



**Faculdade de Ciências do Desporto e de Educação Física
Universidade do Porto**

**Crioterapia: efeitos na homeostasia muscular
após o exercício**

Dissertação apresentada às provas de doutoramento no ramo de Ciência do Desporto, nos termos do Decreto-Lei nº 216/92 13 de Outubro.

Manoel da Cunha Costa

Março de 2002



ÍNDICE GERAL

1. Introdução	1
1.1 Objetivos	3
2. Crioterapia, conceito e modalidades.	4
2.1 Meios comuns de aplicação da crioterapia	4
2.1.1 Bolsa de gelo	4
2.1.2 Sistemas de circulação de água gelada	5
2.1.3 Massagem com gelo	5
2.1.4 Imersão em água gelada e turbilhões	5
2.2 Entropia e Termodinâmica	6
2.2.1 Primeira Lei da Termodinâmica	8
2.2.2 Segunda Lei da termodinâmica	9
2.3 Transferência de energia no corpo humano	10
2.3.1 Condução	10
2.3.2 Convecção	10
2.3.3 Radiação	10
2.3.4 Conversão	11
2.4 Resfriamento e reaquecimento dos tecidos:	11
2.5 Material e métodos	11
2.5.1 Sistema de crioterapia	11
2.5.2 Controle da temperatura dos animais	14
2.5.3 Modelo animal:	15
2.5.4 Procedimento experimental	16
2.5.5 Tratamento dos dados e análise estatística	16
2.5.6 Resultados e discussão	16
3. Miopatia do exercício	27
3.1 Definição	27
3.2 Etiologia/ Fisiopatologia/Miopatia do exercício	27

3.2.1 Etapa inicial:	28
3.2.2 Etapa autogénica	30
3.2.3 Etapa fagocitária	33
3.3 Metodologia	36
3.3.1 Modelo animal	36
3.3.2 Procedimento experimental	36
3.3.3 Crioterapia	37
3.3.4 Exercício	37
3.3.6 Retirada dos músculos	38
3.3.7 Análises bioquímicas	38
3.3.8 Tratamento dos dados e análise estatística	39
3.4 Resultados e discussão	40
3.4.1 Resultado proteínas totais	40
3.4.2 Resultados TBARS	41
3.4.3 Resultados β-Glucuronidase	42
3.4.4 Resultados glutationa	45
3.4.5 Resultados mieloperoxidase	48
3.4.6 Resultados N-Acetyl- Glucosaminidase (NAG)	49
4. Manifestações clínicas da miopatia do exercício (ME)	55
4.1 Indicadores indirectos da miopatia do exercício	55
4.1.1 Perda da funcionalidade	55
4.1.1.1 Alterações na força	55
4.1.1.2 Diminuição da amplitude dos movimentos	55
4.1.2 Dor tardia induzida pelo exercício	56
4.1.3 Alterações morfológicas (edema)	56
4.2 Metodologia	58
4.2.1 Amostra	58
4.2.2 Procedimento experimental	58
4.2.3 Crioterapia	59
4.2.4 Exercício	61

4.2.5 Indicadores indirectos	62
4.2.6 Cronograma de procedimentos	65
4.2.7 Tratamento dos dados e análise estatística	65
4.3 Resultados e discussão	66
4.3.1 Creatina quinase (CK)	66
4.3.2 Resultado escala de dor	68
4.3.3 Variações angulares articulação do tornozelo	71
4.3.4 Pico isométrico de força	72
4.3.5 Perímetros	74
5. Conclusões	76
6. Referências bibliográficas	78

1. Introdução

Todas as funções físicas do corpo implicam actividade muscular. Estas funções incluem movimentos do esqueleto, a contração do coração, a contração dos vasos sanguíneos, o peristaltismo do intestino e muitas outras (91). De um ato explosivo, como um salto, a uma ação mais refinada como a escrita, todas dependem do tecido muscular esquelético, que se constitui como um dos tecidos mais adaptáveis dos seres vivos, devido as suas propriedades intrínsecas.

O tecido muscular esquelético, como qualquer outro tecido humano, está sujeito a agressões de diversas naturezas, mecânicas, metabólicas, térmicas, etc. Destacamos neste estudo o exercício como o agente desta agressão.

Os exercícios intensos ou não habituais produzem no tecido muscular esquelético, um fenômeno designado por comumente por "miopatia do exercício" (68). Esta entidade é, por definição, considerada como um conjunto de alterações transitórias e benignas que ocorrem no tecido muscular quando submetido a níveis elevados de stress. Este acometimento pode ser de ordem morfológica e/ou funcional e o processo patológico ocorre quando um fator, de ordem interna ou externa, degrada ou danifica as células do tecido muscular, impedindo-as de executar suas funções normais. Paralelamente, este quadro é habitualmente acompanhado por um processo inflamatório transitório (citotóxico) (68).

Este tipo de agressão causada pelo exercício, seguida pela quebra de homeostasia nos músculos, tem um caráter biopositivo dentro do processo de treinamento, pois resguarda e faz valer o princípio da sobrecarga no processo de adaptabilidade, já que é necessário que haja a quebra da homeostasia do organismo, para que ele possa se organizar estrutural e funcionalmente a fim de não ser perturbado novamente por estímulos de mesma magnitude. (20, 45, 63, 68, 93).

Sendo assim, o exercício pode ser encarado como uma agressão orgânica particularmente para o tecido muscular, já que perturba, ainda que transitoriamente, a homeostasia das fibras musculares (68). Esta agressão pode

dar-se através de três mecanismos básicos: por via metabólica, mecânica e combinada (14, 68,164).

A miopatia do exercício é caracterizada clinicamente por um processo inflamatório local, dor e perda da função (força e mobilidade articular).

Tradicionalmente são utilizadas diversas estratégias para recuperação ou minimização dos efeitos da miopatia induzida pelo exercício, como por exemplo: massagem, eletroestimulação, ultra-som, infravermelho laser, antiinflamatórios, antioxidantes, crioterapia, alongamento etc. (1, 8, 34, 57, 66, 87, 88, 92, 143, ,144, 171, 174, 182, 217). Ou seja, alguns dos procedimentos utilizados para diminuir a perda da funcionalidade associada a miopatia do exercício, são também os referidos para o tratamento de lesões desportivas. Entre os vários procedimentos, destacamos a forma mais clássica de terapêuticas: Repouso (R), Gelo (I), Fixação (C) e Elevação (E), originando a sigla em língua inglesa RICE (Rest, Ice, Compression, Elevation). Estas condutas têm sido descritas como influenciadoras do edema, dor, e perda de função, desencadeados pelo processo inflamatório (57).

Durante um programa de treinamento fica inviável a aplicação desta eficiente combinação de condutas. Daí surge o desafio de encontrar alternativas, que minimizem o desconforto causado durante o processo inflamatório induzido pelo exercício, sem prejuízo na continuidade dos treinamentos.

Neste estudo foi realizado um corte na terapêutica clássica, selecionando apenas o frio (resfriamento) como forma de tratamento na recuperação da homeostasia muscular. A utilização da crioterapia justifica-se, neste caso, pelo fato de ser aquele procedimento que é mais facilmente utilizável durante um processo de treino intenso. Ou seja, pretendemos, experimentalmente, estudar o efeito do frio na evolução das alterações musculares induzidas por um exercício considerado fisiologicamente intenso e, deste modo, tentar encontrar procedimentos facilmente exequíveis de diminuição da funcionalidade orgânica após agressão muscular.

