## Critical evaluation of Priorities and therapeutic outcome measures

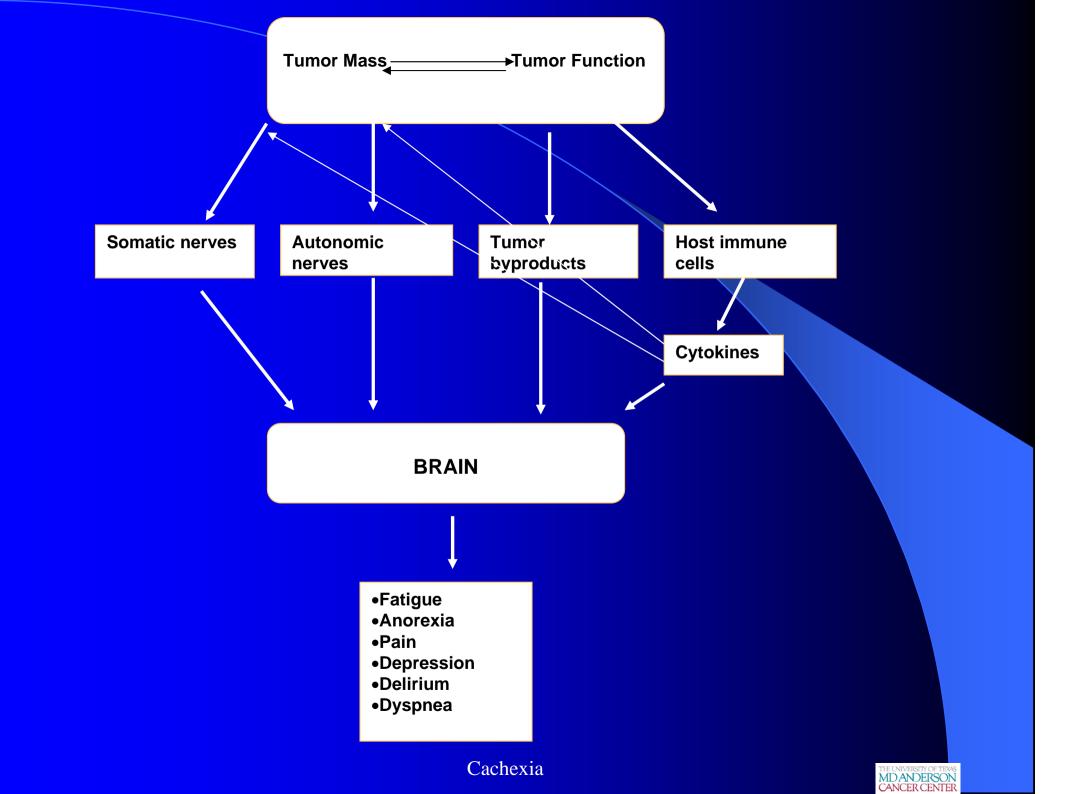
Eduardo Bruera, Houston - USA



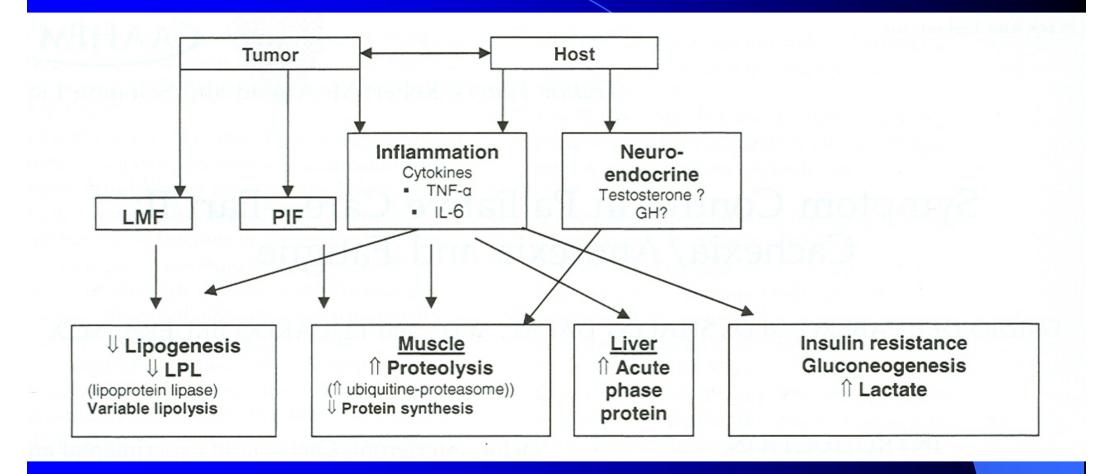
#### **Patient's Problems**

- Pain (80%)
- Fatigue (90%)
- Weight Loss (80%)
- Lack of Appetite (80%)
- Nausea, Vomiting (90%)
- Anxiety (25%)
- Shortness of Breath (50%)
- Confusion-Agitation (80%)





