



84^o **SIC**
SOCIETÀ ITALIANA DI CARDIOLOGIA



14 - 17 Dicembre 2023

Rome Cavalieri

Come, quando e perché CPET + emodinamica non invasiva

Gaia Cattadori

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CORRESPONDENCE

Research Correspondence

Noninvasive Measurement of Cardiac Output During Exercise by Inert Gas Rebreathing Technique: A New Tool for Heart Failure Evaluation

***Piergiuseppe Agostoni, MD, PhD**

Gaia Cattadori, MD

Anna Apostolo, MD

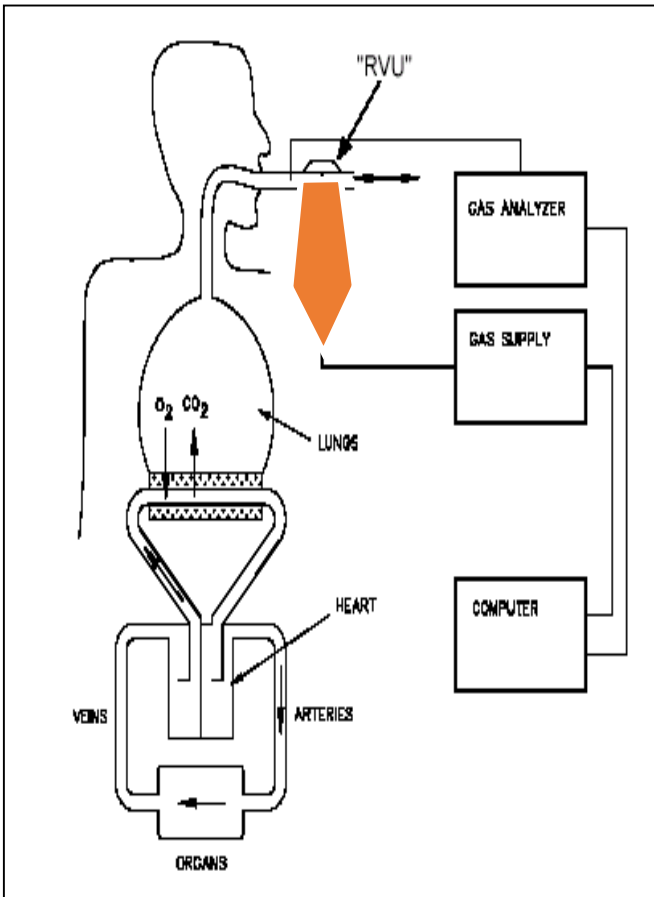
Mauro Contini, MD

Pietro Palermo, MD

Giancarlo Marenzi, MD

Karlman Wasserman, MD, PhD





▪ Respirazione in circuito chiuso connessi a un palloncino di gomma pre-riempito con miscela arricchita di ossigeno con due gas estranei

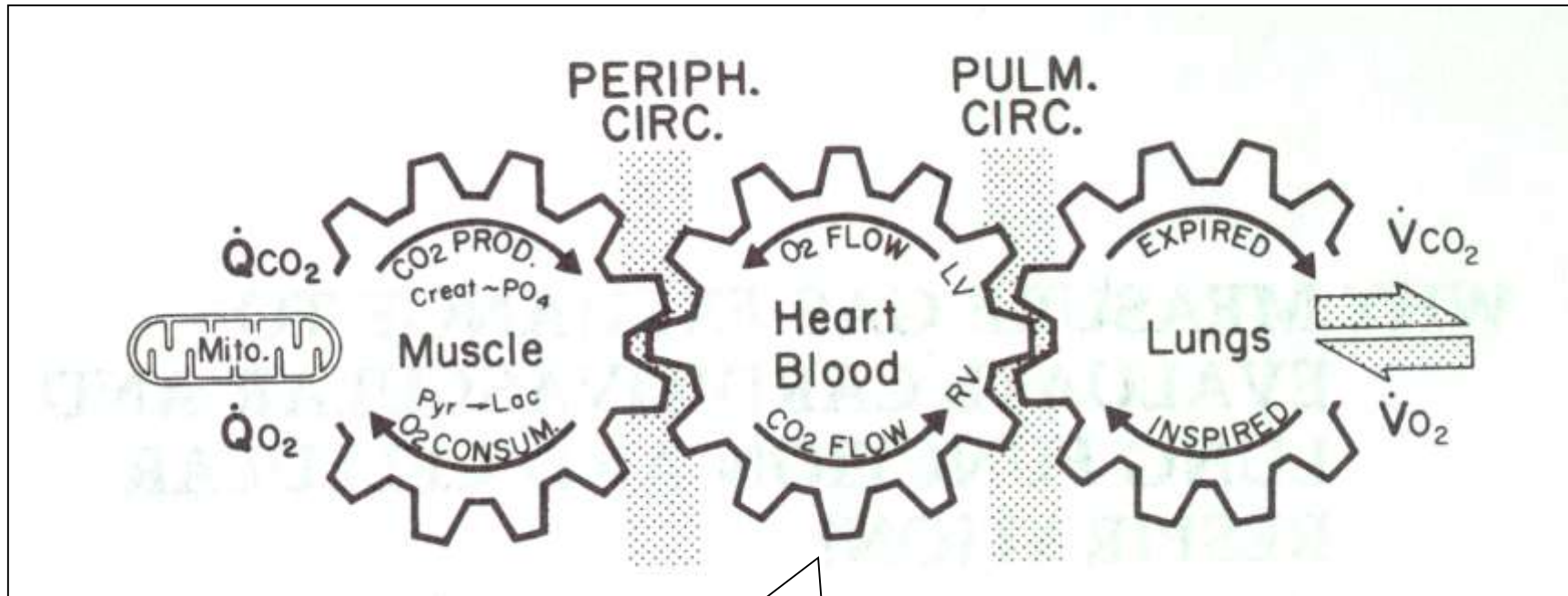
- gas insolubile esafluoruro di zolfo (SF_6) nel sangue
- gas solubile ossido nitroso (N_2O) nel sangue.

▪ Switch automatico di connessione verso aria ambiente alla fine del test

▪ Misurazione continua on-line dei gas con analizzatore fotoacustico alla bocca (più veloce e stabile di uno spettrometro di massa)