O frio isoladamente ou associado ao calor, são as terapêuticas mais comumente utilizadas, devido ao seu caráter prático em suas diversas formas de

aplicação. Basicamente a aplicação do frio produz uma redução na temperatura da pele e de tecidos mais profundos na região da aplicação (57, 110, 144). Apesar deste procedimento ser largamente utilizado, existe uma carência de estudos científicos, que demonstrem de forma objetiva os mecanismos de ação do frio no processo inflamatório, sendo encontradas apenas observações acerca dos resultados alcançados, na redução do edema, dor e recuperação da mobilidade.

Quando abordamos tecnicamente a aplicação do frio de forma clínica, surge um termo de designação, Crioterapia, consiste na associação do termo frio + terapia, ou seja, qualquer procedimento com intenção de tratamento que utilize gelo, água gelada ou qualquer outra forma de frio, como estratégia de diminuição da temperatura tecidual no local de aplicação (110).

Para concretizar os objetivos deste trabalho, subdividimos, o estudo em três capítulos que abordam: (I) o desenvolvimento de tecnologia para crioterapia, no modelo animal, com metodologia de frio por convecção, sem utilização de temperaturas moderadas (10 a 15 graus Celsius); (II) a aplicação desta metodologia em animais com a avaliação de variáveis bioquímicas e histológicas, indicadoras de lesão tecidual e, finalmente, (III) a reprodução do modelo de crioterapia em humanos, com a utilização de marcadores indiretos.

1.1 Objetivos

- Desenvolver sistema para aplicação de crioterapia moderada em animais de laboratório;
- Analisar as interferências da crioterapia moderada na homeostasia muscular, após a agressão tecidual produzida pelo exercício;
- Analisar alterações histológicas e bioquímicas, induzidas pela crioterapia moderada, exercício e a sua combinação;
- Analisar respostas indiretas da miopatia do exercício em humanos, submetidos a exercício, crioterapia moderada, exercício e sua combinação.

6. Referências bibliográficas

- 1-** ABRAMSON, D. I.:Physiologic basis for the use of physical agents in peripheral vascular disorders. **Arch. Phys. Med. Rehabil.** 1995,46:216.
- 2-** ALLEN, P.D., LOPEZ, J.R., SANCHEZ, V. et al.: EU 4093 Decreases intracellular $[Ca^{2+}]$ in skeletal muscle fibers from control and malignant hyperthermia-susceptible swine. In **Anesthesiology**. 76 (1), 1992, 132-138.
- 3 –** ALLESIO, H.M. , et. Al.: generation of reactive oxygen species after exhaustive aerobic and isometric exercise. **Med Sci Sports Exerc.** 32(9): 2000, 1576-81.
- 4-** ALMEKINDERS, L.C., ALMEKINDERS, S.V.: Immune function in exercise-induced injuries. **Exercise and Disease**, [?],1992,149-158.
- 5-** AMELINK, G.J.H., VAN DER KALLEN, C.J.H., WOKKE, J.H.J., et al.: Dantrolene sodium diminishes exercise-induced muscle damage in the rat. **Eur. Journal Pharmacology**, 179, 1990,187-192.
- 6-** AMELINK, G.J., ERICH, W.B.M., KOOT, R.W. et al.: Sex-linked variation in creatine kinase release, and its dependence on oestradiol, can be demonstrated in an in-vitro rat skeletal muscle preparation. **Acta Physiol Scand**, 138, 1990,115-124.
- 7-** APPELL, H.J., SOARES, J.M.C, DUARTE, J.A.R.: Exercise, muscle damage and fatigue. **Sports Medicine**, 13 (2), 1992,108-115.
- 8-** APPELL, H.J., SOARES, J.M.C, DUARTE, J.A.R.:Supplementation of vitamin E may attenuate skeletal muscle immobilization atrophy. **Int. J. Sports Med.** 18, 1997, 157-160.
- 9-** APPELL, H.J., SOARES, J.M.C, DUARTE, J.A.R. et al.: Administration of tourniquet : Prevention of postischemic oxidative stress can reduce muscle edema. **Arch Orthop Trauma Surg.** 255, 1996, 1-5.
- 10-** APPELL, H.J., SOARES, J.M.C, DUARTE, J.A.R. et al.: Skeletal muscle damage during tourniquet-induced ischaemia. **Eur J Appl Physiol**, 67, 1993, 342-347.
- 11-** ARMSTRONG, R.B.: Mechanisms of exercise-induced delayed onset muscular soreness : a brief review. In **Medicine and Science in Sports and Exercise**, 16 (6), 1984, 529-538.

- 12-** ARMSTRONG, R.B., LAUGHLIN, M.H., ROME, L., et al.: Metabolism of rats running up and down an incline. **Journal of Applied Physiology**, 55 (2), 1983, 518-521.
- 13-** ARMSTRONG, R.B.: Initial events in exercise-induced muscular injury. **Medicine and Science in Sports and Exercise**, 22 (4), USA : 1990, 429-435.
- 14-** ARMSTRONG, R.B., WARREN, G.L., WARREN, J.A.: Mechanisms of exercise-induced muscle fibre injury. **Sports Medicine**, 12 (3), 1991, 184-207.
- 15 -** ARMSTRONG, R.B.: Muscle damage and endurance events. **Sports Medicine**, 3, 1986, 370-381.
- 16 -** ARMSTRONG, R.B., DUAN, C., DELP, M.D., et al.: Elevations in rat soleus muscle $[Ca^{2+}]$ with passive stretch. **Journal of Applied Physiology** 74 (6), 1993, 2990-2997.
- 17 -** ARMSTRONG, L.E., MARESH, C. M., GABAREE, C. V. et al.: Thermal and circulatory responses during exercise: effects of hypohydration, dehydration, and water intake. **Journal of Applied Physiology** 82 (6), 1997, 2028-2035.
- 18-** ARMSTRONG, R. B., OGILVE, R. W., SCHWANE, J.A.: Eccentric exercise-induced injury to rat skeletal muscle. **The American Physiological Society**. 54 (1), 1983, 80-93.
- 19 –** ARSLAN, S.; et. Al.: Free radical changes in rat muscle tissue after exercise. **Rheumatol Int.** 20(3), 2001,109-112.
- 20 -** BÄR, P. R., AMELINK, G. J., JACKSON, M. J. et al.: Aspects of exercise-induced muscle damage. In **Elsevier science publishers**. 1990, 1143-1148.
- 21-** BEINER, J.M.; JOKL, P.: Muscle contusion injuries: current treatment options. **J Am Acad Orthop Surg.** 9(4): 2001,227-37.
- 22-** BELCASTRO, A.N.: Skeletal muscle calcium-activated neutral protease (calpain) with exercise. **Journal of Applied Physiology**, 74 (3), 1993, 1381-1386.
- 23 -** BENZI, G: Aerobic performance and oxygen free-radical. **J Sports Med Phys Fit.** 33(3): 1993, 205-222.
- 24 -**BHATTACHARYYA, J., THOMPSON, K.D., SAYEED, M.M.: Skeletal muscle Ca^{2+} flux and catabolic response during sepsis. **The American Physiological Society**, 265 (34), 1993, 487-493.