## Why does cachexia happen?



#### Clinical effects of cachexia

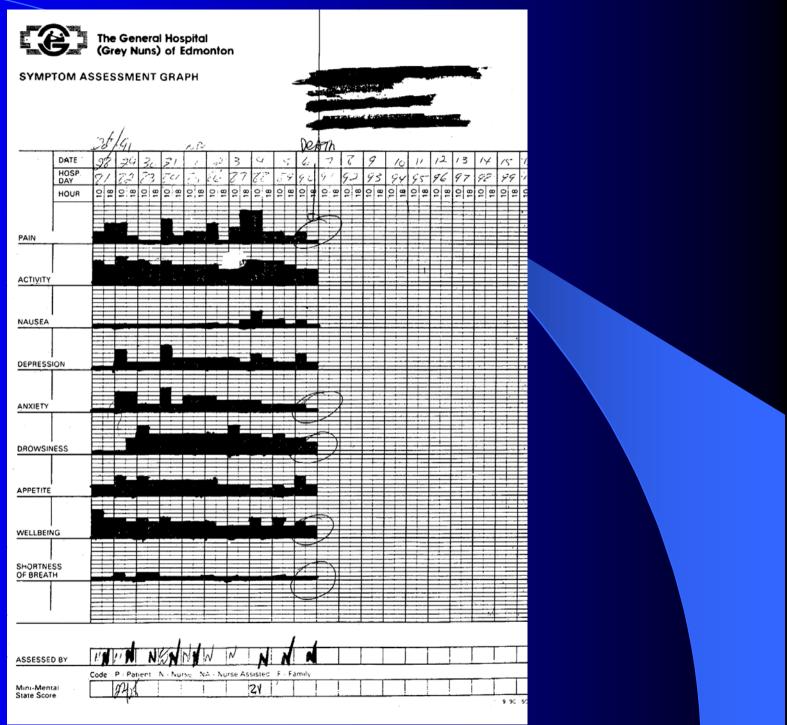
- Symptoms: anorexia, fatigue, nausea
- Psychological distress Pt/family
- Body composition
- Physical function
- Survival



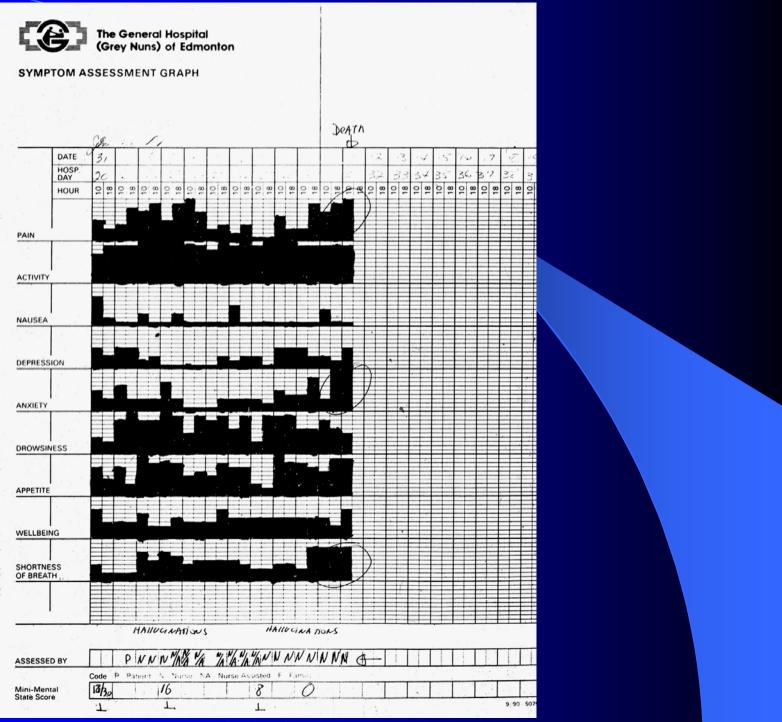
# Symptoms/ distress Pt & family

- Global: ESAS, Q of L scales
- Specific: Fatigue (FACIT-F), mood, nausea, etc
- Family distress tools





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## **Body composition**

- Present/ usual weight (5%, 10%)
- BIE
- Anthropometrics
- Percentual caloric intake
- Albumin, pre-albumin
- Caloric intake ( % intake, diaries)
- Imaging? CT scan MRI muscle/ fat