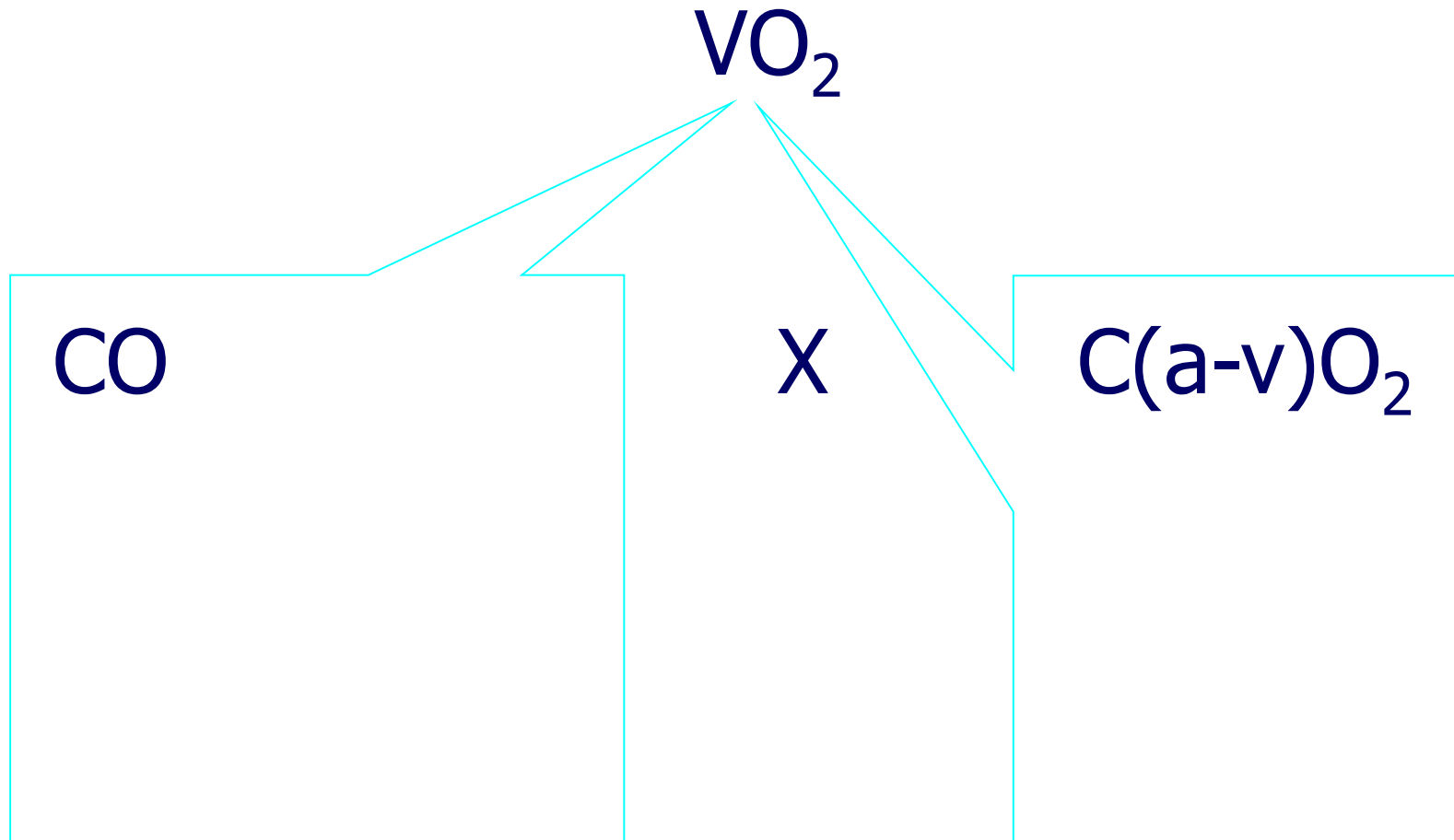
$\dot{V}O_2$



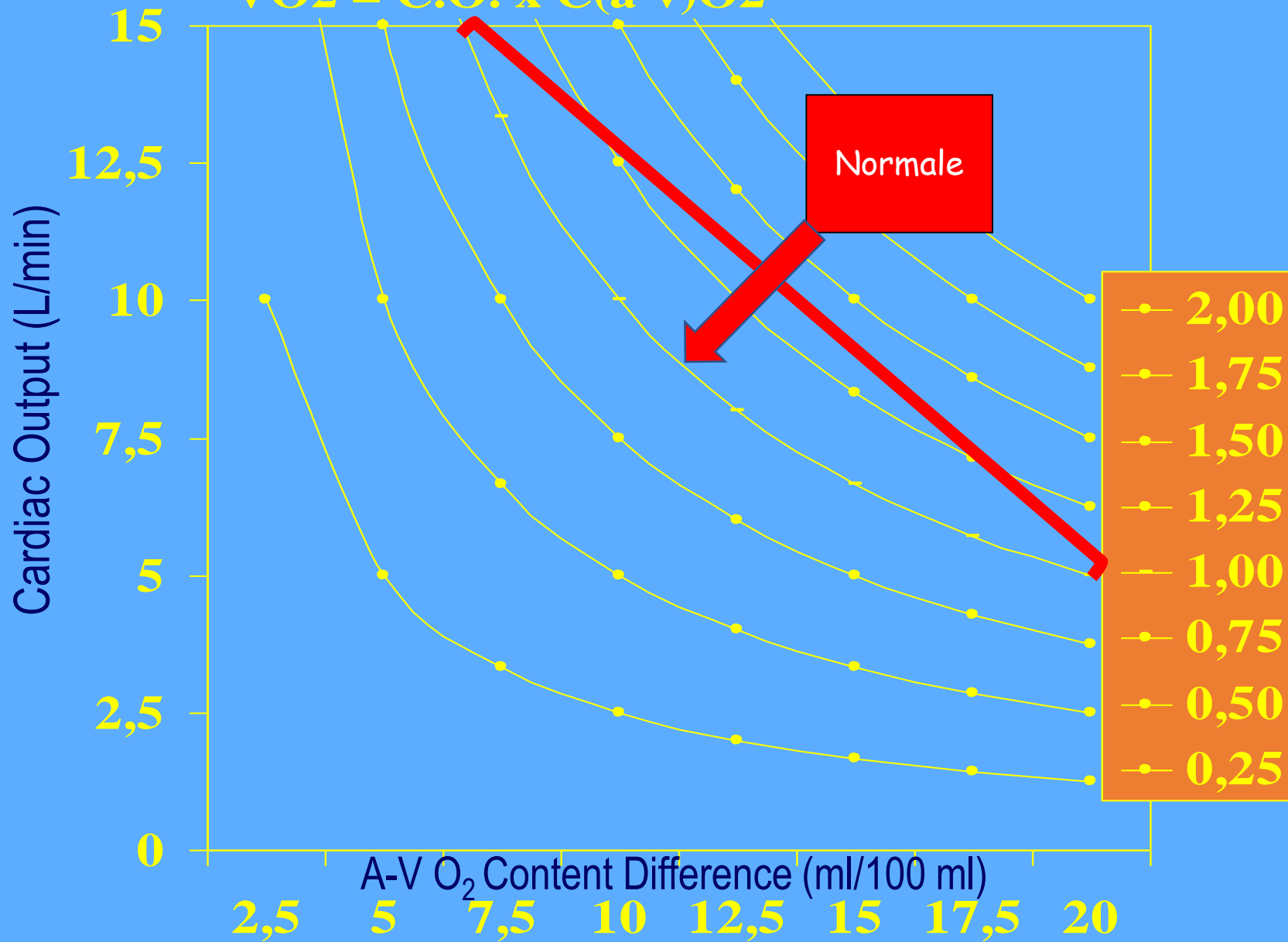
Cardiac Output



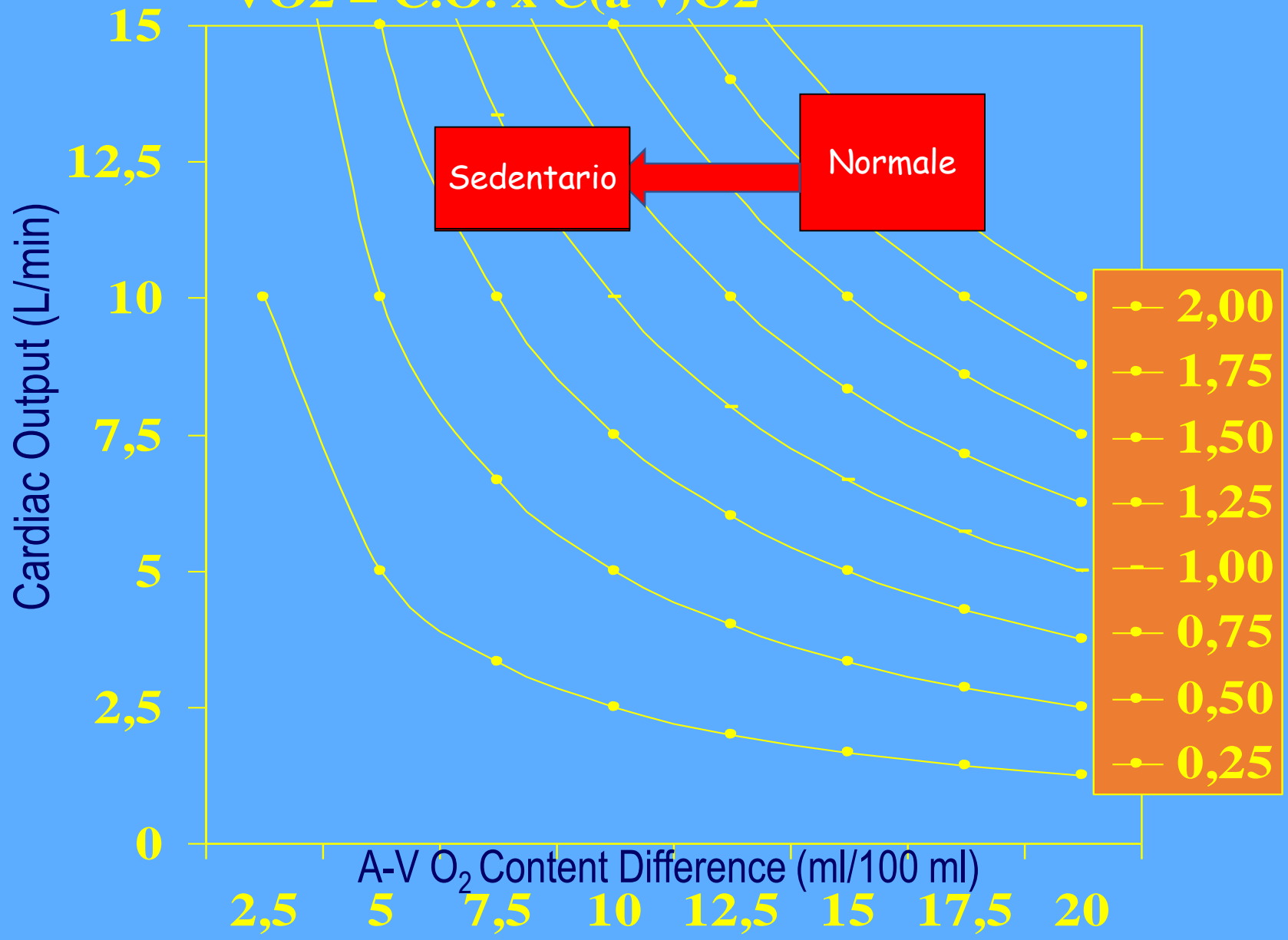
Legge di Fick



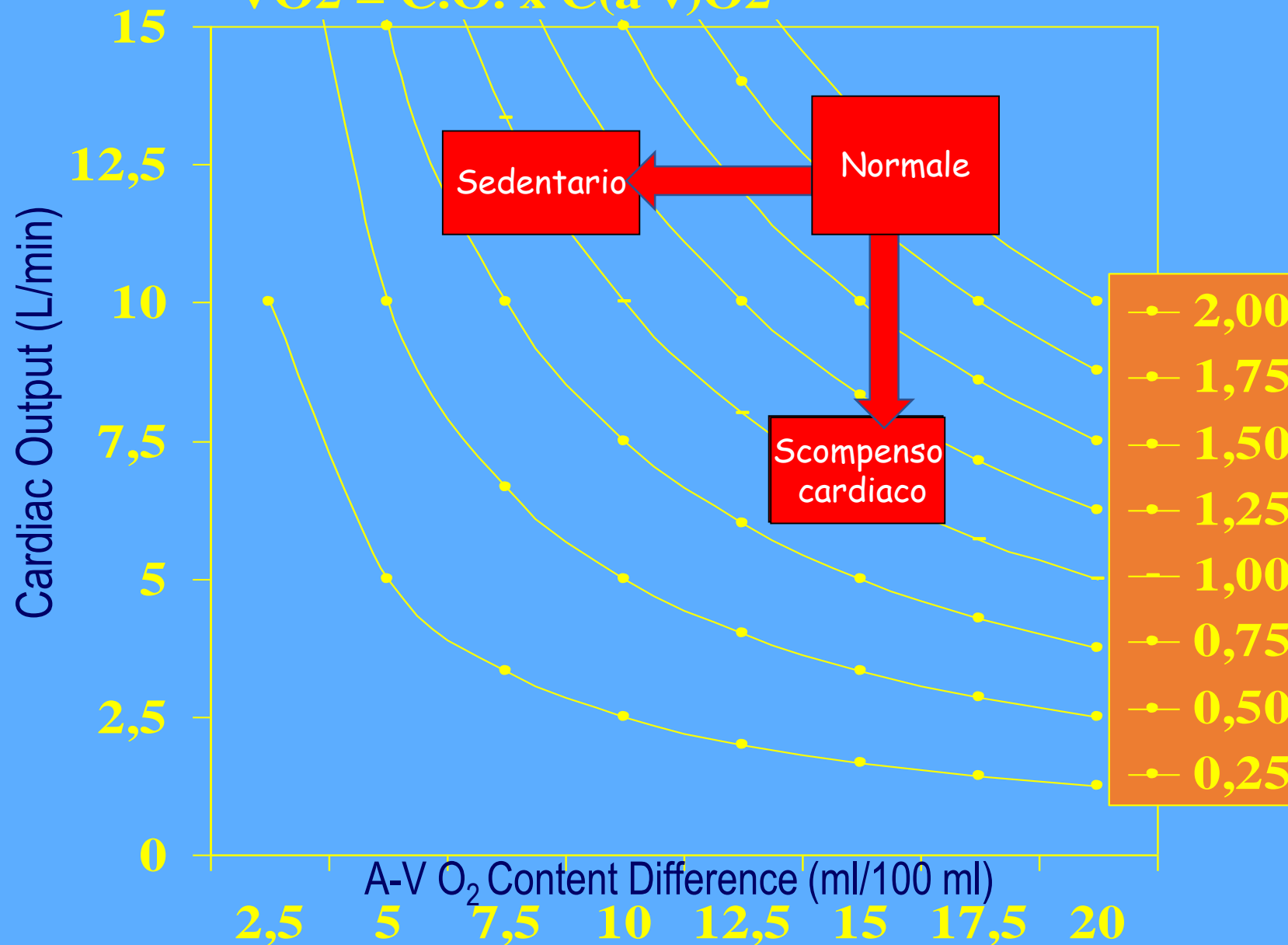
$$VO_2 = C.O. \times C(a-v)O_2$$



$$VO_2 = C.O. \times C(a-v)O_2$$



$$VO_2 = C.O. \times C(a-v)O_2$$



Legge di Fick

VO_2

CO

- Gettata sistolica
- Frequenza cardiaca

X

$C(a-v)O_2$

- Hb
- SatHb O_2
- pO_2



Legge di Fick

VO_2

CO

- Gettata sistolica
- Frequenza cardiaca

X

$C(a-v)O_2$

- **Hb**
- SatHb O_2
- pO_2



Quanto conta l'anemia?

1 gr di Hb porta ai tessuti 1 ml O₂ x dl di sangue

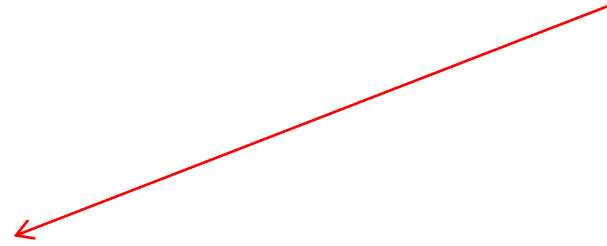
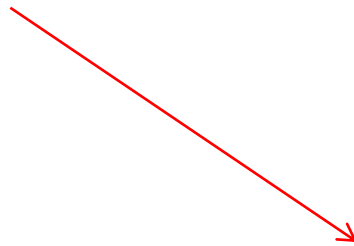


Quanto conta l'anemia?

1 gr di Hb porta ai tessuti 1 ml O₂ x dl di sangue

Se Cardiac Output al picco Ex = 10 L/

Anemia 5 gr/dl



Quanto conta l'anemia?