- 25-** BIERMAN, W.; FRIEDLANDER, M. : The penetrative effect of cold. **Arch Phys Ther.** 1940, 21:585.
- 26-** BISCHOFF, R., HEINTZ, C.: Enhancement of Skeletal Muscle Regeneration. **Developmental dynamics**, 201: 1994, 41-54.
- 27 -** BLOCK, A.B.: Thermogenesis in muscle. **Annu. Rev. Physiol.** 56, 1994, 535-577.
- 28 -** BOONE JR., J.B., LAMBERT, C.P., FLYNN, M.G., et al.: Resistance exercise effects on plasma cortisol, testosterone and creatine kinase activity in anabolic-androgenic steroid users. **Int. J. Sports Med.**, 11 (4), 1990, 293-297.
- 29 -** BOULAY, M.R., SONG, T.M.K., SERRESSE, O., et al.: Changes in plasma electrolytes and muscle substrates during short-term maximal exercise in humans. **Can. Journal of Applied Physiology**, 20 (1), 1995, 89-101.
- 30 -** BRASILEIRO FILHO, G.; PEREIRA, F.E.L. ; PITTELLA, J.E.H. ; BAMBIRRA, E.A. ; BARBOSA, A.J.A. : **Patotogia Geral** ; Guanabara Koogan, - Rio de Janeiro - RJ, 1993.
- 31 -** BOUSQUET, J; CHANEZ, P.; MERCIER, J.; PRÉFAUT, C.: Monocytes, exercise and inflammatory response . In: **Exercise Immunology review**. Shepard, R (ed.) Vol. 2:35-44, 1996.
- 32 -** BROOKS, S.V., FAULKNER, J.A.: Contraction-induced injury : recovery of skeletal muscle in young and old mice. **The American Physiological Society**, 258 (27), 1990, 436-442.
- 33 -** BROTCHIE, D., DAVIES, I., IRELAND, G. et al.: Dual-channel laser scanning microscopy for the identification and quantification of proliferating skeletal muscle satellite cells following synergist ablation. **J. Anat.**, 186, 1995, 97-102.
- 34 -** BUSCHELL A; KLENERMAN L; DAVIES H; GRIERSON I; JACSON M.J.: Ischemia reperfusion induced muscle damage. protective effect of corticosteroids and anti-oxidant in rabbits. **Acta Orthop Scand**, 1996 , 67:4, 393-398.
- 35 -** BYRD, S.K.: Alterations in the sarcoplasmatic reticulum: a possible link to exercise-induced muscle damage. **Méd Sci Sports Exerc.** 24(5) : 1992, 531-536.

- 36-** BYRNES, W.C., CLARKSON, P.M., WHITE, J.S., et al.: Delayed onset muscle soreness following repeated bouts of downhill running. **The American Physiological Society**, 59 (3), 1985, 710-715.
- 37 -** CAMUS, G., PINCEMAIL, J., LEDENT, M. et al.: Plasma levels of polymorphonuclear elastase and myeloperoxidase after uphill walking and downhill running at similar energy cost. **J. Sports Med.**, 13 (6), New York : 1992, 443-446.
- 38 -** CAMUS, G., FELEKIDIS, A., PINCEMAIL, J., et al.: Blood levels of reduced/oxidized glutathione and plasma concentration of ascorbic acid during eccentric and concentric exercises of similar energy cost. **Archives Internationales de Physiologie, de Biochimie et de Biophysique**, 102, 1994, 67-70.
- 39 -** CAMUS, G., DEBY-DUPONT, G., DEBY, C., et al.: Inflammatory response to strenuous muscular exercise in man. In **Mediators of Inflammation**, 2, 1993, 335-342.
- 40-** CANNON, J.G., FIELDING, R.A., FIATARONE, M.A., et al.: Increased interleukin 1 β in human skeletal muscle after exercise. **Am. J. Physiol.**, 257 (26), 1989, 451-455.
- 41 -** CARLSON, B.M., FAULKNER, J.A. The regeneration of skeletal muscle fibers following injury: a review. **Medicine and Science Sports and Exercise**, 15 (3), 1983, 187-198.
- 42 -** CARTER, G.T., KIKUCHI, N., ABRESCH, R.T. et al.: Effects of exhaustive concentric and eccentric exercise on murine skeletal muscle. **Arch. Phys. Med. Rehabil.**, 75, 1994, 555-559.
- 43-** CARVALHO, F.F.; REMIÃO, F. ; VALE, P. ; TIMBRELL, J. A.; BASTOS, M. L.; FERREIRA, M. A. : Glutathione and cysteine measurement in biológica samples by HPLC with a gassy carbon working detector. **Biomed. Chromat..** 8: 134-136 (1994)
- 44 -** CHANCE, B., DAIT, M.T., ZHANG, C., et al.: Recovery from exercise-induced desaturation in the quadriceps muscles of elite competitive rowers. **Am. J. Physiol.**, 262 (31), 1992, 766-775.
- 45 -** CHEN, T.C., HSIEH, S.S.: The effects of repeated maximal voluntary isokinetic eccentric exercise on recovery from muscle damage. **Research Quarterly for Exercise and Sport**, 71 (3), 2000, 260-266.

- 46** - CHEN, Y.J., GUO, Q.F.: Free radical injury in exercise-induced delayed onset muscle damage – A primary study of mechanism of delayed onset muscle damage (Part B). **Hong Kong International Sports Medicine Conference**, [?], 1992, 135-140.
- 47** - CHIN, E.R., GREEN, H.J., GRANGE, F., et al.: Technical considerations for assessing alteration in skeletal muscle sarcoplasmic reticulum Ca^{++} - sequestration function in vitro. **Molecular and cellular biochemistry**, 139, 1994, 41-52.
- 48** - CLARKE, R.S.J.; HELLON, R.F.; LIND, A.R.: Vascular reactions of the human forearm to cold. **Clin Sci.** 195, 17:165, 8.
- 49** - CLARKSON, P.M. :Eccentric exercise and muscle damage. **Sports Med**, 18 (4), 1997, 314-317.
- 50** - CLARKSON, P.M., BYRNES, W.C., McCORMICK, K.M., et al.: Muscle soreness and serum creatine kinase activity following isometric, eccentric and concentric exercise. **J. Sports Med.**, 7 (3), New York : 1986, 152-155.
- 51** - CLARKSON, P.M.: Exercise-induced muscle damage-animal and human models. In **Medicine and Science in Sports and Exercise**, 24 (5), 1991, 510-511.
- 52** - COBBOLD , A. F. ; LEWIS, O. J.: Blood flow to the knee joint of the dog: Effect of heating, cooling and adrenaline. **J Physiol** 132:279, 1956.
- 53** - COSTILL, D.L., PASCOE, D.D., FINK, W.J., et al.: Impaired muscle glycogen resynthesis after eccentric exercise. In **J. Appl., Physiol.**, 69 (1), 1990, 46-50.
- 54** – CRENSH AW, A.G., THORNELL, L.-E., FRIDÉN, J.: Intramuscular pressure, torque and swelling for the exercise-induced sore vastus lateralis muscle. **Acta Physiol. Scand.**, 152, 1994, 265-277.
- 55** - RENSHAW, A.G., KARLSSON, S., FRIDÉN, J.: Knee extension torque and intramuscular pressure of the vastus lateralis muscle during eccentric and concentric activities. **Eur. Journal of Applied Physiology**, 70, 1995, 13-19.
- 56** - DAS, D.K. **Pathophysiology of reperfusion injury**. Boca Raton, CRC Press, 1993.
- 57** - DENEGAR, C.R.: **Therapeutic modalities for athletic injuries**: Athletic training education series. USA, Human kinetics, 2000.