### Physical function

- Subjective description of performance
- Subjective status upon performance
- Tests: 6 min walk, stand up and go,
  Simmonds, actigraph, pedometer
- O2 consumption, heart rate



## Anorexia/Cachexia Decision-Making

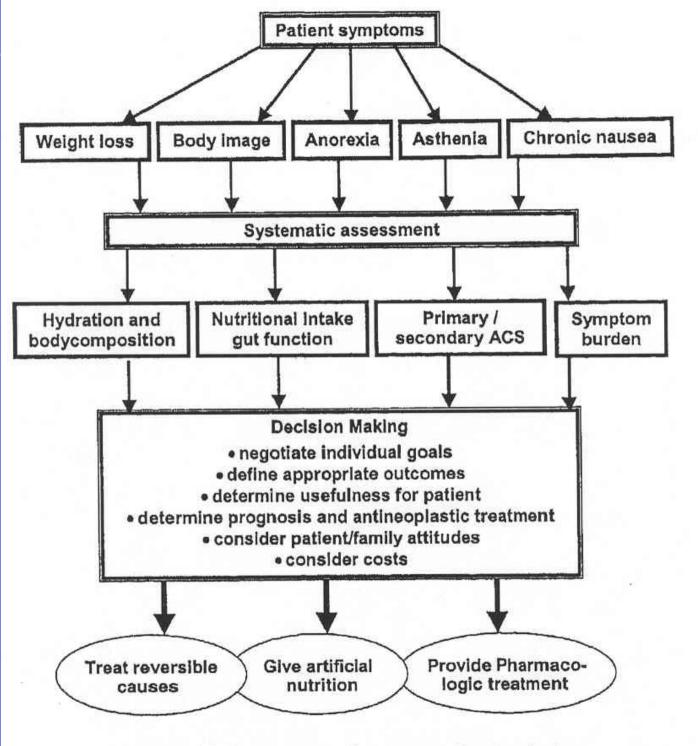


Fig. 3. Clinical management of cancer anorexia and cachexia.

#### Interventions

- Treat primary cachexia
- Nutrition
- Exercise
- Treat secondary cachexia (nausea, taste, dysphagia)
- Treat associated symptoms ( depression, pain)



## Treat primary cachexia

- Tumor byproducts (targeted therapies, hormones)
- Proteasome inhibitors (bortezomib, others)
- Cytokine block (thalidomide, monoclonal Abs), anti-inflammatory
- Target organs (appetite, muscle)



#### Nutrition

- Oral Vs parenteral/ enteral
- Energy Vs protein
- Differential nutrition (PUFAs, branched chain AAs, arginine)- what is really lacking?
- Stimulate intake: ghrelin, pro-motility
- Treat secondary cachexia: dysphagia, depression, meal preparation