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Se Cardiac Output al picco Ex = 10 L/'

Anemia 5 gr/dl

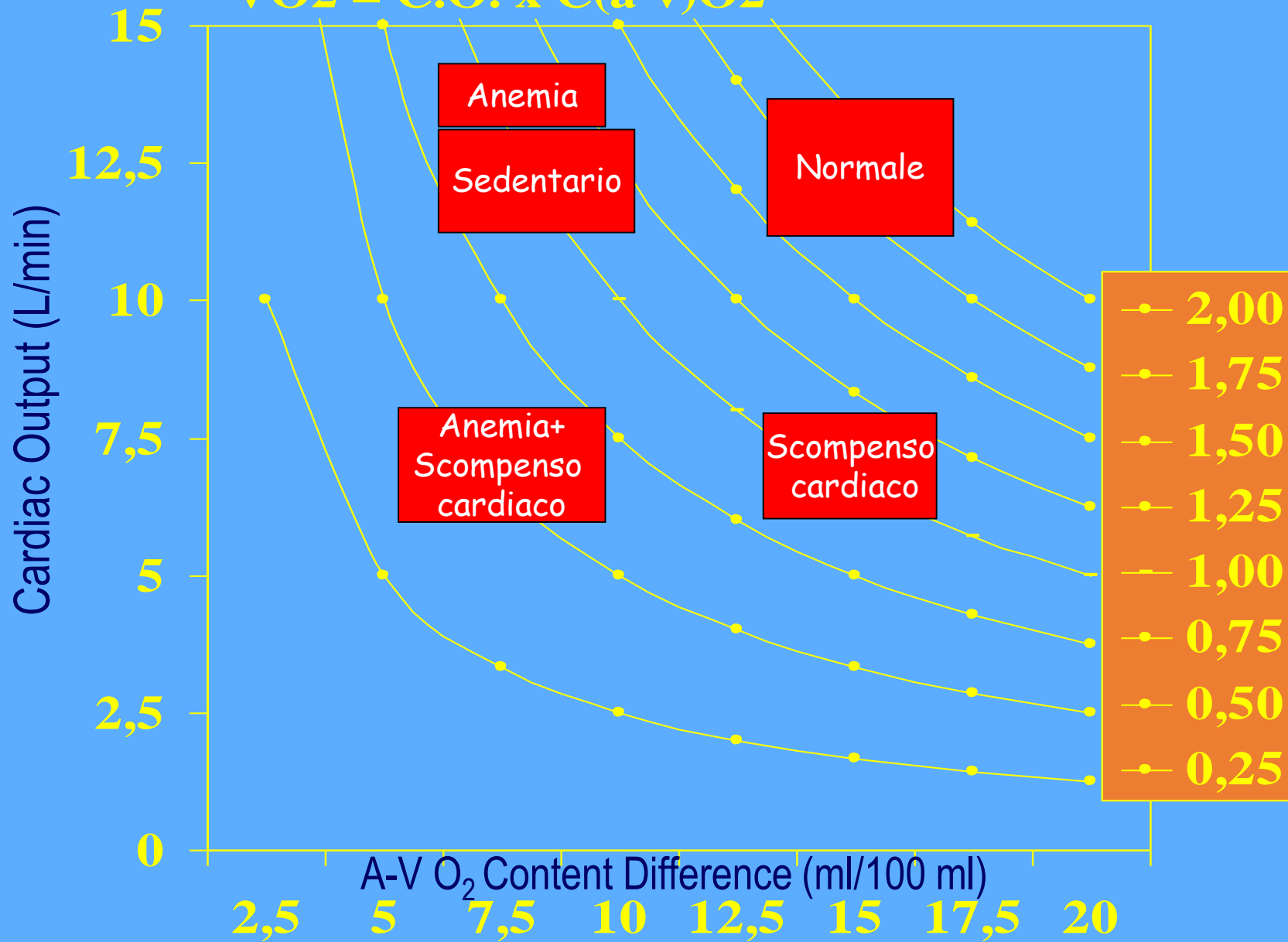
VO₂Max ridotto di

5x100 = 500 ml/'

solo per anemia



$$VO_2 = C.O. \times C(a-v)O_2$$



Teoria

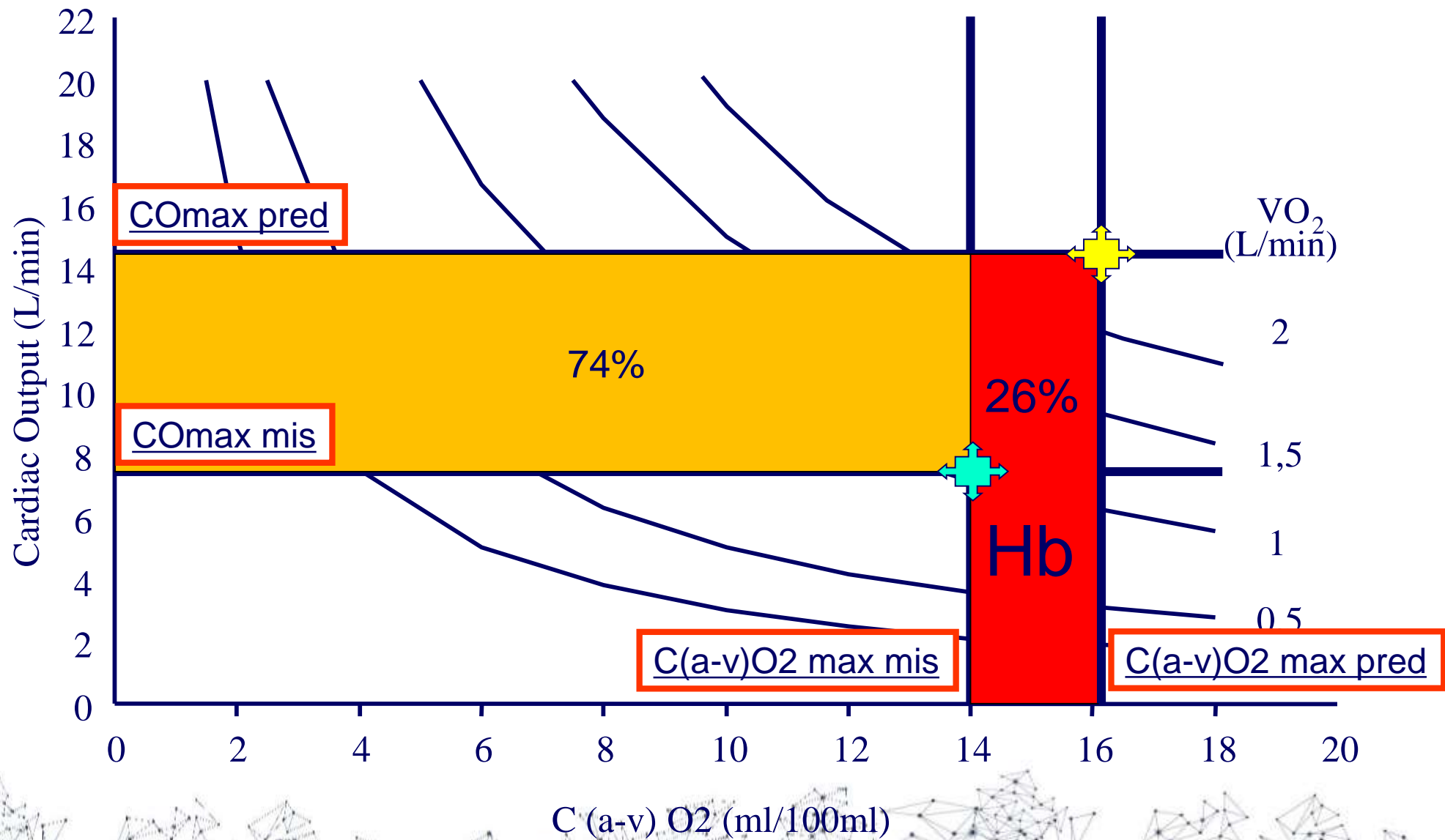


Teoria → Applicazione clinica

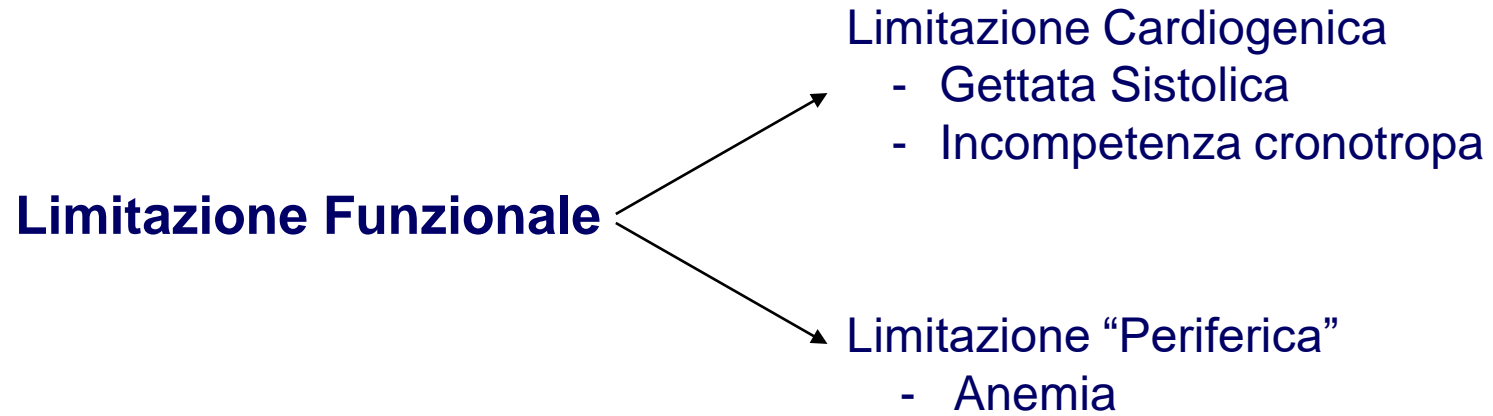


VO_2 Pred = 2120 ml/min

VO_2 Measured 1080 ml/min (51%pred)



Come, quando e perché
CPET + emodinamica non invasiva



Use of Cardiopulmonary Exercise Testing With Hemodynamic Monitoring in the Prognostic Assessment of Ambulatory Patients With Chronic Heart Failure

Marco Metra, MD,* Pompilio Faggiano, MD,† Antonio D'Aloia, MD,* Savina Nodari, MD,* Anna Gualeni, MD,* Domenica Raccagni, MD,* Livio Dei Cas, MD*

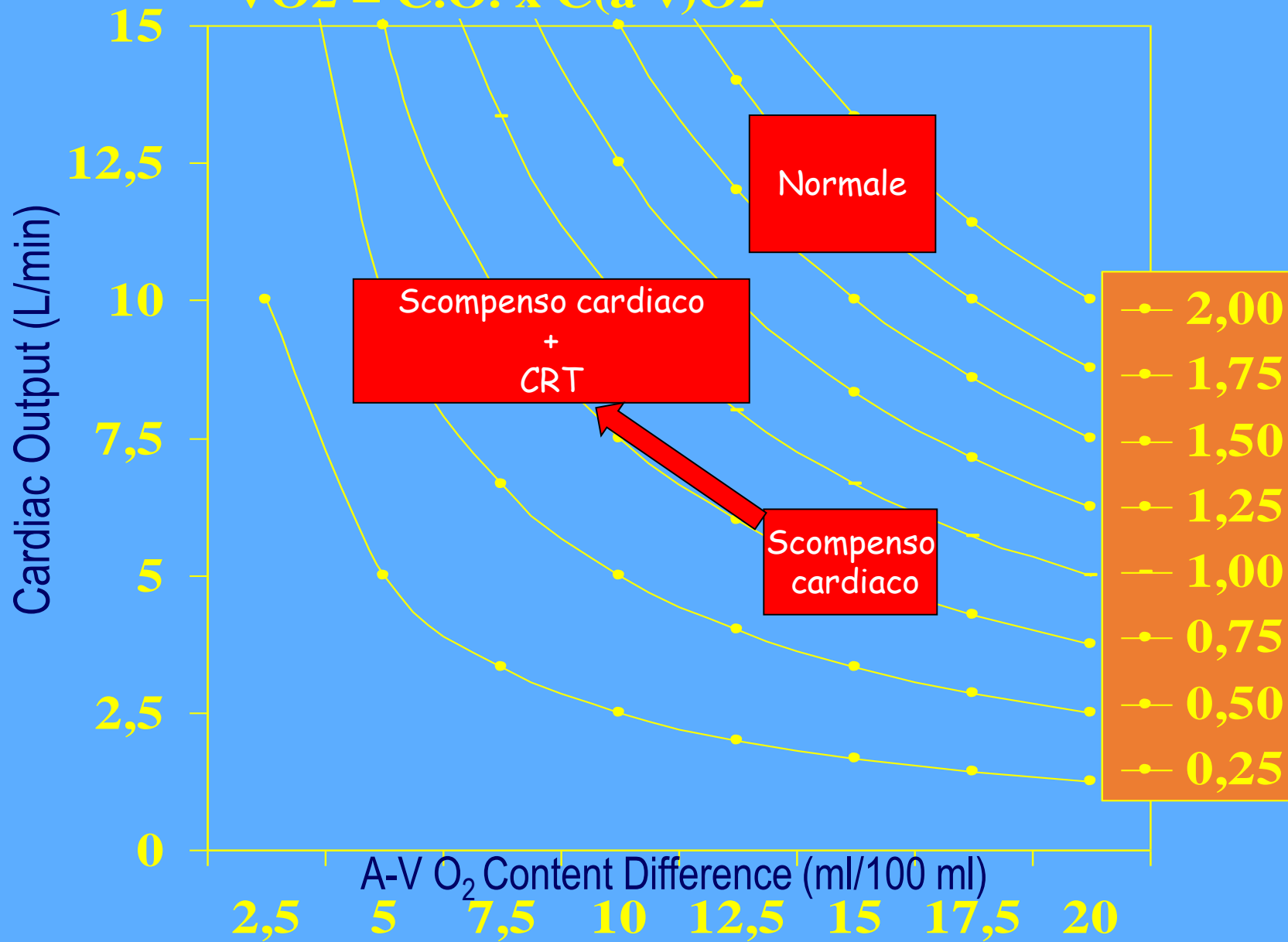
Brescia, Italy

- OBJECTIVES** We studied whether direct assessment of the hemodynamic response to exercise could improve the prognostic evaluation of patients with heart failure (HF) and identify those in whom the main cause of the reduced functional capacity is related to extracardiac factors.
- BACKGROUND** Peak exercise oxygen consumption (VO_2) is one of the main prognostic variables in patients with HF, but it is influenced also by many extracardiac factors.
- METHODS** Bicycle cardiopulmonary exercise testing with hemodynamic monitoring was performed, in addition to clinical evaluation and radionuclide ventriculography, in 219 consecutive patients with chronic HF (left ventricular ejection fraction, $22 \pm 7\%$; peak VO_2 , 14.2 ± 4.4 ml/kg/min).
- RESULTS** During a follow-up of 19 ± 25 months, 32 patients died and 6 underwent urgent transplantation with a 71% cumulative major event-free 2-year survival. Peak exercise stroke work index (SWI) was the most powerful prognostic variable selected by Cox multivariate analysis, followed by serum sodium and left ventricular ejection fraction, for one-year survival, and peak VO_2 and serum sodium for two-year survival. Two-year survival was 54% in the patients with peak exercise SWI ≤ 30 gm/m² versus 91% in those with a SWI >30 gm/m² ($p < 0.0001$). A significant percentage of patients (41%) had a normal cardiac output response to exercise with an excellent two-year survival (87% vs. 58% in the others) despite a relatively low peak VO_2 (15.1 ± 4.7 ml/kg/min).
- CONCLUSIONS** Direct assessment of exercise hemodynamics in patients with HF provides additive independent prognostic information, compared to traditional noninvasive data. (J Am Coll Cardiol 1999;33:943-50) © 1999 by the American College of Cardiology