- 58** - DOLENC, I., CRNE-FINDERLE, N., ERZEN, I. et al.: Satellite cells in slow and fast rat muscle differ in respect to acetylcholinesterase regulation mechanisms they convey to their descendant myofibers during regeneration. **Journal of Neuroscience Research**, 37, 1994, 236-246.
- 59** - DONNELLY, A. E., CLARKSON, P. M., MAUGHAN, R. J.: Exercise-induced muscle damage: effects of light exercise on damaged muscle. **Eur. Journal of Applied Physiology**, 64, 1992, 350-353.
- 60** - DONAVAN, C. M., FAULKNER, J.A.: Plasticity of skeletal muscle regenerating fibers adapted more rapidly than surviving fibers. **The American Physiological Society**, 62 (6), 1987, 2507-2511.
- 61** - DOOLEY, P.C., BACH, T.M., LUFF, A.R.: Effect of vertical jumping on the medial gastrocnemius and soleus muscles of rats. **Journal of Applied Physiology**, 69 (6), 1990, 2004-2011.
- 62** - DOYLE, J.A., SHERMAN, W.M., STRAUSS, R.L.: Effects of eccentric and concentric exercise on muscle glycogen replenishment. **Journal of Applied Physiology**, 74 (4), 1993, 1848-1855.
- 63** - DRESSENDORFER, R.H., WADE, C.E., CLAYBAUGH, J., et al.: Effects of 7 successive days of unaccustomed prolonged exercise on aerobic performance and tissue damage in fitness joggers. **Int. J. Sports Med.**, 12, 1991, 55-61.
- 64** - DUAN, C., DELP, M. D., HAYES, D. A. et al.: Rat skeletal muscle mitochondrial $[Ca^{2+}]$ and injury from downhill walking. **Journal of Applied Physiology**, 68 (3), 1990, 1241-1251.
- 65** - DUARTE, J. A., SOARES, J.M.C., APPELL, H.J. et al.: Do invading leucocytes contribute to the decrease in glutathione concentration indicating oxidative stress in exercised muscle, or are they important for its recovery ? In **Eur. Journal of Applied Physiology**, 68, 1994, 48-53
- 66** - DUARTE, J. A., SOARES, J.M.C., APPELL, H.J. et al.: Nifedipine diminishes exercise-induced muscle damage in mouse. **Int. J. Sports Med.**, 13 (3), 1992, 274-277.
- 67** - DUARTE, J. A., SOARES, J.M.C., APPELL, H.J. et al. Endothelium-derived oxidative stress may contribute to exercise induced muscle damage. In **Int. J. Sports Med.**, 14 (8), 1993, 440-443.
- 68** - DUARTE, J. A.: Lesões celulares do músculo esquelético induzidas pelo exercício físico. Dissertação apresentada às provas de doutoramento no ramo de

Ciências do Desporto, especialidade de Biologia do Desporto. Faculdade de Ciências do Desporto e de Educação Física, **Universidade do Porto**, 1993.

69 - EBBELING, C.; CLARKSON, P.M. : Muscle adaptation prior to recovery following eccentric exercise . **Eur J Appl Physio**. 60: 1990, 26-31,

70 - ENGEL, A.G., ARAHATA, K., BIESECKER, G.: Mechanisms of muscle fiber destruction. **Neuromuscular Diseases**, 1984, 137-141.

71 - EPSTEIN, H.F. Skeletal muscle structure and function. **Principles of molecular medicine**, Totowa, NJ : 1998, 851-857.

72 - EVANS, W.; CANNON, J.: The metabolic effects of exercise-induced muscle damage. **Exerc Sports Rev**. 19:99-125, 1991

73 - EVORA, P.R.B., PEARSON, P.J., SECCOMBE, J.F., et al.: Lesão de isquemia-reperfusão. Aspectos fisiológicos e a Importância da função endotelial. In **Arquivos Brasileiros de Cardiologia**, 66 (4), 1996, 1-14.

74 - FAGAN, J.M., WAJNBERG, E.F., CULBERT, L., et al.: ATP depletion stimulates calcium-dependent protein breakdown in chick skeletal muscle. **Am. J. Physiol.** 262 (25), 1992, 637-643.

75 - FARBER, J.L. :The role of calcium in lethal cell injury.\b>American Chemical Society, 1990, 503-508.

76 - FARRY, P.J., PRENTICE, N.G.: Ice treatment of injured ligaments: An experimental model. **NZ Med J**. 9:12, 1980.

77 - FAVERO, T.G., ZABLE, A.C., BOWMAN, M.B.: Metabolic end products inhibit sarcoplasmatic reticulum Ca^{2+} release and [^3H]ryanodine binding. **Journal of Applied Physiology** 78 (5), 1995, 1665-1672.

78 - FEHR, H.G., LÖTZERICH, H., MICHNA, H.: Human macrophage function and physical exercise: phagocytic and histochemical studies. **Eur. Journal of Applied Physiology** 58, 1989, 613-617.

79 - FERRY, A., AMIRIDIS, I., RIEU, M.: Glycogen depletion and resynthesis in the rat after downhill running. **Eur. Journal of Applied Physiology**, 64, 1992, 32-35.

80 – FOLKOW, B. , et. Al.: Studies on the reaction of the cutaneous vessels to coul exposure. **Acta Physiol Scan.** 58:342, 1963.

- 81** - FREIBURG, A., TROMBITAS, K., HELL, W., et al. :Series of exon-skipping events in the elastic spring region of titin as the structural basis for myofibrillar elastic diversity. **Circ. Res.**, 86, 2000, 1114-1121.
- 82** - FRIDÉN, J.:Muscle soreness after exercise : Implications of morphological changes. In **Sports Med**, 5, New York : 1984, 57-66.
- 83** - FRIDÉN, J., SJÖSTRÖM, M., EKBLOM, B.: Myofibrillar damage following intense eccentric exercise in man. In **Int. J. Sports Med.**, 4 (3), 1983, 170-176.
- 84** - GIBALA, M.J., McDougall, J.D., TARNOPOLSKY, M.A., et al.: Changes in human skeletal muscle ultrastructure and force production after acute resistance exercise. **Journal of Applied Physiology**, 78 (2), 1995, 702-708.
- 85** - GIDDINGS, C.J., NEAVES, W.B., GONYEA, W.J.: Muscle fiber necrosis and regeneration induced by prolonged weight-lifting exercise in the cat. **The anatomical record**, 211, 1985, 33-141.
- 86** - GIULIANI, A.; CESTARO, B.: Exercise, free radical generation and vitamins. **Eur. Cancerr Prev**, 1997 Mar, 6 Suppl 1:, S55-67.
- 87** -GLASGOW, P.D., HILL, I.D., McKEVITT, A.-M., et al.: Low intensity monochromatic infrared therapy : A preliminary study of the effects of a novel treatment unit upon experimental muscle soreness. **Lasers in Surgery and Medicine**, 28 (1), 2001, 33-39.
- 88** - GLEESON, M., BLANNIN, A.K., WALSH, N.P.: Overtrainning, imunosupression, exercise-induced muscle damage and anti-inflammatory drugs. In **The clinical pharmacology of sport and exercise**, 1997, 47-57.
- 89** - GOSSEN, E.R., ALLINGHAM, K., SALE, D.G.: Effect of temperature on post-tetanic potentiation in human dorsiflexor muscles. **Can. J. Physiol. Pharmacol.**, 79, 2001, 49-58.
- 90** - GULICK, D., KIMURA, I.F. :Delayed onset muscle soreness : What is it and How do we treat it ? **Journal of Applied Physiology Journal of sport rehabilitation**, 5, 1996, 234-243.
- 91** – GUYTON, A. C. : **Fisiologia Humana** . Guanabara Koogan, Rio de Janeiro – RJ, 1998.
- 92** - HAAPANIEMI, T., NYLANDER, G., SIRSJÖ, A., et al.: Hyperbaric oxygen reduces ischemia-induced skeletal muscle injury. **The American Society of Plastic and Reconstructive Surgeons**, 1996, 602-607.

