GIGANTIC INFLUENCES HORMONE IMBALANCES, PITUITARY ADENOMA, GENETIC MUTATIONS AND SYMPTOMS, DIAGNOSIS, DIFFERENTIAL DIAGNOSIS AS WELL AS TREATMENT OF GIGANTISM

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ABSTRACT

Gigantism is a rare endocrine disorder manifested by excessive growth in children and adolescents due to an overproduction of growth hormone. Genetic mutations also lead to gigantism. Symptoms include facial changes. Large hands as well feet, muscle weakness, cardiovascular issues, joint problems. Diagnosis involves measuring the level of GH and IGF-1. Treatment concludes surgical reboval of the tumor, medications, long-term follow-Up as well as hormone replacement treatment (HRT).

INTRODUCTION

Gigantism is a rare endocrine disorder characterized by excessive growth in children and adolescents due to an overproduction of growth hormone (GH). This condition leads to abnormal height and various health complications. In this article, we will delve into the clinical aspects of gigantism, exploring its causes, symptoms, diagnosis, and treatment.
Causes of Gigantism

1. **Hormonal Imbalance:**
   Gigantism primarily results from an overproduction of growth hormone (GH) by the pituitary gland. This excessive GH secretion often occurs due to a tumor or adenoma in the pituitary gland, known as a pituitary adenoma. These tumors stimulate the gland to release GH uncontrollably.

2. **Pituitary Adenoma**
   **Definition:** Pituitary adenomas are benign tumors of the pituitary gland.
   **Cause:** Most cases of gigantism result from a pituitary adenoma that produces excess GH.
   **Mechanism:** The tumor stimulates the pituitary gland to produce an abundance of GH, leading to excessive growth.

3. **Genetic Mutations**
   **Definition:** Rare genetic mutations can cause gigantism.
   **Cause:** Inherited genetic mutations affecting genes related to GH regulation.
   **Mechanism:** These mutations disrupt normal GH regulation, causing uncontrolled growth.

4. **McCune-Albright Syndrome**
   **Definition:** McCune-Albright syndrome is a genetic disorder.
   **Cause:** Activating mutations in the GNAS1 gene.
   **Mechanism:** These mutations lead to an overproduction of GH, causing gigantism as one of the symptoms.

5. **Carney Complex**
   **Definition:** Carney complex is a rare genetic disorder.
   **Cause:** Genetic mutations, often in the PRKAR1A gene.
   **Mechanism:** These mutations can lead to excessive GH secretion and gigantism.

6. **Ectopic GH Secretion**
   **Definition:** Rare cases involve tumors outside the pituitary gland.
   **Cause:** Tumors in other parts of the body can produce GH.
   **Mechanism:** These tumors release GH, leading to gigantism.

7. **Rare Syndromes**
   **Definition:** Gigantism can be associated with certain syndromes.
   **Cause:** Syndromes like Sotos syndrome or Beckwith-Wiedemann syndrome may involve excessive growth.
   **Mechanism:** Altered genetic regulation within these syndromes can lead to gigantism as a feature.

8. **Unknown Causes**
   **Definition:** In some cases, the exact cause of gigantism remains unknown.
   **Cause:** Despite extensive testing, no specific genetic or tumor-related cause can be identified.
   **Mechanism:** It is a challenging subset to diagnose and manage.

9. **Complication**
   **Definition:** Gigantism can lead to various health issues.
   **Cause:** Excessive GH can affect multiple organs and systems.
   **Mechanism:** Complications may include cardiovascular problems, joint issues, and metabolic abnormalities.

**Symptoms of Gigantism**

**Excessive Growth:**
The hallmark symptom of gigantism is excessive height. Affected individuals often grow much taller than their peers. This accelerated growth typically begins in childhood and continues until the growth plates in the bones close.

**Facial Changes:**
Gigantism can lead to distinct facial changes, including an enlarged jaw (prognathism) and thickening of the nose and lips. This can result in a coarser facial appearance.

**Large Hands and Feet:**
Hands and feet may also enlarge significantly. This can cause difficulties in finding appropriately sized footwear and gloves.

**Muscle Weakness:**
Excessive growth can strain muscles, leading to muscle weakness and reduced physical endurance.
Cardiovascular Issues:
Gigantism can increase the risk of cardiovascular problems such as hypertension and cardiomegaly (enlarged heart).

Joint Problems:
The rapid growth can put stress on joints, leading to joint pain and osteoarthritis.

Headaches and Vision Problems:
The pituitary adenoma responsible for gigantism may cause headaches and vision problems due to its proximity to the optic nerve.