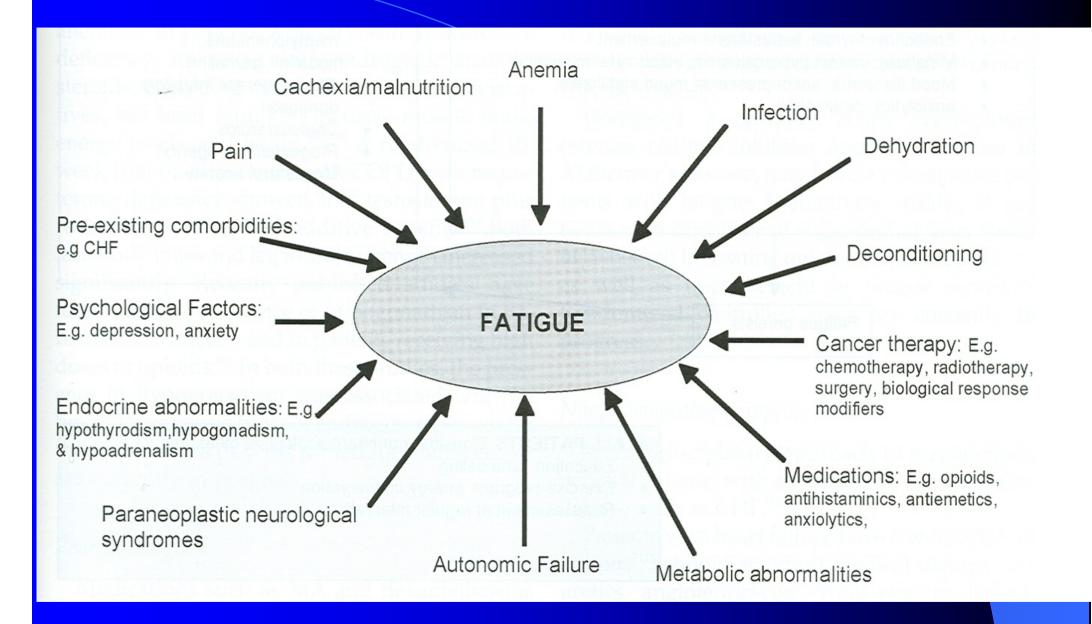


#### Exercise

- ADLs as exercise
- Increased walking (pedometer, actigraph)
- Upper limb exercise
- Individualized plan
- Treat fatigue!!
- Add testosterone/ steroids?

**Essential for increased LBM** 





#### Fatigue 8/10

#### Patient 1

Patient 2

**Depression** 

**60%** 

10%

Cachexia

**10%** 

50%

Anemia

**10%** 

• 30%

**Opioids** 

**20%** 

0%

**Autonomic** 

0%

10%



## Treat secondary cachexia

Starvation component



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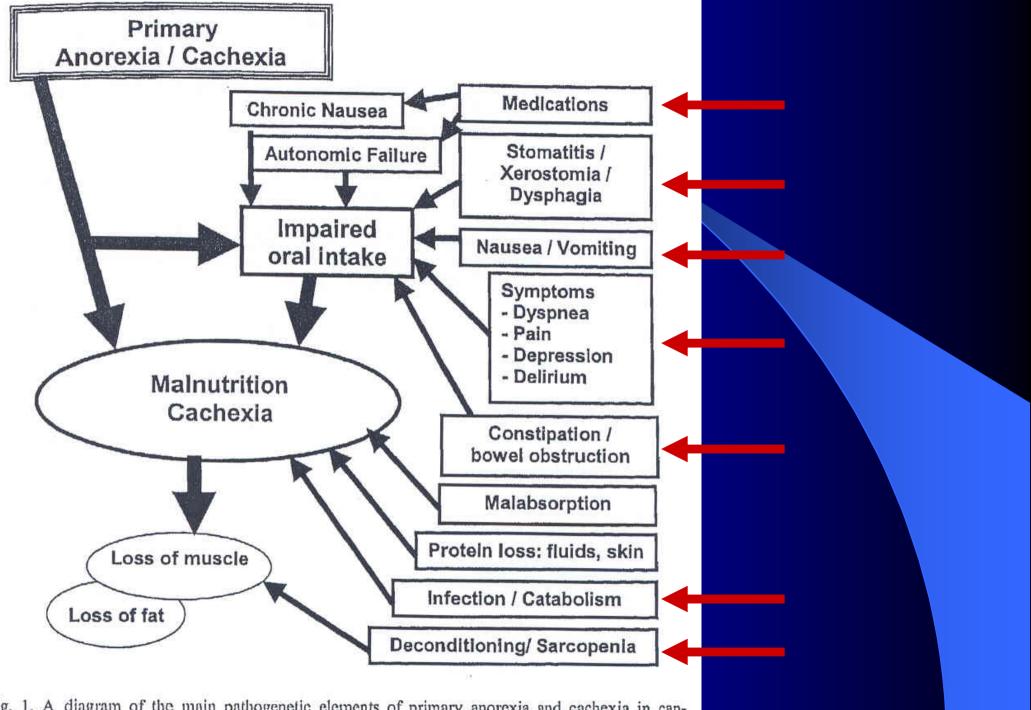
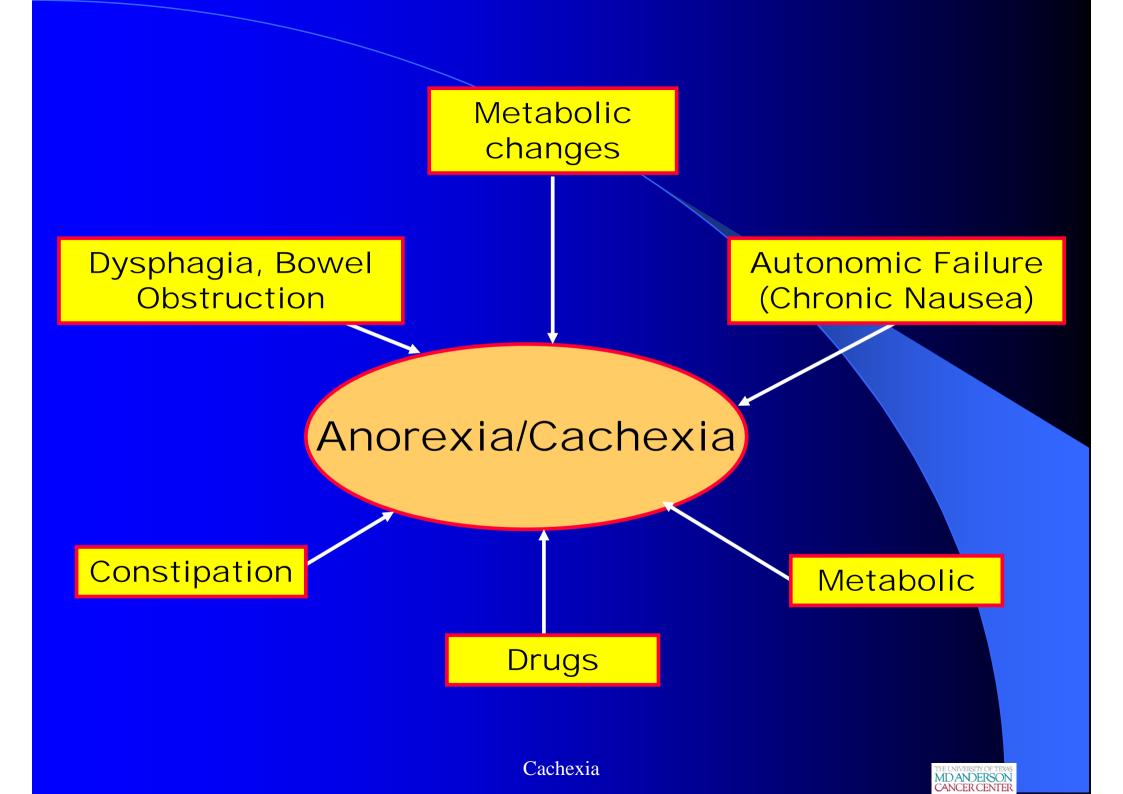