- 93** - HAMILTON, M.T., BOOTH, F.W.: Skeletal muscle adaptation to exercise: a century of progress. **Journal of Applied Physiology**, 88, 2000, 327-331.
- 94** - HARGENS, A.R., TIPTON, C.M., GOLLNICK, P.D. et al.: Fluid shifts and muscle function in humans during acute simulated weightlessness. **Journal of Applied Physiology**, 54 (4), 1983, 1003-1009.
- 95** - HOHLFELD, R.; ENGEL, A.G.: The immunobiology of muscle. **Immunology Today**, 15 (6), 1994, 269-274.
- 96**- HOPPELER, H.: Exercise-induced ultrastructural changes in skeletal muscle. **Int. J. Sports Med.**, 7, 1986, 187-204
- 97** - HORSWILL, C.A., LAYMAN, D.K., BOILEAU, R.A., et al.: Excretion of 3-methylhistidine and hydroxyproline following acute weight-training exercise. **Int. J. Sports Med.**, 9 (4), 1988, 245-248.
- 98** - JANSSEN, G.M.E., KUIPERS, H., WILLEMS, G.M., et al.: Plasma activity of muscle enzymes: quantification of skeletal muscle damage and relationship with metabolic variables. **J. Sports Med.**, 10, 1989, 160-168.
- 99** - JASSON, E.: Methodology and actual perspectives of the evaluation of muscular enzymes in skeletal muscle by biopsy during rest, exercise and detraining. **Medicina Dello Sport**. 47 (3), 1994, 377-383.
- 100** - JÄRVINEN, M.: Muscle Injuries in sports. **Sports Exercise and Injury**. 3, 1997, 53-61.
- 101** - JENKINS, R.; KRAUSE, K; SHOFIELD, S.: Influence of exercise on clearance of oxidant stress products and loosely bound iron. **Med. Sci.Sports Exerc.** 25 (2): 1993, 213-217.
- 102**- JI, L.L., FU, R.: Responses of glutathione system and antioxidant enzymes to exhaustive exercise and hydroperoxide. **Journal of Applied Physiology**, 72 (2), 1992, 549-554.
- 103** - JI, L.L.: Oxidative stress during exercise: Implication of antioxidant nutrients. **Free Radical Biology & Medicine**, 18 (6), 1995, 1079-1086.
- 104** – JOHNSON , D. J. ; Et. Al. : Effect of cold submersion on intramuscular temperature of the gastrocnemius muscle. **Phys. Ther.** 59: 1979,1238.

105 - KAKKAR, P., MEHROTRA, S.: VISWANATHAN, P.N. Interrelation of active oxygen species, membrane damage and altered calcium functions. **Molecular and Cellular Biochemistry**, 111, 1992, 11-15.

106 - Kanter, M.: Free radicals, exercise, and antioxidant supplementation. **Int. J. Sports Nutri.** 4, 1994, 205-220.

107 - KASEMKIJWATTANA, C., MENETREY, J., BOSCH, P. et al.: Use of Growth Factors to Improve Muscle Healing After Strain Injury. **Clinical orthopaedics and related research.** 370, Jan. USA: 2000, 272-285.

108 - KEMP, G.J., TAYLOR, D.J., RADDA, G.K.: et al. Bio-energetic changes in human gastrocnemius muscle 1-2 days after strenuous exercise. **Acta Physiol. Scand.** 146, 1992, 11-14.

109- KHASSAF, M., CHILD, R.B., McARDLE A., et al.: Time course of responses of human skeletal muscle to oxidative stress induced by nondamaging exercise. **Journal of Applied Physiology**, 90, 2001, 1031-1035.

110 - KNIGHT, K.L. **Cryotherapy in sport injury management.** USA, Human kinetics, 1995.

111 - KNUTTGEN, H.G.: Human performance in high-intensity exercise with concentric and eccentric muscle contractions. In **Int. J. Sports Med.**, 7, 1986, 6-9.

112 - KOLTYN, K.F.: Analgesia Following Exercise. **Sports Med.** 29 (2), 2000, 85-98.

113 - KOMULAINEN, J., KYTÖLÄ, J., VIJKO, V.: Running-induced muscle injury and myocellular enzyme release in rats. **The American Physiological Society**, 77 (5), 1994, 2299-2304.

114 - KUIPERS, H.: Exercise-Induced Muscle Damage. **J. Sports Med.** 15, 1994, 132-135.

115 - KULIG,K., POWERS, C.M., SHELLOCK, F.G., et al. The effects of eccentric velocity on activation of elbow flexors: Evaluation by magnetic resonance imaging. **Medicine and Science in Sports and Exercise**, 33 (2), 2001, 196-200.

116 - KUMAMOTO, T., UEYAMA, H., WATANABE, S., et al. Muscle fiber degradation in distal myopathy with rimmed vacuoles. **Acta Neuropathol.** 87, 1994, 143-148.

- 117** - KUNTER, M.M., LESMES, G.R., KAMINSKY, L.A., et al. Serum creatine kinase and lactate dehydrogenase changes following an eighty kilometer race. **Eur. Journal of Applied Physiology**, 57, 1988, 60-63.
- 118** - KUTRYK, M.J.B., MADDAFORD, T.G., RAMJIAWAN, B. et al. Oxidation of membrane cholesterol alters active and passive transsarcolemmal calcium movement. **Circulation Research**, 68 (1), Jan, 1991, 18-26.
- 119** - KYLE, B.G. ; **Chemical and process thermodynamics**: USA-NJ; Englewood Cliffs. 1992
- 120** - LAMB, G.D. Ca²⁺ inactivation, Mg²⁺ inhibition and malignant hyperthermia. **Journal of muscle research and cell motility**. 14, 1993, 554-556.
- 121** - LAPIER, T.K., BURTON, H.W., ALMON, R., et al. Alterations in intramuscular connective tissue after limb casting affect contraction-induced muscle injury. **Journal of Applied Physiology** 78 (3), 1995, 1065-1069.
- 122** - LEEUWENBURGH, C., JI, L.L. Glutathione depletion in rested and exercise mice biochemical consequence and adaptation. **Archives of Biochemistry and Biophysics**, 316 (2), 1995, 942-949.
- 123** - Lekil J. : **Entropy and Thermodynamics**, Springfield, IL : CCT Publisher, 1975
- 124** - LEHTO, M.U.K., JÄRVINEN, M.J., Muscle injuries, their healing process and treatment. **Annales Chirurgiae et Gynaecologiae**. 80, 1991, 102-108.
- 125** - LIEBER, R.L., SCHMITZ, M.C., MISHRA, D.K., et al. Contractile and cellular remodeling in rabbit skeletal muscle after cyclic eccentric contractions. **Journal of Applied Physiology**, 77 (4), 1994, 1926-1934.
- 126** - LOCKE, M., NOBLE, E.G. Stress proteins: The exercise response. **Can. Journal of Applied Physiology**, 20 (2), 1995, 155-167.
- 127** - LÓPEZ, J.R., SÁNCHEZ, V., LIZARRAGA, G., et al. Modificaciones en la concentración basal de calcio libre mioplasmico inducida por dantroleno sodico en sujetos susceptibles a hipertermia maligna. **Rev. Arg. Anest.** 46 (4), 1988, 149-156.
- 128** - LÓPEZ, J.R., PARRA, L. ositol 1,4,5-trisphosphate increases myoplasmic [Ca²⁺] in isolated muscle fibers. Despolarization enhances its effects. **Cell Calcium**, 12, 1991, 543-557.