$$VO_2 = C.O. \times C(a-v)O_2$$



Hemodynamic Effects of Exercise Training in Heart Failure

GAIA CATTADORI, MD,¹ JEAN-PAUL SCHMID, MD,² NICOLAS BRUGGER, MD,² ERICA GONDONI, MD,¹
PIETRO PALERMO, MD,¹ AND PIERGIUSEPPE AGOSTONI, MD, PhD^{1,3,4}

Milan, Italy; Bern, Switzerland; and Seattle, Washington

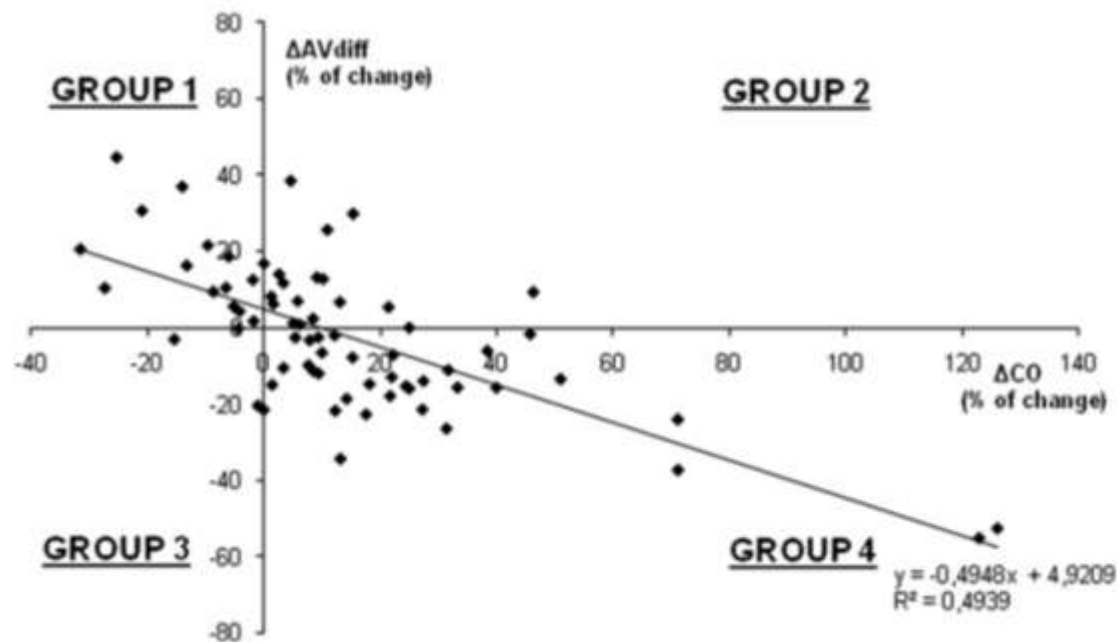
ABSTRACT

Background: Exercise performance improvement after training in heart failure (HF) can be due to central or peripheral changes.

Methods and Results: In 70 HF stable patients we measured peak VO_2 and cardiac output (CO, inert gas rebreathing technique) and calculated arteriovenous O_2 differences (a-v O_2 -diff) before and after an 8-week training program. Peak VO_2 changed from 1111 ± 403 mL/minute to 1191 ± 441 ($P < .001$), peak workload from 68 ± 29 watts to 76 ± 32 ($P < .0001$), peak CO from 6.6 ± 2.2 L/minute to 7.3 ± 2.5 ($P < .0001$), and peak a-v O_2 -diff from 17.5 ± 5.1 mL/100 mL to 16.6 ± 4.1 ($P = .081$). Changes in peak CO and a-v O_2 -diff allowed to identify 4 behaviors: group 1: ($n = 15$) reduction in peak CO and increase in a-v O_2 -diff (peak VO_2 unchanged, peak workload +9.5%); group 2: ($n = 16$) both peak CO and a-v O_2 -diff increased as well as peak VO_2 (23%) and workload (18%); group 3: ($n = 4$) peak CO and a-v O_2 -diff reduced as well as peak VO_2 (-18%) and workload (-5%); group 4: ($n = 35$) peak CO increased with a-v O_2 -diff reduced (increase in peak VO_2 by 5.5 and workload by 8.4%).

Conclusions: Exercise training improves peak VO_2 by increasing CO with unchanged a-v O_2 -diff. A reduction after training of a-v O_2 -diff with an increase in CO is frequent (50% of cases), is suggestive of blood flow redistribution and, per se, not a sign of reduced muscle performance been associated with improved exercise capacity. (*J Cardiac Fail* 2011;17:916–922)

Key Words: Training, heart failure, cardiac output.



ESC HEART FAILURE

ESC Heart Failure 2021; **8**: 4915–4924

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ORIGINAL ARTICLE

Rest and exercise oxygen uptake and cardiac output changes 6 months after successful transcatheter mitral valve repair

Carlo Vignati^{1,2}, Fabiana De Martino¹, Manuela Muratori¹, Elisabetta Salvioni¹, Gloria Tamborini¹, Antonio Bartorelli^{1,3}, Mauro Pepi¹, Francesco Alamanni^{1,2}, Stefania Farina¹, Gaia Cattadori⁴, Valentina Mantegazza¹ and Piergiuseppe Agostoni^{1,2*}

¹Centro Cardiologico Manzino, IRCCS, Milan, Italy; ²Department of Clinical Sciences and Community Health, Cardiovascular Section, University of Milan, Milan, Italy;

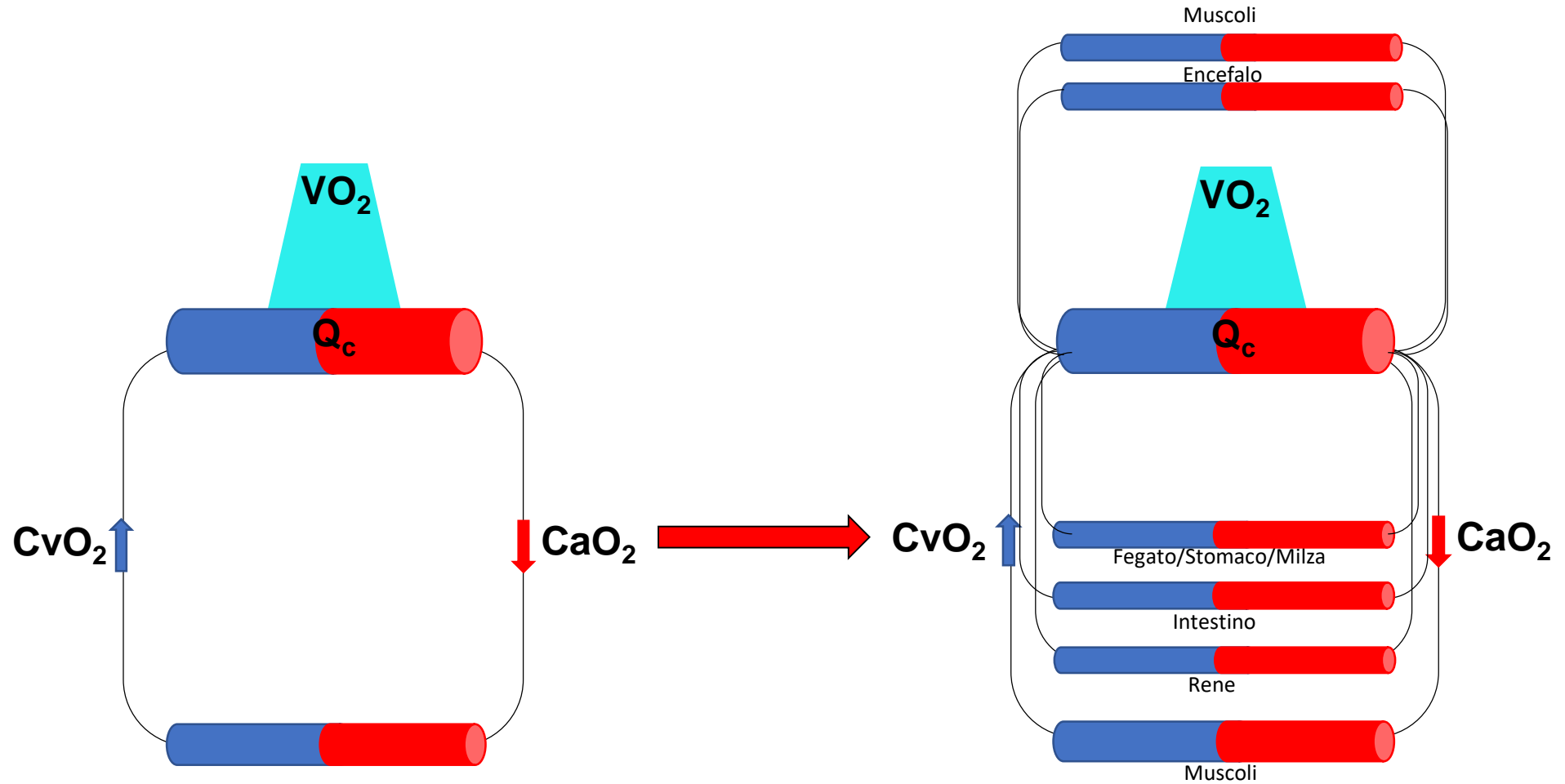
³Department of Biomedical and Clinical Sciences "Luigi Sacco", University of Milan, Milan, Italy; and ⁴IRCCS Multimedica, Milan, Italy



LEGGE DI FICK

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DISTRIBUZIONE



...grazie per l'attenzione.



Quanto conta l'anemia?

