

# **Ketogenic diet and cancer patients: can it evolve into a synergistic adjuvant avenue in the therapeutic algorithm?**

S.Zouganeli<sup>1</sup>, A.P. Stefanoyiannis<sup>2</sup>

<sup>1</sup> Department of Clinical Nutrition, University General Hospital of Athens "Attikon", Athens, Greece

<sup>2</sup> 2<sup>nd</sup> Department of Radiology, Nuclear Medicine Division, University General Hospital of Athens "Attikon", Athens, Greece

# 1. Background-Aim

---

**Ketogenic Metabolic Therapy (KMT):** Ketogenic Diet (KD) and other low-glycemic diets

- potential **anti-tumor effects**
- positive impacts on quality of life (**QoL**) and **survival**

**KMT in combination with radiotherapy**

- emerging field of investigation
- **potential complementary therapy** for various types of cancer

***Aim:** a review of recent evidence about **the role of KMT as an adjuvant to radiotherapy for cancer patients***

## 2. Materials & Methods

---

A review was conducted using the **PubMed platform**, with an advanced search employing the terms:

**“ketogenic diet” AND**

**“radiotherapy” AND**

**“cancer” OR “oncological patient”**

### 3. Results

---

✓ **Increasing number of pre-clinical and clinical studies**

*(mostly case reports /pilot studies, small number of RCTs) exploring the role of Ketogenic Metabolic Therapy(KMT) as an adjuvant in traditional therapies of certain cancer types (brain, breast, lung, head and neck, pancreatic, colorectal)*

✓ **KMT: various regimes applied during chemotherapy and/or radiotherapy**

- ketogenic diet in different ratios
- caloric restricted diet
- low carbohydrate diet
- fasting protocols

### 3. Results

---

✓ **multiple mechanisms of KMT to anti-cancer effect**

- targeting glucose (Warburg effect) and amino acid metabolism of cancer cells
- interfering with ROS production in cancer cells
- special actions of ketone bodies (e.g. epigenetic alterations, molecular signaling, anti-inflammatory effects)

✓ **significant KD anti-tumor effects concerning**

tumor growth, survival rate, cancer initiation, cancer-induced cachexia,  
enhancement of targeted therapy efficacy, sensitization of cancer cells to classic chemo or radiotherapy

✓ **different response to KD according to tumor special characteristics**

(e.g. genetic alterations, glucose uptake on PET scan)

## 4. Conclusions

---

- small number of RCTs
- heterogeneity in study design (different types of cancer, variety of chemotherapy and RT schedules, differences in diet composition, exogenous ketone esters, etc)
- questions about optimal KD ratio, time point, caloric restriction, fasting
- low compliance rate due to KD low palatability and tumor growth related complications

- ✓ **No robust evidence to form guidelines for the use of KMT in cancer patients**
- ✓ **Need for RCTs with protocol standardization**
- ✓ **KMT improves overall health, body composition and QoL in certain cancer patients**
- ✓ **KMT is a cost-effective adjuvant therapy to chemotherapy and RT with no serious adverse effects or toxicity**
- ✓ **KMT could be included in the therapeutic algorithm with careful implementation and monitoring by a multidisciplinary team of experts to increase radiotherapy efficiency and preserve patient's nutritional status**

## 5. References

---

1. Icard P. et al: Perspective: Do Fasting, Caloric Restriction, and Diets Increase Sensitivity to Radiotherapy? A Literature Review, *Adv Nutr* 2020;11:1089–1101
2. Klement R.J.: Fasting, Fats, and Physics: Combining Ketogenic and Radiation Therapy against Cancer, *Complement Med Res* 2018;25:102–113
3. Klement R.J et al: Ketogenic diets consumed during radio-chemotherapy have beneficial effects on quality of life and metabolic health in patients with rectal cancer, *Eur J Nutr* 61, 69–84 (2022)
4. Weber D.D et al: Ketogenic diet in the treatment of cancer. Where do we *stand*? *Molecular Metabolism* 33 (2020) 102-121