- 129** - LOWE, D.A., WARREN, G.L., HAYES, D.A.F., et al. Eccentric contraction-induced injury of mouse soleus muscle: Effect of varying [Ca²⁺]0. **The American Physiological Society.** 161 (7567), 1994, 1445-1453.
- 130** - LYNCH, G.S., WILLIAMS, D.: The effect of lowered pH on the Ca²⁺-activated contractile characteristics of skeletal muscle fibres from endurance-trained rats. **Experimental Physiology,** 79, 1994, 47-57.
- 131** - MACAULEY, D: Do textbooks agree on their advice on ice ?. **Clin. J. Sport Med.;** 11(2): 2001,67:72.
- 132** - MACINTYRE, D. L.; REID, W.D.; MCKENZIE, D. C.: Delayed soreness: The Inflammatory response to muscle injury and its clinical implications. **Sports Med.** 20(1): 1995,24-40.
- 133** - MACKINNON, L.; TOMASI, T. : **Immunology of exercise.: Sports Medicine, Fitness, Training and Injuries.** Appenzeller, o (Ed.). New Mexico University , Baltimore, Munich, pp. 273-289, 1988.
- 134** - MANTZ, J., HINDELANG, C. MANTZ, J.M., et al.: Muscle regeneration after exercise- induced electron microscopic study. **Virchows Archiv. A Pathol. Anat.,** 423, 1993, 91-95.
- 135** - MARGARITIS, I., TESSIER, F., VERDERA, F., et al.: Muscle enzymes release does not predict muscle function impairment after triathlon. **The Journal of Sports Medicine and Physical Fitness,** 39 (2), 1999, 133-139.
- 136** - MAJOMAKI, V. ; SALMINEN, A. : Morfological and enzymatic heterogeneity of suramin-induced lysosomal storage disease in some tissues of mice and rats. **Experimental and molecular pathology,** 45 , 1986,76-83.
- 137** - MARTIN, S.S. et. Al.: Cryotherapy: an effective modality for decreasing intraarticular temperature after knee arthroscopy. **Am. J. Sports Med.** 29(3): 2001,288-91.
- 138** - MATSEN, F.A.; QUESTD, K; MATSEN, A.L.: The effect of local cooling on post fracture swelling. **Clin. Orthop.** 1975,109:201.
- 139** - McCOMAS, A.J. **Skeletal muscle:** Form and funtion. USA, Human kinetics, 1996.
- 140** - McMASTER, W. C., LIDLLE, S.: Cryotherapy influence on post traumatic limb edema. **Clin. Orthop.** 1980,150:238.

- 141** - McCULLY, K., SHELLOCK, F.G., BANK, W.J., et al. The use of nuclear magnetic resonance to evaluate muscle injury. **Medicine and Science in Sports and Exercise**, 24 (5), 1991, 537-542.
- 142** - MEHROTRA, S., KAKKAR, P., VISWANATHAN, P.N. Mitochondrial damage by active oxygen species in vitro. **Free Radic. Biol. Med.**, 10, 1991, 277-285.
- 143** - MEYDANE, M., EVANS, W.J., HANDELMAN, G., et al. Protective effect of vitamin E on exercise-induced oxidative damage in young and older adults. **Am. J. Physiol.**, 264, 1993, 992-998.
- 144** - MICHLOVITZ, S.L. **Thermal agents in rehabilitation**. Philadelphia, F.A. Davis company, 1996.
- 145** - MILES, M.P., CLARKSON, P.M. Exercise-induced muscle pain, soreness, and cramps. **The journal of sports medicine and physical fitness**. 34 (3), 1994, 203-216.
- 146** - MORGAN, D.L., ALLEN, D.G. Early events in stretch-induced muscle damage. **J. Appl. Physiol**, 87 (6), 1999, 2007-2015.
- 147** – MONTEIRO, M. R. M. : Agressão/lesão muscular esquelética em crianças induzida pelo exercício; influência do tipo predominante de contracções na manifestação de diferentes indicadores indirectos. Dissertação apresentada às provas de mestrado no ramo de Ciências do Desporto,. Faculdade de Ciências do Desporto Educacão Física, **Universidade do Porto**. 1996.
- 148** - MOSCOVITZ, B., KATZ, Y., SINGER, P., et al. Glutamine metabolism and utilization : relevance to major problems in health care. **Pharmacological Research**, 30 (1), 1994, 61-71.
- 149** - MOXLEY, R.P. Evaluation of neuromuscular function in inflammatory myopathy. **Rheumatic Disease Clinics of North America**, 20 (4), Nov, 1994, 827-843.
- 150** -NAGAMATSU, M., SCHMELZER, J.D., ZOLLMAN, P.J., et al. Ischemic reperfusion causes lipid peroxidation and fiber degeneration. **Muscle & Nerve**, 19, Jan, 1996, 37-47.
- 151** – NAVÉ, C.R.; NAVÉ, B. C. : **Physics for the health sciences**, ed 2. WB Saunders, Philadelphia , 1980.

152 - NEWHAM, D.J. The consequences of eccentric contractions and their relationship to delayed onset muscle pain. **Eur. Journal of Applied Physiology**, 57, 1988, 353-359.

153 - NIEMAN, D.C., MILLER, A.R., HENSON, D.A., et al. Effects of high-vs moderate- intensity exercise on natural killer cell activity. **Med. Sci. Sports Exerc.**, 25 (10), 1993, 1126-1134.

154 - NOSAKA, K., CLARKSON, P.M. Relationship between post-exercise plasma CK elevation and muscle mass involved in the exercise. **Int. J. Sports Med.**, 13 (6), 1992, 471-475.

155 - NOSAKA, K., CLARKSON, P.M. Effect of eccentric exercise on plasma enzyme activities previously elevated by eccentric exercise. **Eur. Journal of Applied Physiology**, 68, 1994, 492-497.

156 - NOSAKA, K., CLARKSON, P.M. Muscle damage following repeated bouts of high force eccentric exercise. **Med. Sci. Sports Exec.**, 27 (9), 1995, 1263-1269.

157 - NOSAKA, K., KURAMATA, T. Muscle soreness and serum enzyme activity following consecutive drop jumps. **J. Sports Sciences**, 9, 1991, 213-220.

158 - OHNO, H., GASA, S., HABARA, Y., et al. Effects of exercise stress and cold stress on glutathione and γ - glutamyltransferase in rat liver. **Biochimica et Biophysica Acta**, 1033, 1990, 19-22.

159 - PACKER, L. : **Oxidants and antioxidants and the biological effects of physical exercise.: Biological Effects of Physical Activity.** Williams, R.S. & Wallace , A.G. (Eds.) HKP Sport Science Monograph Series. Vol 2 . Human kinetics Books, Champaign, Illinois. USA. Pp. 85-90, 1989.

160 - PALMO, J., ASP, S., DAUGAARD, J.R., et al: Effect of eccentric exercise on natural killer cell activity. **Journal of Applied Physiology**, 78 (4), 1995, 1442-1446.

161 - PEDERSON, B. : **Exercise and immunity – mechanisms of action in: Integration of Medical and Sports Sciences** . Sato Y., Poortmans, J. Hashimoto, I., Oshida, I (Eds.). Basel . Karger . vol 17. pp 33-39, 1992

162 - PEREIRA, B., COSTA ROSA, L.F.B., SAFI, D.A., et al. Superoxide dismutase, catalase, and glutathione peroxidase activities in muscle and lymphoid organs of sedentary and exercise-trained rats. **Physiology & Behavior**, 56 (5), 1994, 1095-1099.

163 - PEREIRA, B., COSTA ROSA, L.F.B., SAFI, D.A., et al. Antioxidant enzyme activities in the lymphoid organs and muscle of rats fed fatty acids-rich diets subjected to prolonged physical exercise-training. **Physiology & behavior**, 56 (5), 1994, 1049-1055.

164 - PEREIRA, J.F.M.P. : Exercícios físicos inabituais e exaustivos em crianças. Fluência do tipo predominante de contrações em indicadores indirectos de stress oxidativo e de agressão muscular esquelética. Dissertação apresentada às provas de mestrado no ramo de Ciências do Desporto,. Faculdade de Ciências do Desporto Educacão Física, **Universidade do Porto**. 1996.