Table 2-1. Concepts and Calculations Pertaining to Oxygen Utilization, Content, Transport, and Extraction

$\frac{O_2 \text{ utilization}}{250 \text{ ml/min}}$	$= \text{Cardiac output} \cdot (\text{arterial } O_2 \text{ content} - \text{venous } O_2 \text{ content})$ $= 5000 \text{ ml/min} \cdot (19 \text{ ml/dl} - 14 \text{ ml/dl})$
$\frac{\text{Arterial } O_2 \text{ content}}{19 \text{ ml/dl}}$	$= \text{Hemoglobin} \cdot \% \text{ saturation} \cdot O_2 \text{ combining capacity}$ $= 14 \text{ gm/dl} \cdot 0.96 \cdot 1.34 \text{ ml/gm}$
$\frac{\text{Venous } O_2 \text{ content}}{14 \text{ ml/dl}}$	$= 14 \text{ gm/dl} \cdot 0.75 \cdot 1.34 \text{ ml/gm}$
$\frac{\text{Arteriovenous } O_2 \text{ difference}}{5 \text{ ml/dl}}$	$= \text{Arterial } O_2 \text{ content} - \text{venous } O_2 \text{ content}$ $= 19 \text{ ml/dl} - 14 \text{ ml/dl}$
$\frac{O_2 \text{ transport}}{950 \text{ ml/min}}$	$= \text{Cardiac output} \cdot \text{arterial } O_2 \text{ content}$ $= 5000 \text{ ml/min} \cdot 19 \text{ ml/dl}$
$\frac{O_2 \text{ extraction}}{25\%}$	$= \frac{\text{Arteriovenous } O_2 \text{ difference}}{\text{Arterial } O_2 \text{ content}} \cdot 100\%$ $= \frac{19 - 14}{19} \cdot 100\%$

$$\underline{CaO_2 = 1.34 \text{ ml} \times \text{gr Hb}}$$

$$\underline{\text{Estrazione periferica } O_2 = 75\%}$$

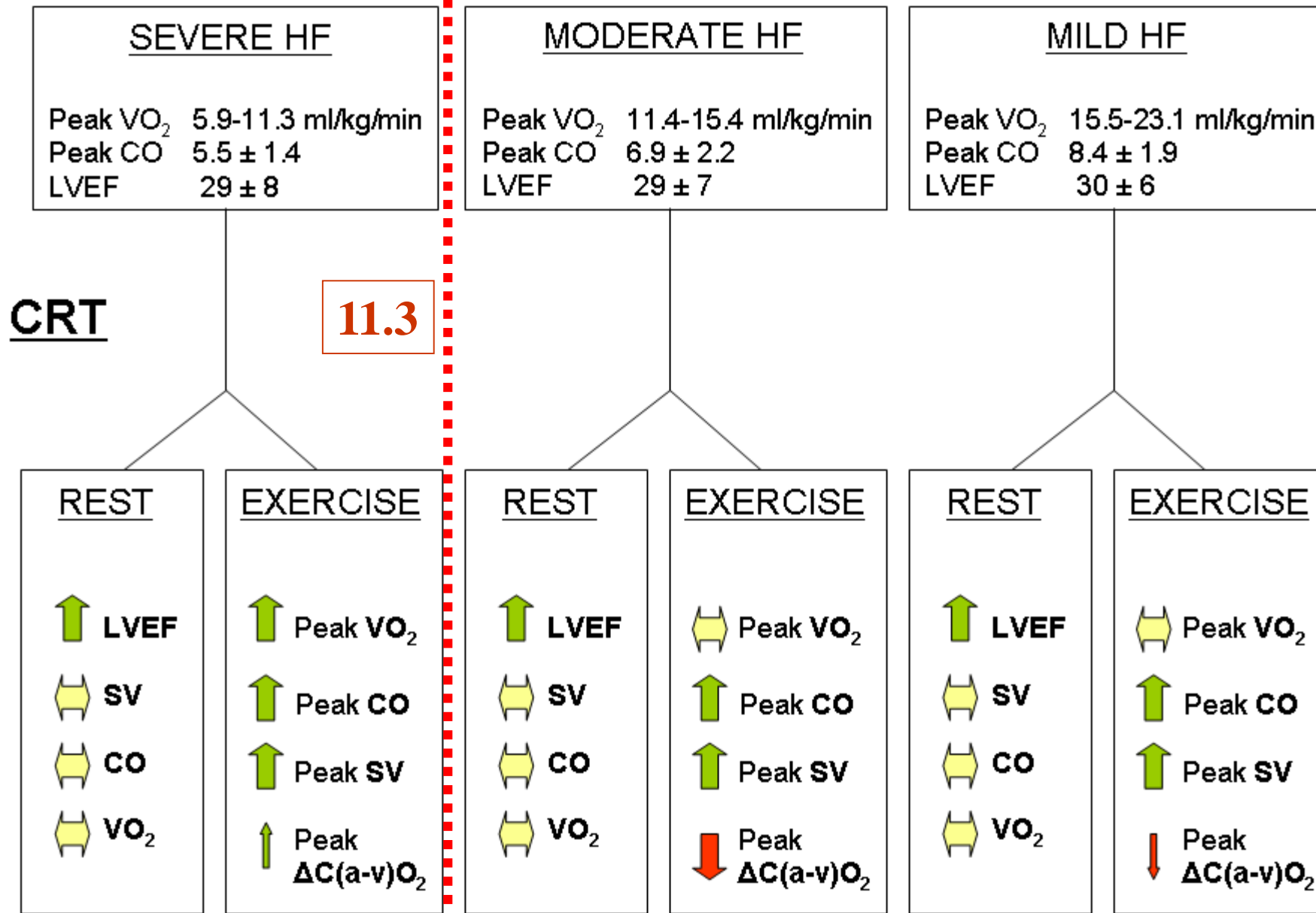
1 gr di Hb porta ai tessuti 1 ml O_2 x dl di sangue



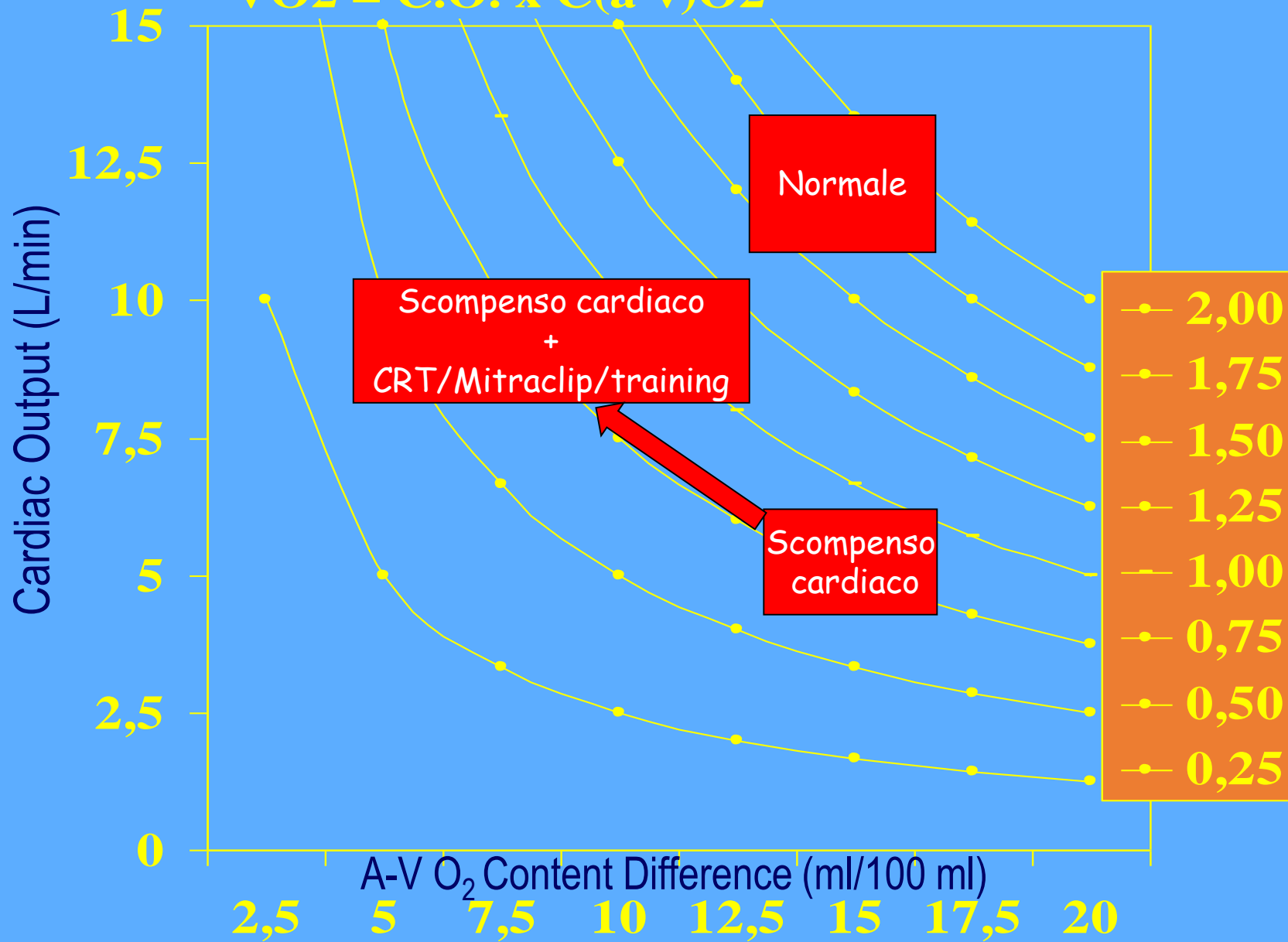
Research Article

**Peak Exercise Cardiac Output but Not Oxygen Uptake Increases in
All Heart Failure Patients After Successful Resynchronization
Therapy**

**Gaia Cattadori^{1#}, Carlo Vignati^{2,5#}, Alice Bonomi², Massimo Mapelli^{3,5}, Susanna Sciomer³,
Mauro Pepi², Claudio Tondo², Giuseppe Ambrosio⁴, Silvia Di Marco¹, Massimo Baravelli¹,
Piergiuseppe Agostoni^{2,5*}**



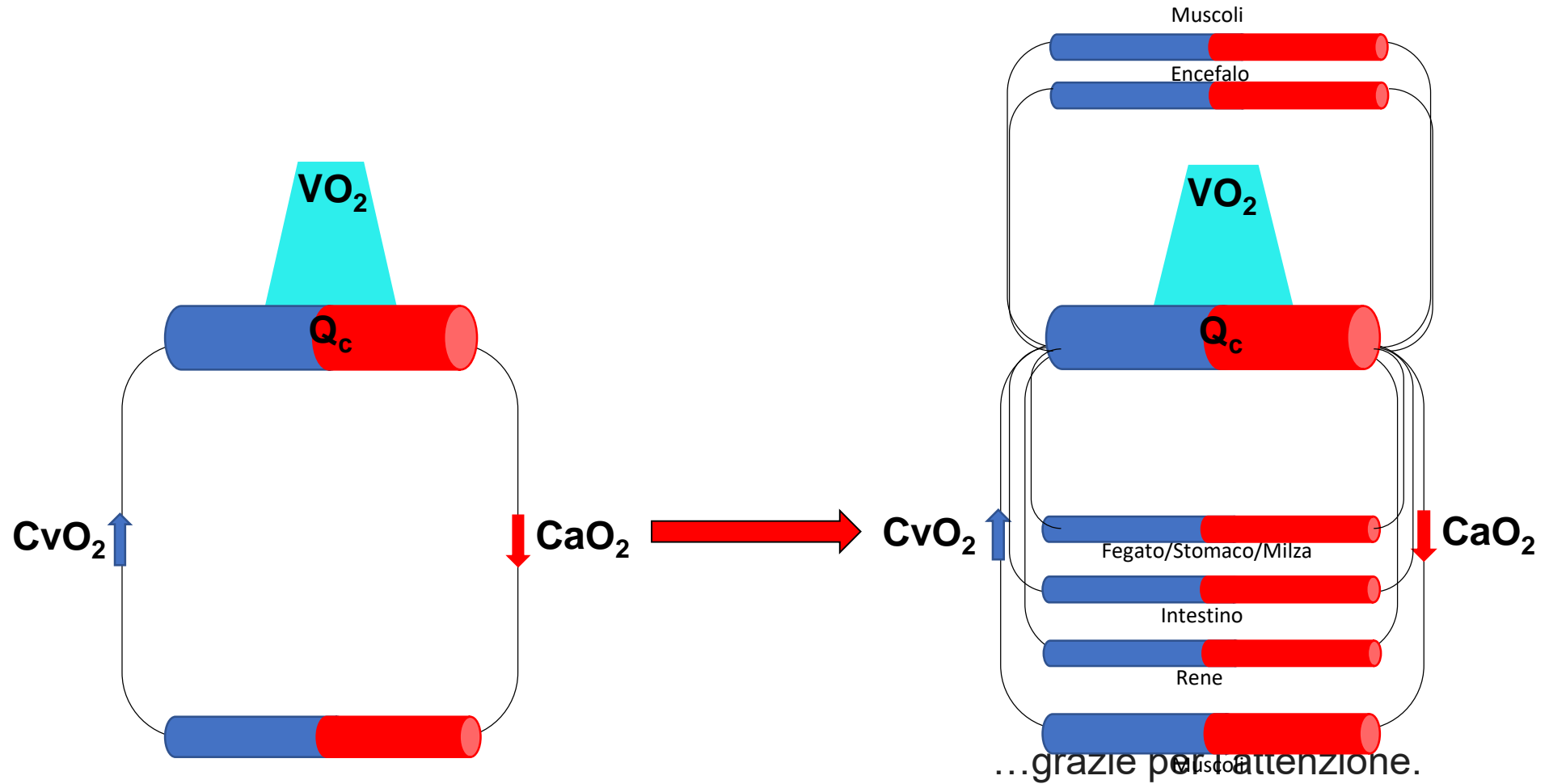
$$VO_2 = C.O. \times C(a-v)O_2$$



LEGGE DI FICK

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DISTRIBUZIONE



...grazie per l'attenzione.

