The term Globus was first described by Purcell in 1707 who coined the term globus hystericus, the word globus originating from the Latin meaning “ball” and “hystericus” reflecting the then assumed psychological component of the disorder [1]. Globus is a persistent or intermittent non-painful sensation of a lump or foreign body in the throat. It is a commonly encountered clinical condition that is usually long-lasting, difficult to treat, and has a tendency to recur. It is a common disorder of indeterminate origin and constitutes about 5% of all new ENT referrals [2]. The literature reports a slight female preponderance [3]. The disorder was renamed globus pharyngeus in 1968 [4]. Globus Pharyngeus has more recently been defined as

a. A persistent or intermittent sensation of a lump or foreign body in the throat for at least 12 weeks,

b. Occurrence of the sensation between meals,

c. Absence of dysphagia and odynophagia,

d. Absence of pathological gastroesophageal reflux (GERD), achalasia, or other motility disorder with a recognized pathological basis (e.g., scleroderma of the esophagus) [5].

Potential Causes of Globus

Most of the time we do not use any scoring system to grade Globus Pharyngeus. The reflux symptom index and the reflux finding score are not particularly valid diagnostic tools when used in globus patients [8]. The Glasgow Edinburgh Throat Score (GETS) has been validated for use in globus but is not widely used [9].

The common causes attributed in the etiology of Globus Pharyngeus could be:

- Gastroesophageal Reflux Disease: There is a considerable debate over the involvement of gastroesophageal reflux in the etiopathogenesis of Globus Pharyngeus. A number of studies are both for and against the fact. Malcolmson was the first to link GERD to the globus sensation through the use of barium swallow. Chevaller et al. [10] looked at globus patients with and without typical GER symptoms. They found that 66.6% of the non-reflux globus group and 80% of the GER globus group had significant episodes of reflux (based on pH monitoring). Whereas in direct contrast, Chen et al. in a similar study found no evidence of reflux in globus patients based on ambulatory pH monitoring [11].

Despite this gastroesophageal reflux (GER) has been suggested to be a major etiology of this symptom, potentially accounting for 23%-68% of globus patients [10]. Two basic mechanisms have been proposed to explain the association between GERD and the globus sensation [10]:

a. Direct irritation and inflammation of the laryngopharynx by retrograde flow of gastric contents, also known as laryngopharyngeal reflux (LPR);
Factors favoring the involvement of GORD in etiopathogenesis of Globus Pharyngeus include:

a) Significant episodes of reflux based on pH monitoring studies.

b) Globus symptom score was significantly higher in patients with GERD than in those without.

c) Globus sensation improved after 8 wks of proton pump inhibitor (PPI) therapy.

d) Several population-based surveys have supported such a potential link between GERD and the globus symptom by demonstrating an increased risk of globus among patients with GERD symptoms.

e) Endoscopic evaluation also reveals a congested post cricoid area and posterior laryngitis in patients of Globus Pharyngeus. Although not always.

Psychological Factors and Stress: Globus Pharyngeus has traditionally been associated with Psychological problems either as causative factor or triggering factor. The association was initially thought to be so strong that the disease wads earlier known as Globus Hystericus. Many of these patients are suffering from cancer phobias because someone close developed a cancer in the near past and hence they develop this phobia, this is also the primary reason for investigating these patients only to rule out malignancy, however even after investigations many of these (44% after 5 years of follow up - [11,12] were symptomatic on long term followup. It is the fourth most discriminating symptom of a somatization disorder after vomiting, aphony, and painful extremities [11]. Personality studies have found higher levels of alexithymia, neuroticism, and psychological distress (including anxiety, low mood, and somatic concerns) and lower levels of extraversion in patients presenting with globus [10].

Many patients report stressful life events preceding symptom onset, suggesting that life stress might be a cofactor in symptom of the condition and in exacerbation. Indeed, up to 96% of patients with globus report symptom exacerbation during periods of high emotional intensity [10]. There is also a correlation between psychological factors and globus pharyngeus patients of the two types - one with Gastroesophageal reflux positive and the other with a negative Gastroesophageal reflux. Globus patients with LPR exhibited weaker psychological symptoms than non-LPR globus patients, and globus patients who did not respond to PPI had significantly higher anxiety scores [10]. Although etiological role of psychological factors have not yet been established but these factors must always be kept in mind while evaluating these patients.

Abnormal Upper Esophageal Sphincter Function: Contradictory studies are available for the role of USE function in the etiology of Globus pharyngeus. Some studies suggest that abnormal USE function has to be a cause of globus sensation. Elevated USE pressure has been found to be much more frequent in patients with globus sensation than in controls (28% vs. 3%), suggesting that hypertensive USE is a background factor for globus [10]. Additionally, injection of botulinum toxin into the cricopharyngeal muscle in a patient with both globus and extremely high USE pressure led to a resolution of the globus symptom and a decrease in USE pressure [10].

In a study conducted by Cook IJ [13] they examined the psychological profile and effect of acute mental stress (dichotic listening task) on USE tone in patients reporting to the clinic with globus sensation and concluded that in patients with a history of globus sensation, resting USE pressure and its response to stress is normal. They also suggested that USE hyper responsiveness to other stimuli or subjective intolerance to changes in USE pressure could account for symptoms of globus sensation.

Tokashiki et al. [13], however, showed that perfusion of HCl into the distal esophagus was related to a sensation of globus associated with a rise in UOS pressure. This rise in pressure was independent of the detection of a rise in pH in the hypopharynx. S. Korteque et al. [11] found that videofluoroscopic evidence of pharyngeal dysfunction especially laryngeal penetration had a strong association with globus.

Other Causes of Globus Pharyngeus

I. Pharyngeal inflammatory causes.

II. Cervical Osteophytes.

III. Esophageal motor disorders.

IV. Hypertrophy of the tongue base.

V. Upper aerodigestive tract malignancy.

VI. Retroverted epiglottis.

VII. Thyroid diseases.

VIII. Cervical heterotopic gastric mucosa.

IX. Rare tumors.

X. Others. Temporo mandibular joint disorders, hyper viscosity of the nasopharyngeal mucosa, Eagle’s syndrome, excessive laryngeal and pharyngeal tension, and salivary hypo function [10].
Conclusion

Globus Pharyngeus is a pharyngeal disorder of unknown etiology, the diagnosis of which is clinical. So far there are no studies that show that these patients eventually develop pharyngeal malignancy. A number of studies demonstrate that rigid endoscopy and barium swallow add little valuable information in evaluating these patients and investigations are done to rule out malignancy and patient satisfaction but unnecessary investigations must not be done, because of this reason there is no standard protocol for its diagnosis and management.

A number of studies suggest that GERD is a major cause of globus, though this remains under considerable debate. A number of other disorders like such as abnormal UES function, esophageal motility disorders, structural head and neck diseases, and psychological factors, have been suggested as potential causes of globus but none have been proved as the etiological factor. A complete head and neck examination including fibreoptic laryngoscopy can be done if patient’s symptoms are not relieved with routine management. It’s advisable that over investigating these patients can often add unnecessary stress to a group of these patients who already seem to have higher levels of depression, anxiety, and other somatic concerns. If the standard PPI therapy fails to relieve symptoms then other treatment modalities like speech therapy, anti-depressants, and cognitive-behavioral therapy, should be considered. A well-designed, randomized controlled study is required to establish the cause of the disease.

References