165 - PERKINS, J., et. Al.: Cooling and contraction of smooth muscle. **Am. J. Physiol.** 1950, 163:14.

166 - PETTE, D., VRBOVÁ, G. Adaptation of mammalian skeletal muscle fibers to chronic electrical stimulation. **Rev. Physiol. Biochem. Pharmacol.** 120, 1992, 115-202.

167 - PIZZA, F.X., MITCHELL, J.B., DAVIS, B.H., et al. Exercise-induced muscle damage: Effect on circulating leukocyte and lymphocyte subsets. **Med. Sci. Sports Exerc.**, 27 (3), 1995, 363-370.

168 - PIZZA, F.X.; MITCHELL, J.B.; DAVIS, B.H.; STARLING, R.D.; HOLTZ, R.W.; BIGELOW, N. : Exercise-induced muscle damage: effect on circulating leukocyte and lymphocyte subsets. **Med. Sci. Sports Exerc.** 27 (3) 1995, 363-370.

169 - PLAGHKI, E., SCHOOR, M.C.V., BLEUKX, G.B., et al.: Creatine, creatine kinase and glycolytic enzymes in regenerating rat casctrocnemius. **Adv. Physiol Sci.**, 24, [?].

170 - POWERS, S.K., CRISWELL, D., LAWLER, J., et al.: Fluence of exercise and fiber type on antioxidant enzyme activity in rat skeletal muscle. **Am. J. Physiol.**, 266 (35), 1994, 375-380.

171 - RANTANEN, J., THORSSON, O., WOLLMER, P., et al.: Effects of therapeutic ultrasound regeneration of skeletal myofibers after experimental muscle injury. **The American Journal of Sports Medicine**, 27 (1), 1999, 54-59.

172 - RASMUSSEN, U.F., RASMUSSEN, H.M. Human skeletal muscle mitochondrial capacity. **Acta Physiol Scand.**, 168, 2000, 473-480.

173 - ROBERTSON, T.A., PAPADIMITRIOU, J.M., GROUNDS, M.D. Fusion of myogenic cells to the newly sealed region of damaged myofibres in skeletal muscle regeneration. **Neuropathology and Applied Neurobiology**, 19, 1993, 350-358.

174 - RODENBURG, J.B., STEENBEEK, D., SCHIEREK, P., et al. Warm-up, stretching and massage diminish harmful effects of eccentric exercise. **Int. J. Sports Med.**, 15, 1994, 414-419.

175 - RODENBURG, J.B., BOER, R.W., SCHIERECK, P., et al. Changes in phosphorus compounds and water content in skeletal muscle due to eccentric exercise. **Eur. Journal of Applied Physiology**, 68, 1994, 205-213.

176 - RODENBURG, J.B., DE GROOT, M.C.H., VAN ECHTELD, C.J.A., et al. Phosphate metabolism of prior eccentrically loaded vastus medialis muscle during exercise in humans. **Acta Physiol. Scand.**, 153, 1995, 97-108.

177 - SAHLIN, K., HARRIS, R.C., HULTMAN, E. Creatine Kinase equilibrium and lactate content compared with muscle pH in tissue samples obtained after isometric exercise. **Biochem. J.** 152, 1975, 173-180.

178 - SALMINEN, A., HONGISTO, K., VIJKO, V. Lysosomal changes related to exercise injuries and training-induced protection in mouse skeletal muscle. **Acta Physiol. Scand.**, 120, 1984, 15-19.

179 - SALMINEN, A., K., VIJKO. Lipid peroxidation in exercise myopathy. **Exper. and Molec. Patho.**, 38, 1983, 380-388.

180 - SALMINEN, A., K., VIJKO. Autophagic response to strenuous exercise in mouse skeletal muscle fibers. **Virchows Arch [Cell Pthol]**, 45, 1984, 97-106.

181 - SASTRE, J.; ASENSI, M.; GASCÓ, E.; PALLARDÓ , F. ; FERRERO, J; FURUKAWA, T .; VIÑA, J. : Exhaustive physical exercise causes oxidation of glutathione status in blood; prevention and antioxidant administration. **Am. J. Physiol.** 263: 1992, R992- R995.

182 - SAYERS, S.P., CLARKSON, P.M., LEE, J. Activity and immobilization after eccentric exercise: Recovery of muscle function. **Med. Sci. Sports Exerc.**, 32 (9), 2000, 1587-1592.

183 - SCHÄFER, M., SCHLEGEL, C., KIRLUM, H.-J., et al. Monitoring of damage to skeletal muscle tissues caused by ischemia. **Bioelectrochemistry and Bioenergetics**, 45, 1998, 151-155.

184 - SEJERSTED, O.M., HARGENS, A.R., KARDEL, K.R., et al. intramuscular fluid pressure during isometric contraction of human skeletal muscle. **Journal of Applied Physiology**, 56 (2), 1984, 287-295.

185 - SEN, G.K., MARIN, E., KRETZSCHMAR, M., et al. Skeletal muscle and liver glutathione homeostasis in response to training, exercise, and immobilization. **Journal of Applied Physiology**, 73 (4), 1992, 1265-1272.

186 - SEN, G.K., ATALAY, M., HÄNNINEN, A.O. Exercise- induced oxidative stress: Glutathione supplementation and deficiency. **Journal of Applied Physiology**, 77 (5), 1994, 2177-2187.

187 - SEN , C.K. : Oxidants and antioxidants in exercise . **Journal of Applied Physiology**. 79(3): 1995, 675-686.

188 - SHELLOCK, F.G., FUKUNAGA, T., MINK, J.H., et al. Acute effects of exercise on MR imaging of skeletal muscle: Concentric vs eccentric actions. **AJR**, 156, 1991, 765-768.

189 - SHELLOCK, F.G., FUKUNAGA, T., MINK, J.H., et al. Exertional muscle injury: Evaluation of concentric versus eccentric actions with serial MR imaging. **Radiology**, 179, 1991, 659-664.

190 - SHEPARD, R.; VERDE, T.; THOMAS, S.; SHECK, P : Physical activity and the immune system . **Can. J. Sports. Sci.** 16 (3): 1991,163-185.

191 - SHRIER, I. Stretching before exercise does not reduce the risk of local muscle injury : a critical review of the clinical and basic science literature. **Clinical Journal of Sport Medicine**, 9, 1999, 221-227.

192 -SIMON,H.B. The immunology of exercise: A brief review. **JAMA**, 252 (19), 1984, 2735-2738.

193 - SINERT, R., KOHL, L., RAINONE, T., et al. Exercise- induced rhabdomyolysis. **Annals of Emergency Medicine**, 23 (6), 1994, 1301-1306.

194 - SMITH, J.K., CARDEN, D.L., KORTHUIS, R.J. Role of xanthine oxidase in postischemic microvascular injury in skeletal muscle. **Am. J. Physiol.**, 257, 1989, 782-789.

195- SMITH, T. L. ; et. Al.: New skeletal muscle model for the longitudinal study of alterations in microcirculation following contusion and cryotherapy. **Microsurgery** 14: 1993, 487-492.