Step	CO L/min	HR b/min	SV ml/b	VO2 ml/min	C(a-v)O ₂ ml/dl
Basal	4.5	75	60	220	5
Peak	7.9	109	72	1080	14
Peak pred	13.6	154	88	2120	16
Peak %	58	71	82	51	88

Hb 13.5 gr/dl

- 1) Limitazione Cardiogenica (mlVO2)
- 2) Limitazione "Periferica" (mlVO2)

Step	CO L/min	HR b/min	SV ml/b	VO2 ml/min	C(a-v)O ₂ ml/dl
Basal	4.5	75	60	220	5
Peak	7.9	109	72	1080	14
Peak pred	13.6	154	88	2120	16
Peak %	58	71	82	51	88

Hb 13.5 gr/dl

1040 ml/min

- 1) Limitazione Cardiogenica (mlVO2)
- 2) Limitazione "Periferica" (mlVO2)

Step	CO L/min	HR b/min	SV ml/b	VO2 ml/min	C(a-v)O ₂ ml/dl
Basal	4.5	75	60	220	5
Peak	7.9	109	72	1080	14
Peak pred	13.6	154	88	2120	16
Peak %	58	71	82	51	88

Hb 13.5 gr/dl

919 ml/min

1040 ml/min

- 1) Limitazione Cardiogenica (mlVO2)
- 2) Limitazione "Periferica" (mlVO2)

Step	CO L/min	HR b/min	SV ml/b	VO2 ml/min	C(a-v)O ₂ ml/dl
Basal	4.5	75	60	220	5
Peak	7.9	109	72	1080	14
Peak pred	13.6	154	88	2120	16
Peak %	58	71	82	51	88

Hb 13.5 gr/dl

919 ml/min ← 74% → 1040 ml/min

- 1) Limitazione Cardiogenica (mlVO2) → 74%
- 2) Limitazione "Periferica" (mlVO2)

Step	CO L/min	HR b/min	SV ml/b	VO2 ml/min	C(a-v)O ₂ ml/dl
Basal	4.5	75	60	220	5
Peak	7.9	109	72	1080	14
Peak pred	13.6	154	88	2120	16
Peak %	58	71	82	51	88

919 ml/min

←

74%

1040 ml/min

→

26%

121 ml/min

Hb 13.5 gr/dl

- Hb
 - SatHb O₂
 - pO₂

Step	CO L/min	HR b/min	SV ml/b	VO2 ml/min	Diff AV
Basal	4.5	75	60	220	5
Peak	7.9	109	72	1080	14
Peak pred	13.6	154	88	2120	16
Peak %	58	71	82	51	88

Hb 13.5 gr/dl

1 gr Hb porta ai tessuti 1 ml O₂ x dl di sangue

2) Limitazione (mlVO2) dovuta all'anemia = CO x anemia = CO x (15-13.5)

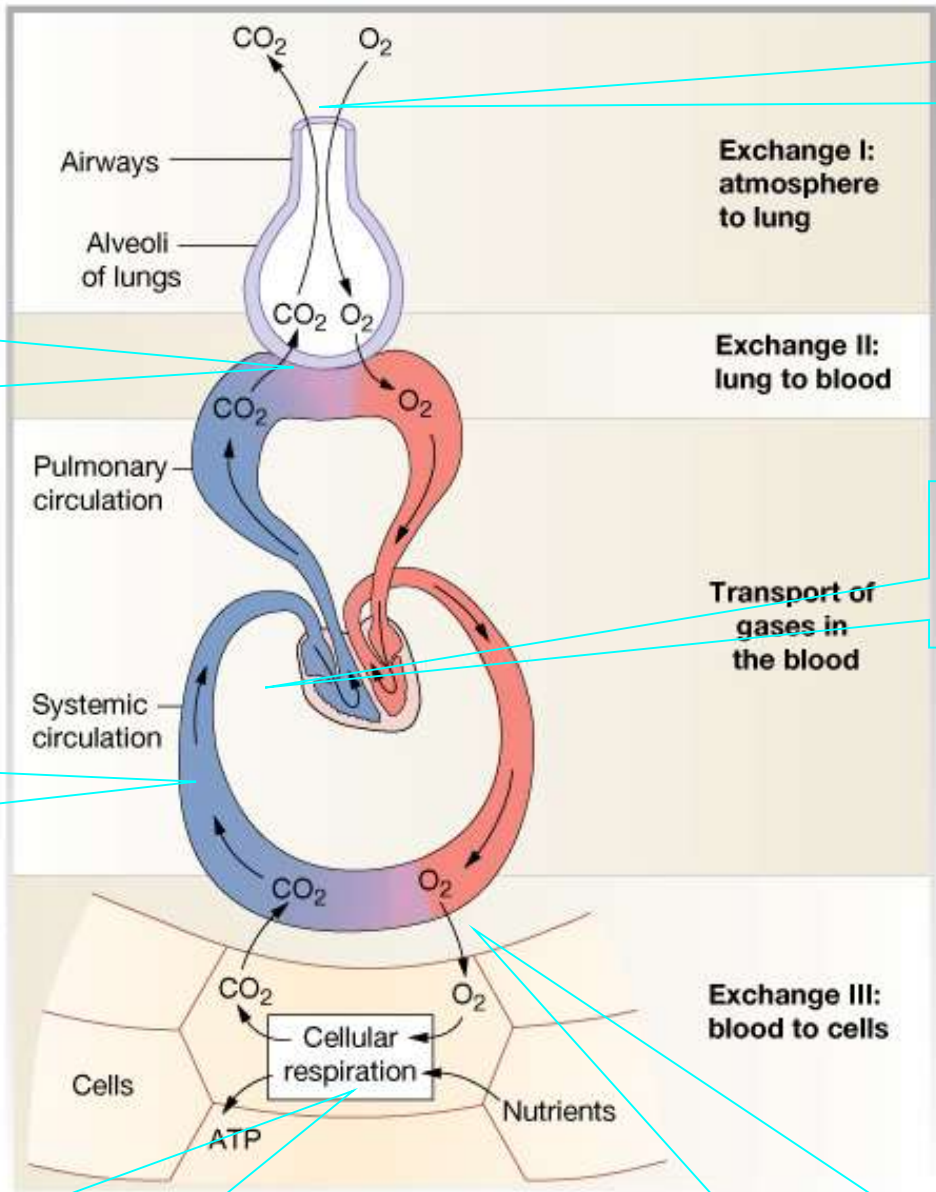
$$79 \times 1.5 = 119 \text{ mlVO2}$$

VO_2

Diffusione O_2

Trasporto O_2

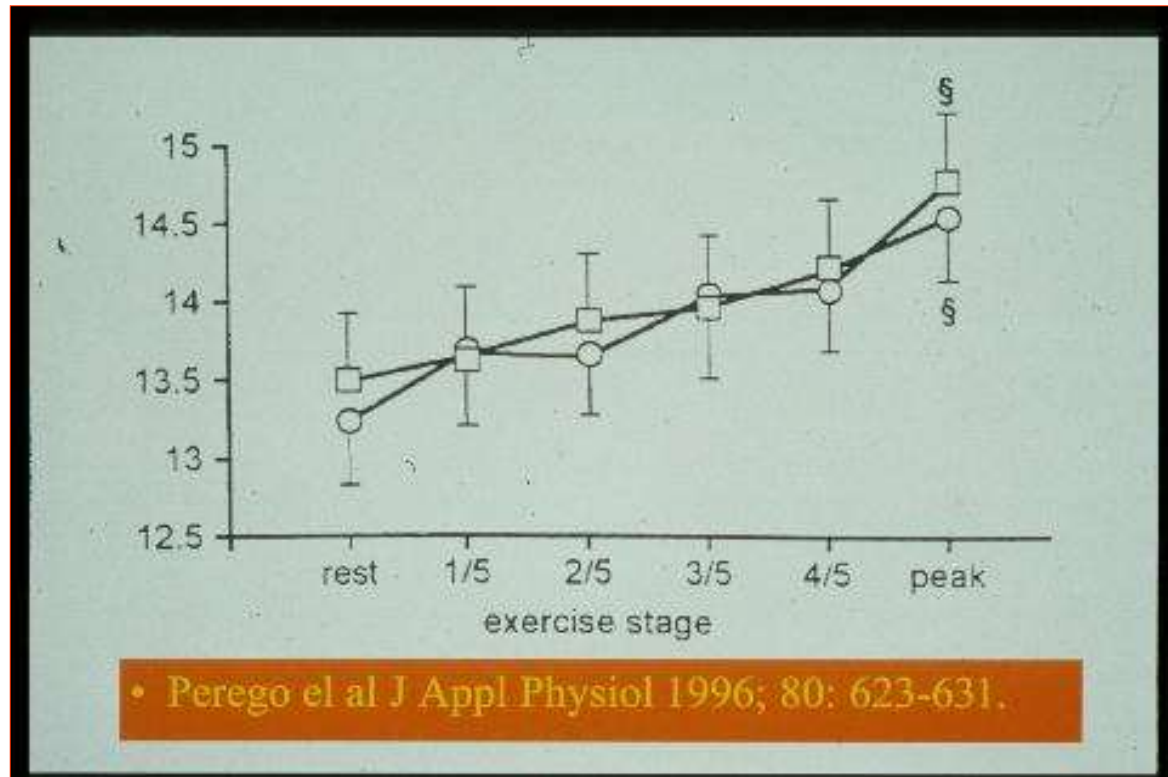
Utilizzo muscolare O_2



Ventilazione

Gettata Cardiaca

Estrazione periferica O_2



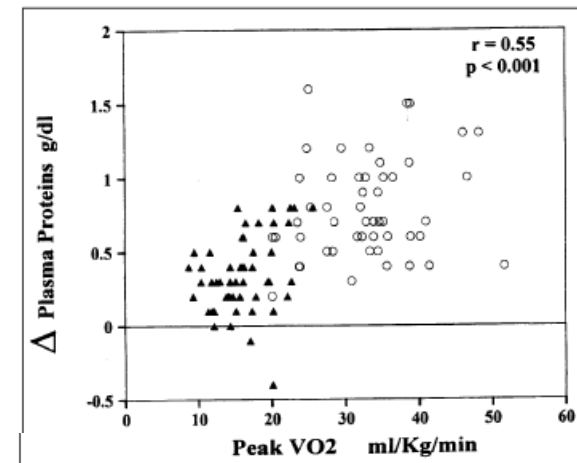
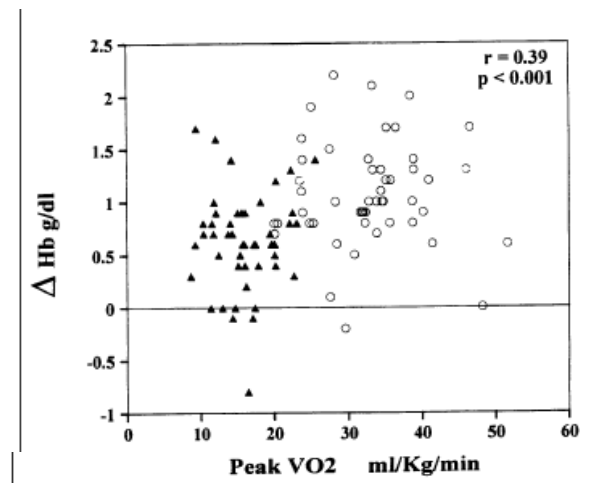
Due possibili meccanismi per incremento Hb:

1. Riduzione plasma
2. Incremento globuli rossi

1. Riduzione plasma

Exercise-Induced Hemoconcentration in Heart Failure Due to Dilated Cardiomyopathy

Piergiuseppe Agostoni, MD, Karlman Wasserman, MD, Marco Guazzi, MD,
Gaia Cattadori, MD, Pietro Palermo, MD, Giancarlo Marenzi, MD, and
Maurizio D. Guazzi, MD