Thickened Skin
Skin may thicken and become coarse.

Enlarged Organs
Internal organs may also enlarge, leading to complications.

Enlarged heart can result in cardiovascular issues.

Enlarged liver and spleen may cause abdominal discomfort.

Diagnosis of Gigantism

Hormone Testing:
The initial step in diagnosing gigantism is hormone testing. This involves measuring the levels of growth hormone (GH) and insulin-like growth factor 1 (IGF-1) in the blood. Elevated GH and IGF-1 levels are indicative of gigantism.

Imaging:
Imaging studies such as MRI or CT scans are performed to locate and assess the size of the pituitary adenoma responsible for the excess GH production.

Genetic Testing:
In some cases, genetic testing may be necessary to rule out genetic causes of excessive growth.

Differential diagnosis of Gigantism

Growth Hormone-Producing Pituitary Adenoma (Acronegadyl):
Elevated levels of insulin-like growth factor 1 (IGF-1). Symptoms of acral overgrowth (enlarged hands and feet), coarsened facial features, and organomegaly.

Pituitary Gigantism:
Typically occurs in children or adolescents due to excessive growth hormone secretion.

Rapid linear growth, delayed puberty, and enlarged features.

McCune-Albright Syndrome:
Associated with café-au-lait spots, polyostotic fibrous dysplasia, and precocious puberty.

Gigantism-like growth patterns in some cases.

Multiple Endocrine Neoplasia Type 1 (MEN1):
May present with pituitary adenomas causing gigantism.

Associated with other endocrine tumors (parathyroid, pancreatic).

Carney Complex:
May involve pituitary adenomas leading to gigantism. Often associated with cardiac and cutaneous myxomas.

Non-Pituitary Tumors:
Rarely, gigantism can result from non-pituitary tumors, such as lung or pancreatic tumors, secreting growth hormone-releasing hormone (GHRH).

Primary Hypothyroidism:
Severe and untreated hypothyroidism in children can cause excessive linear growth.

Genetic Syndromes:
Marfan syndrome, Sotos syndrome, and Beckwith-Wiedemann syndrome may exhibit gigantism-like features.

Exogenous Hormone Exposure:
Gigantism can occur in cases of excessive use of growth hormone or anabolic steroids.

Rare Genetic Disorders:
Conditions like Weaver syndrome or Simpson-Golabi-Behmel syndrome can cause overgrowth.

Isolated Growth Hormone Deficiency:
Rarely, gigantism-like features can occur if there’s a deficiency of factors inhibiting growth hormone release.

Psychosocial Factors:
Gigantism-like appearance can result from psychological issues causing exaggerated concern about one’s height.

Treatment of Gigantism

Surgical Removal of the Tumor:
The primary treatment for gigantism is the surgical removal of the pituitary adenoma.
removal of the pituitary adenoma. This procedure is known as transsphenoidal surgery. It aims to remove the tumor while preserving normal pituitary function.

**Medications:**
In some cases, medications like somatostatin analogs or growth hormone receptor antagonists may be used to lower GH levels before surgery or if surgery is not possible.

**Radiation Therapy:**
Radiation therapy may be considered when surgery and medication are ineffective or not feasible.

**Multidisciplinary Care:**
A team of specialists, including endocrinologists, neurosurgeons, and radiologists, is often involved in the treatment plan.

Psychological support and counseling may be necessary for patients dealing with the emotional and social challenges of gigantism.

**Long-Term Follow-Up:**
Regular follow-up appointments are essential to monitor hormone levels, growth, and potential complications.

Treatment adjustments may be needed as the patient’s condition evolves.

**Potential Complications:**
Gigantism can lead to various health issues, including cardiovascular problems, joint pain, and diabetes.

Treating and managing these complications is an integral part of the overall treatment plan.

**Patient Education:**
Educating patients and their families about the condition, treatment options, and potential side effects is vital for informed decision-making.

**Hormone Replacement Therapy:**
After successful treatment, some patients may require hormone replacement therapy to address deficiencies caused by the treatment.

**Support Groups and Resources:**
Encourage patients to connect with support groups and organizations dedicated to rare conditions like gigantism for emotional and informational support.

**CONCLUSION**
Gigantism is a rare condition with significant clinical implications. Early diagnosis and appropriate treatment are crucial to manage its symptoms and prevent complications. With advances in medical science, individuals with gigantism can receive effective treatment and lead healthier lives. Awareness of this condition is essential among healthcare professionals and the general public to ensure timely intervention and support for affected individuals.

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