes of cardiac function during ultradistance trail running. Am J Cardiol. 2015;116(8):1284–1289.
15. Klinkenberg LJ, Luyten P, van der Linden N, et al. Cardiac Troponin T and I release after a 30-km run. Am J Cardiol. 2016;118(2):281–287.
16. Seedhouse EL, Walsh ML, Blaber AP. Heart rate, mean arterial blood pressure, and pulmonary function changes associated with an ultradistance triathlon. Wilderness Environ Med. 2006;17(4):240–245.
17. Michard F, Teboul JL. Predicting fluid responsiveness in ICU patients: a critical analysis of the evidence. Chest. 2002;121(6):2000–2008.
18. Maitland K, Levin M, English M, et al. Severe P. falciparum malaria in Kenyan children: evidence for hypovolaemia. QJM. 2003;96(6):427–434.
19. Sanchez LD, Corwell B, Berkoff D. Medical problems of marathon runners. Am J Emerg Med. 2006;24(5):608–615.