2. Incremento globuli rossi

bjh research paper

Exercise capacity in patients with β -thalassaemia intermedia

Piergiuseppe Agostoni^{1,2} Mario Cerino,³
Pietro Palermo,¹ Alessandra Magini,¹
Michele Bianchi,¹ Maurizio Bussotti,¹
Gemino Fiorelli³ and Maria D.
Cappellini³

¹Centro Cardiologico Monzino, IRCCS, Istituto di
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²Division of Respiratory and Critical Care
Medicine, Department of Medicine, University of
Washington, Seattle, WA, USA, and ³Centro
Anemie Congenite, Ospedale Maggiore, IRCCS,
Università di Milano, Milan, Italy

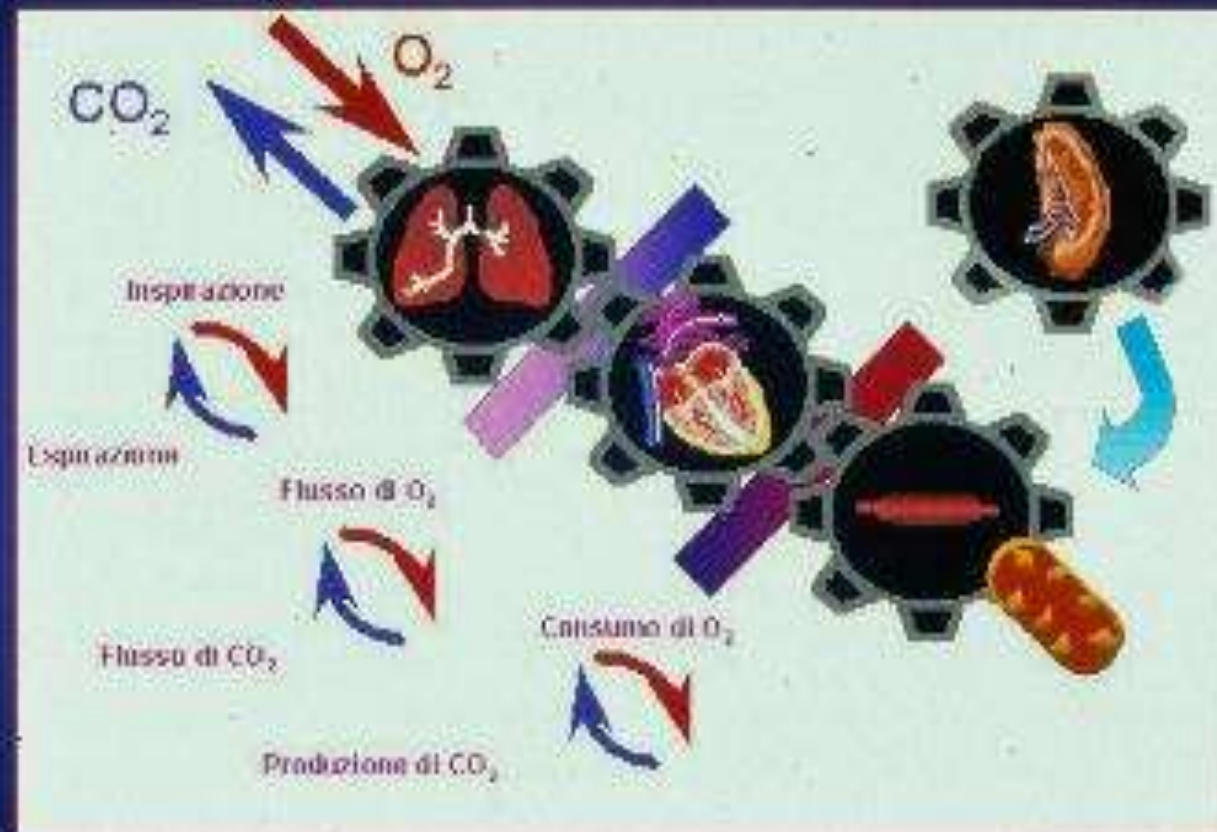
Received 2 June 2005; accepted for publication
9 August 2005

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Summary

Thalassaemia intermedia patients can suffer fatigue and exercise capacity reduction, possibly because of anaemia, deconditioning and lack of exercise-induced haemoconcentration. We studied 21 β -thalassaemia intermedia patients, 10 splenectomised (group A) and 11 not splenectomised (group B). Patients were evaluated by cardiopulmonary exercise test with blood sampling for haemoglobin and plasma protein measurements at rest and peak. During exercise, an isolated increase of haemoglobin suggested spleen contraction while a parallel increase of haemoglobin and proteins suggested fluid filtration through capillary wall. Groups were homogeneous for age and gender. Peak oxygen consumption (VO_2) was 22.5 ± 4.4 ml/min/kg ($51 \pm 14\%$) and 24.3 ± 7.0 ($53 \pm 12\%$) in groups A and B respectively [not significant (NS)]. At rest, haemoglobin was 8.8 g/dl in both groups. Exercise-induced increment was 0.4 ± 0.2 and 1.0 ± 0.4 g/dl ($P < 0.001$) for haemoglobin and 4.0 ± 3.0 and 5.0 ± 4.0 g/l (NS) for proteins, in groups A and B respectively. Anaemia was the major cause of peak VO_2 reduction (1097 ± 260 ml/min). However, anaemia did not explain the entire exercise capacity reduction, suggesting the presence of muscular deconditioning. Exercise capacity is reduced in β -thalassaemia intermedia because of anaemia and muscular deconditioning. Spleen contraction does not significantly influence exercise capacity although exercise-induced haemoconcentration was greater in patients with spleen.

Keywords: anaemia, haemoglobin, oxygen consumption, muscular deconditioning, spleen contraction.



Ex 2) Rehabilitation Patients



CARDIOTHORACIC ANESTHESIOLOGY:

The *Annals of Thoracic Surgery* CME Program is located online at <http://cme.ctsnetjournals.org>. To take the CME activity related to this article, you must have either an STS member or an individual non-member subscription to the journal.

Postoperative Anemia and Exercise Tolerance After Cardiac Operations in Patients Without Transfusion: What Hemoglobin Level Is Acceptable?

Marco Ranucci, MD, Maria Teresa La Rovere, MD, Serenella Castelvecchio, MD, Roberto Maestri, MS, Lorenzo Menicanti, MD, Alessandro Frigiola, MD, Andrea Maria D'Armini, MD, Claudio Goggi, MD, Roberto Tramarin, MD, and Oreste Febo, MD

Department of Cardiothoracic and Vascular Anesthesia, ICU & Cardiac Surgery, IRCCS Policlinico San Donato, Milan; Departments of Cardiology and Biomedical Engineering, Fondazione Salvatore Maugeri, IRCCS Istituto Scientifico di Montescano, Montescano; Division of Cardiac Surgery, IRCCS Fondazione Policlinico San Matteo, Pavia; and Cardiac Rehabilitation Unit, Fondazione Europea per la Ricerca Biomedica, Cernusco S/N, Milan, Italy

Background. Restrictive transfusion strategies have been suggested for cardiac surgical patients, leading to various degrees of postoperative anemia. This study investigates the exercise tolerance during rehabilitation of cardiac surgical patients who did not receive transfusions, with respect to their level of postoperative anemia.

Methods. This observational study started in January 2010 and ended in May 2010 in 2 rehabilitation hospitals and 2 large-volume cardiac surgical hospitals. The study population was 172 patients who did not receive transfusions during cardiac surgical operations with cardiopulmonary bypass and subsequently followed a rehabilitation program in 1 of the 2 rehabilitation hospitals. No patient received a transfusion during the rehabilitation hospital stay. Exercise tolerance was measured using the 6-minute walk test at admission and discharge from the rehabilitation hospital. The level of anemia at admission

to the rehabilitation hospital was tested as an independent predictor of exercise tolerance within a model inclusive of other possible confounders.

Results. Patients with values of hemoglobin less than 10 g/dL at admission to the rehabilitation institute had a significantly ($p = 0.007$) worse performance on the 6-minute walk test than patients with higher values (258 ± 106 vs 306 ± 101 meters). This functional gap was completely recovered during a normal rehabilitation period. Other independent factors affecting exercise tolerance were age, sex, and albumin concentration.

Conclusions. Postoperative anemia with hemoglobin levels of 8 to 10 g/dL is well tolerated in patients who have not received a transfusion and induces only a transient impairment of exercise tolerance.

(*Ann Thorac Surg* 2011;92:25–31)

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Ex 2)

Rehabilitation Patients

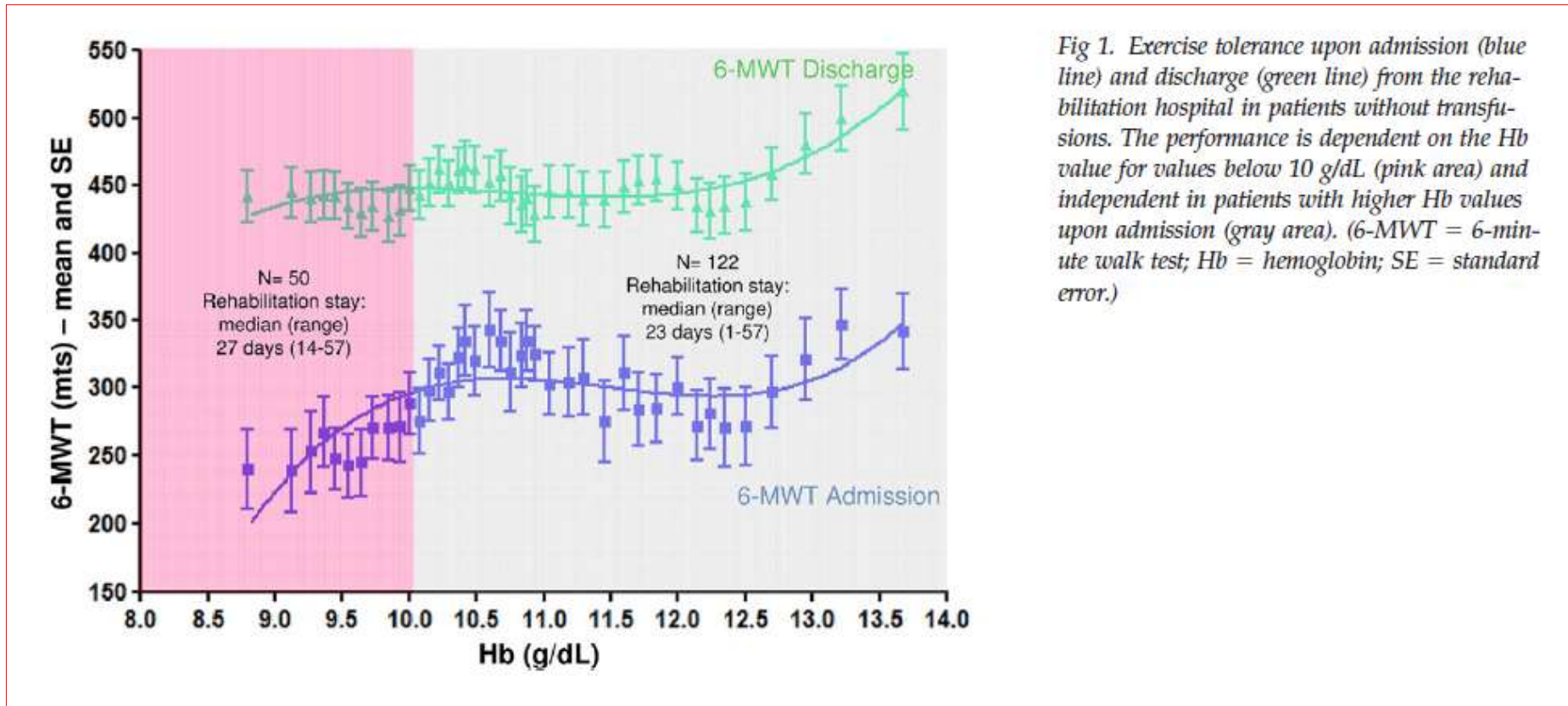
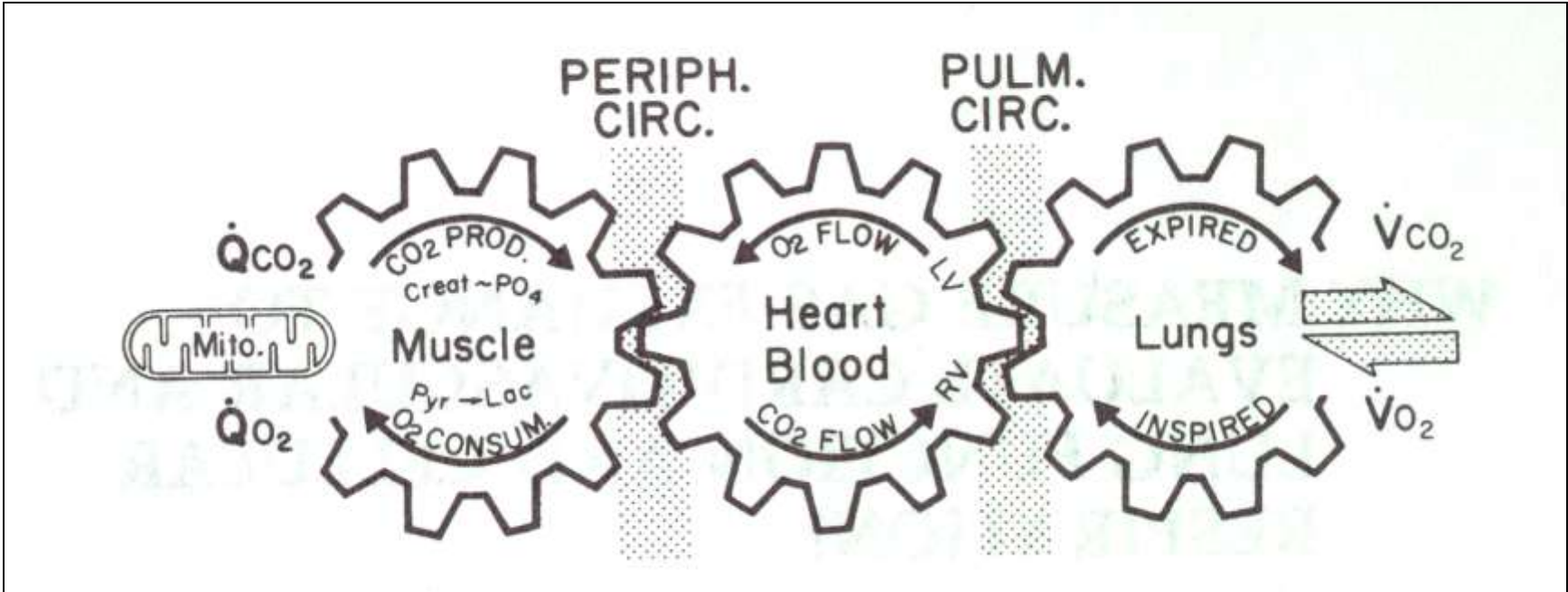