Fig. 1. A diagram of the main pathogenetic elements of primary anorexia and cachexia in caner patients.



## Primary outcome: depends on the intervention!

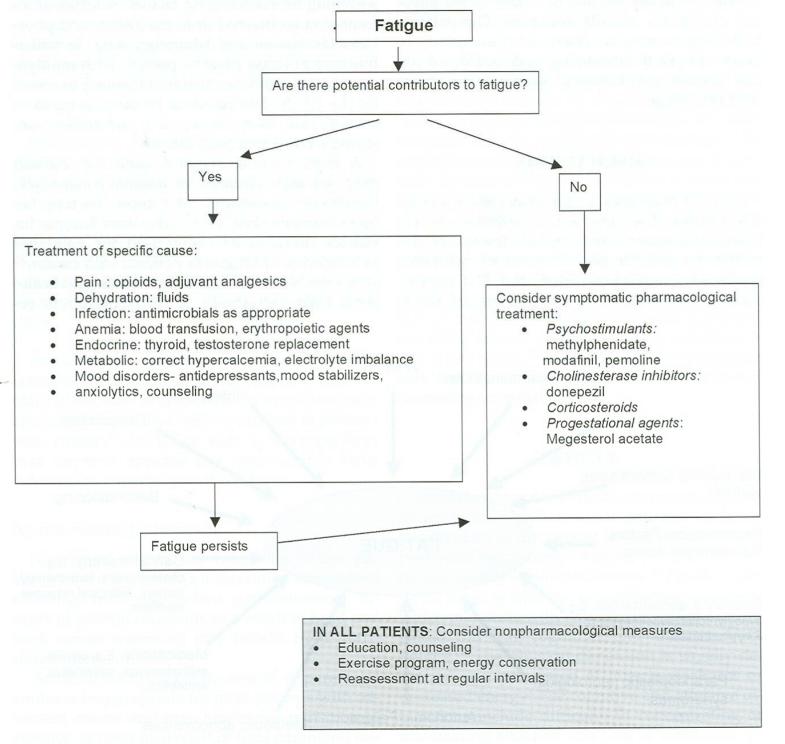
- Different outcomes need different time:
- 1. Symptoms are legitimate outcomes: anorexia, early satiety, family distress
- 2. Weight/ fat gain: body image, QoL
- 3. LBM/ body cell mass
- 4. Function!, fitness.
- Complications/ infections
- 6. Response to treatment: cancer/AIDS/leprosy,TBC
- 7. Survival



### **Combined therapy!**

- Primary cachexia : anti-proteasome, anti-cytokine, anti-inflammatory
- Nutritional support: Energy (+?)
- Exercise
- Target organs: appetite (ghrelin), muscle (testosterone)
- Secondary/ associate: depression, taste, nausea, constipation





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