- 196** - SOARES, J.M.C., MOTA, P., DUARTE, J.A.R., et al. Children are less susceptible to exercise - induced muscle damage than adults : A preliminary investigation. **Pediatric Exercise Science**, 8 (4), 1996, 361-367.
- 197** - SOARES, J.M.C. Effects of training on muscle capillary pattern : intermittent vs. Continuous exercise. **J. Sports Med. Phys. Fitness**, 32, 1992, 123-127.
- 198** - SOARES, J.M.C., DUARTE, J.A.R., CARVALHO, J., et al. The possible role of intracellular Ca^{2+} accumulation for the development of immobilization atrophy. **J. Sports Med.**, 14 (8), 1993, 437-439.
- 199** - STAUBER, W.T., CLARKSON, P.M., FRITZ, V.K., et al. Extra cellular matrix disruption and pain after eccentric muscle action. **Journal of Applied Physiology**, 69 (3), 1990, 868-874.
- 200** - STAUBER, W.T., FRITZ, V.K., VOGELBACH, D.W., et al. Characterization of muscle injured by forced lengthening. I. Cellular infiltrates. **Med. Sci. Sports Exerc.**, 20 (4), 1988, 345-353.
- 201** - STÄUBLI, M., ROESSLER, B., KÖCHLI, H.P., et al. Creatine Kinase and creatine kinase MB in endurance runners and patients with myocardial infarction. **Eur. Journal of Applied Physiology**, 54, 1985, 40-45.
- 202** - STEIGER, F.: **Thermodynamics**. USA, Boston Press, 1995.
- 203** - STITT, J.T., SCHIMADA, S.G. Site of action of calcium channel blockers in inhibiting endogenous pyrogen fever in rats. **Journal of Applied Physiology**, 71 (3), 1991, 956-960.
- 204** - SULLIVAN, T.E., ARMSTRONG, R.B. Rat locomotory muscle fiber activity during trotting and galloping. **Journal of Applied Physiology**, 44 (3), 1978, 358-363.
- 205** - SYLVEDÉN, C., JANSSON, E., KALLNER, A., et al. Human Creatine Kinase. **Scand. J. Clin. Lab.vest.**, 44, 1984, 611-615.
- 206** - TAKAHASHI, H., KUNO, S., MIYAMOTO,T., et al. Changes in magnetic resonance images in human skeletal muscle after eccentric exercise. **Eur. Journal of Applied Physiology**, 69, 1994, 408-413.
- 207** - TAKEKURA, H., SUN, X., FRANZINI-ARMSTRONG, C. Development of the excitation-contraction coupling apparatus in skeletal muscle : peripheral and internal calcium release units are formed sequentially. **Journal of Muscle Research and Cell Motility**, 15, 1994, 102-118.

- 208** - TESSIER, F., MARGARITIS, I., RICHARD,M., et al. Selenium and training effects on the glutathione system and aerobic performance. **Med. Sci. Sports Exerc.**, 27 (3), 1995, 390-396.
- 209** - THAYER, R.E., RICE, C.L., PETTIGREW, F.P., et al. The fibre composition of skeletal muscle. **Med. Sport Sci.**, 38, 1993, 25-50.
- 210** - THIEL, M., BARDENHEUER, H. Regulation of oxygen radical production of human polymorphonuclear leukocytes by adenosine : the role of calcium. **Eur. J. of Physiol.**, 420, 1992, 522-528.
- 211** - TIDBALL, J.G., SALEM, G., ZERNICKE, R. Site and mechanical conditions for failure of skeletal muscle in experimental strain injuries. **Journal of Applied Physiology**, 74 (3), 1993, 1280-1286.
- 212** - TIIDUS, P.M. Can strogens diminish exercise induced muscle damage ? In **Can. Journal of Applied Physiology**, 20 (1), 1995, 26-38.
- 213** - TIIDUS, P.M. Strogen and gender effects on muscle damage, inflammation, and oxidative stress. **Can. Journal of Applied Physiology**, 25 (4), 2000, 274-287.
- 214** - TURINSKY, J., O'SULLIVAN, D.M., BAYLY, B.P. Modulation of prostaglandin E₂ synthesis in rat skeletal muscle. **Am. J. Physiol.**, 262 (25), 1992, 476-482.
- 215** - VANDENBURGH, H.H., HATFALUDY, S., SOHAR, I., et al. Stretch-induced prostaglandins and protein turnover in cultured skeletal muscle. **Am. J. Physiol.**, 259 (28), 1990, 232-240.
- 216** - VECCHIET, L., VECCHIET, J., BELLOMO, R. Muscle pain from physical exercise. **Journal of Musculoskeletal pain**, 7 (1/2), 1999, 43-53.
- 217** - VERDUCCTI, F.M.terval cryotherapy decreases fatigue during repeated weight lifting. **Journal of Athletic Training**, 35 (4), 2000, 422-426.
- 218** - VIRTANEN, P., VIITASALO, J.T., VUORI, J., et al. Effect of concentric exercise on serum muscle and collagen markers. **Journal of Applied Physiology**, 75 (3), 1993, 1272-1277.
- 219** - VOLFINGER, L., LASSOURD , V., MICHAUX, J.M., et al. Kinetic evaluation of muscle damage during exercise by calculation of amount of creatine kinase released. **Am. J. Physiol.**, 266 (35), 1994, 434-441.

220 - WAKIM, K. G. , PORTER, A.N., KRUSEN, K. H. : Influence of physical agents of certain drugs on intra articular temperature. **Arch. Physc. Med.** 1953, 2:714, 1.

221 - WALKER, J.A., CERNY, F. J., COTTER, J.R., et al. Attenuation of contraction-induced skeletal muscle injury by bromelain. **Med. Sci. Sports Exerc.**, 24 (1), 1992, 20-25.

222 -WARREN, G.L., LOWE, D.A., ARMSTRONG, R.B. Measurement tools used in the study of eccentric contraction-induced injury. **Sports Med.**, 27 (1), 1999, 43-59.

223 - WATERMAN-STORER, C.M. The cytoskeletal of skeletal muscle : Is it affected by exercise ? a brief review. **Med. Sci. Sports Exerc.**, 23 (11), 1991, 1240-1249.

224 - WELSH, D.G., LINDINGER, M.I. L-type Ca^{2+} channel and $\text{Na}^+ / \text{Ca}^{2+}$ exchange inhibitors reduce Ca^{2+} accumulation in reperfused skeletal muscle. **Journal of Applied Physiology**, 80 (4), 1996, 1263-1269.

225 / 3012 - WEISS, L. M. : Oxygem, isquemia and inflammation. **Acta Physiol. Scand. Suppl.** 548: 9-36, 1986

226 - WESTERBLAD, H., DUTY, S., ALLEN, D.G. tracellular calcium concentration during low-frequency fatigue in isolated single fibers of mouse skeletal muscle. **Journal of Applied Physiology**, 75 (1), 1993, 382-388.

227 - WILLIANS, J.H., KLUG, G.A., Calcium exchange hypothesis of skeletal muscle fatigue: A brief review. **Muscle & Nerve**, 18, 1995, 421-434.

228 -WILSON, G.J., MURPHY, A.J., PRYOR, J.F. Musculotendinous stiffness: its relationship to eccentric, isometric, and concentric performance. **Journal of Applied Physiology**, 76 (6), 1994, 2714-2719.

229 - WOOD, S.A., MORGAN, D.L., PROSKE, U. Effects of repeated eccentric contractions on structure and mechanical properties of toad sartorius muscle. **Am. J. Physiol.**, 265 (34), 1993, 792-800.

230 - YTREHUS, K., REIKERÅS, O., HUSEBY, N., et al. Ultrastructure of reperfused skeletal muscle: The effect of oxygen radical scavenger enzymes. **Int. J. Microcirc.**, 15, 1995, 155-162.

231 - ZIMMERMANN, M. Components of pain and pain- associated phenomena in the musculoskeletal system. **The Parthenon Publishing Group**, [?], 1990, 29-52.

232 - ZIMÁNYI, I., BUCK, E., ABRAMSON, J.J., et al. Ryanodine induces persistent inactivation of the Ca^{2+} release channel from skeletal muscle sarcoplasmic reticulum. **Molecular Pharmacology**, 42, 1992, 1049-1057.