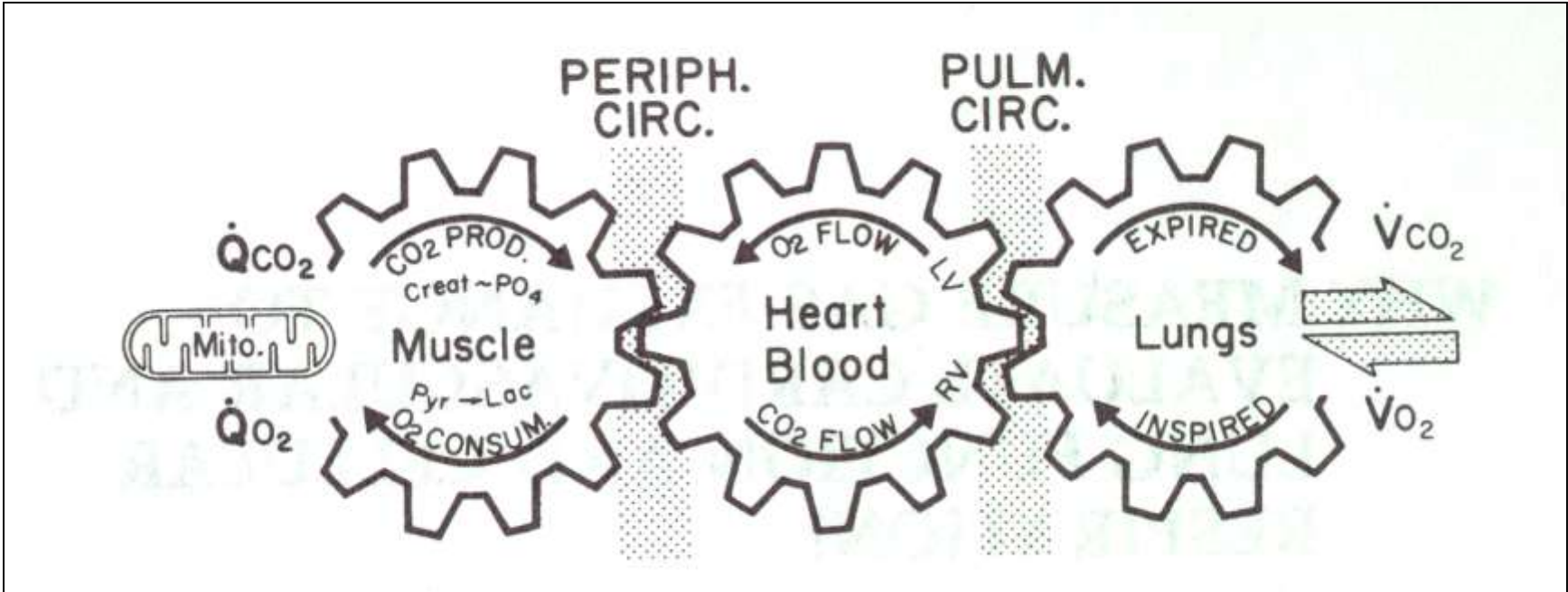
Fig 1. Exercise tolerance upon admission (blue line) and discharge (green line) from the rehabilitation hospital in patients without transfusions. The performance is dependent on the Hb value for values below 10 g/dL (pink area) and independent in patients with higher Hb values upon admission (gray area). (6-MWT = 6-minute walk test; Hb = hemoglobin; SE = standard error.)

Ranucci M. et al. Ann Thorac Surg 2011; 92:25-31

$\dot{V}O_2$

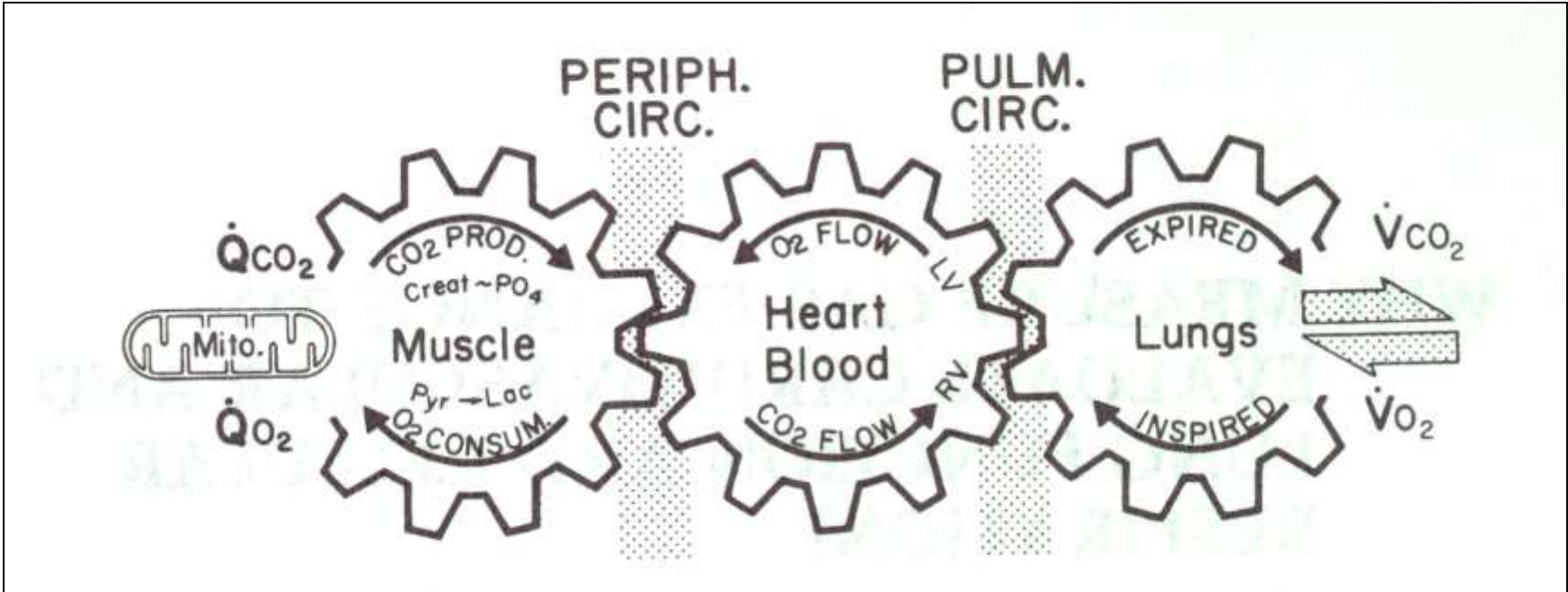


VO₂



$$VO_2 = VE (FiO_2 - FeO_2)$$

VO₂

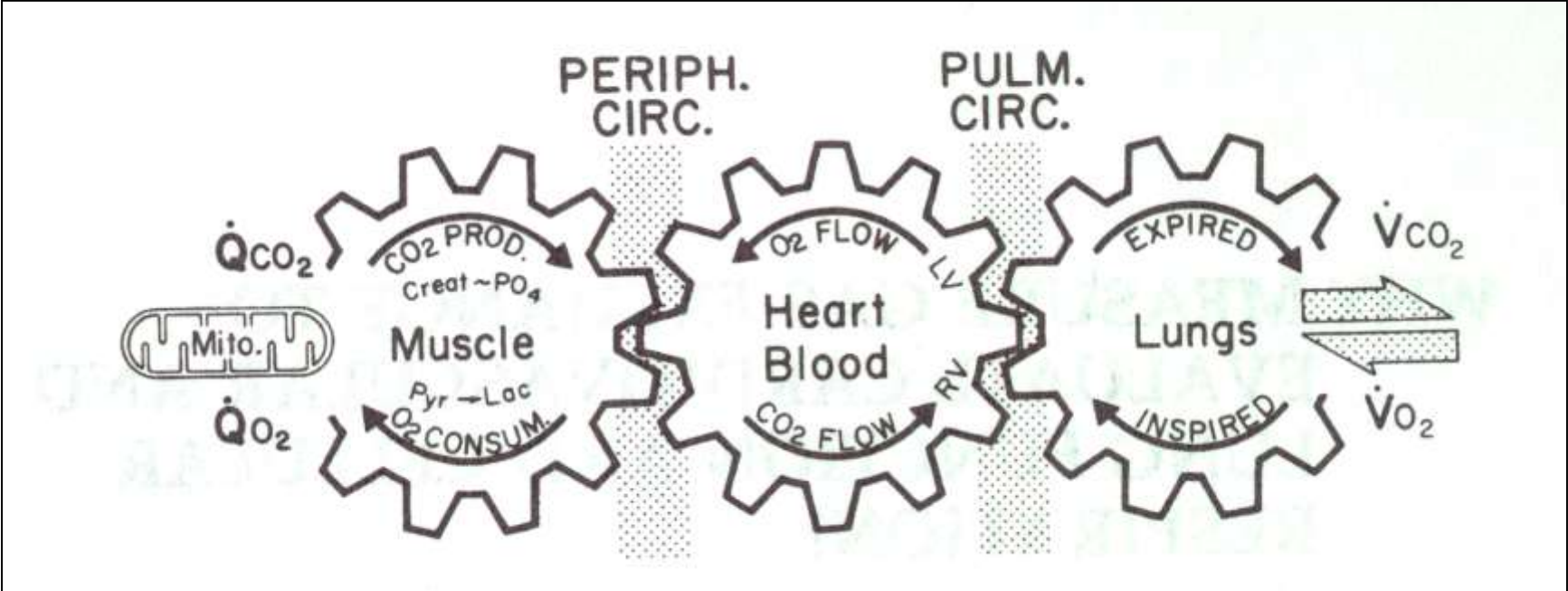


$$VO_2 = D(p_{capO_2} - p_{mitO_2})$$

$$VO_2 = VE (F_{iO_2} - F_{eO_2})$$

VO₂

$$VO_2 = CO \times C(a-v)O_2$$



$$VO_2 = D(pcapO_2 - pmitO_2)$$

$$VO_2 = VE (FiO_2 - FeO_2)$$

