What you need to know

- Dystonia is a neurological condition characterised by abnormal postures and movements resulting from abnormal neural control of muscles.
- The most common forms of isolated dystonia in adults are focal, affecting the neck (cervical dystonia), eyes (blepharospasm), or associated with a task (eg, writer's cramp).
- Acute and tardive dystonia can occur as complications of medications such as dopamine receptor blockers.
- Neurophysiotherapy, botulinum toxin injections, and deep brain stimulation are effective treatments.
- Management of non-motor symptoms such as depression, anxiety, and associated pain are also important.

Dystonia

Dystonia is an abnormality of movement and posture caused by the abnormal neural control of muscle contractions. It is a highly stigmatising long term condition associated with a reduced quality of life. Management is a multidisciplinary partnership between general practitioners, specialists, therapists, and patients and seeks to alleviate motor and non-motor symptoms and the wider psychosocial repercussions of dystonia. This practice pointer highlights the common forms of dystonia, flags rarer causes which can have significant repercussions if not recognised, and offers a clinical roadmap for patient management.

What is dystonia?

Dystonia is defined as “a movement disorder characterised by sustained or intermittent muscle contractions causing abnormal, often repetitive, movements, postures, or both. Dystonic movements are typically patterned, twisting, and may be tremulous. Dystonia is often initiated or worsened by voluntary action and associated with overflow muscle activation.” A dystonic body part usually remains mobile and movements are often slow and effortful, with a reduced range. The most common forms of dystonia in adults are focal, affecting the neck (cervical dystonia), eyes (blepharospasm), or associated with a task (eg, writer’s cramp). However, dystonia can also present with a generalised pattern, which is more common with childhood onset forms. Dystonia is a subcategory of the hyperkinetic movement disorders in which too much or disordered movement is seen. Other subtypes include chorea, myoclonus, tics, and tremor. Pathophysiology and clinical descriptors of dystonia are outlined in boxes 1 and 2.

Box 1: Pathophysiology of dystonia

Dystonia was originally thought to be caused by disrupted basal ganglia function, as lesions in the basal ganglia (particularly the putamen) frequently resulted in dystonia. However, later studies showed that it is better conceptualised as a network disorder that also includes the cerebellum, thalamus, and cortex. Genetic mutations can cause generalised isolated dystonia (eg, TOR1A), focal isolated dystonia (eg, GNAL), and combined dystonia. Some evidence suggests that monogenic mutations converge on common molecular pathways. Other subtypes of dystonia, particularly focal dystonias, have stronger environmental influences (such as intensive stereotyped practice in musician’s dystonia). Traditionally, isolated dystonia has been used to infer the essential mechanisms that underwrite dystonia. Classic neurophysiological markers of dystonia, such as disordered plasticity responses or reduced inhibition, have limited utility as they are variable across individuals and are not specific to dystonia.

Box 2: Categories of key clinical descriptors for dystonia

| Age of onset | Body distribution | Temporal features | Associated features |
|--------------|-------------------|-------------------|-------------------|
| Infancy: birth to 2 | Focal: one body region | Disease course |
| Childhood: 3 to 12 | Segmental: two or more contiguous body regions | Static |
| Adolescence: 13 to 20 | Multifocal: two or more non-contiguous body regions | Progressive |
| Early adulthood: 21 to 40 | Generalised: two or more contiguous regions plus trunk | Variability |
| Late adulthood: over 40 | | Persistent |

Variability
- Persistent
- Diurnal
- Task specific
- Paroxysmal

Associated features
- Isolated dystonia
- Combined dystonia
- Coexists with another movement disorder, or
Confusingly, the term dystonia can be used interchangeably to refer to a range of related clinical entities. Dystonia can refer to a type of movement disorder (ie, pattern of abnormal movement, phenomenology), a clinical syndrome (eg, adult onset focal dystonia), and/or a specific diagnosis (eg, \textit{TORT1A}-related dystonia).\textsuperscript{3}

To move beyond such a non-specific use of the term dystonia, stratifying patients by clinical features often reveals likely causes and guides investigation.

\textbf{General approach}

When a patient presents with a movement disorder consistent with dystonia, a targeted history can help pick out the key clinical features (box 2). Consider the age of onset, body distribution, course of disease, variability, and whether dystonia is associated with other features (box 2). For example, dystonia that begins in childhood is more likely to have a discoverable acquired cause and is more likely to progress from focal to generalised. In contrast, a typical adult onset focal dystonia is likely to be a subtype of isolated dystonia and will rarely spread. The family history reveals the likelihood of a genetic cause. Ask if patients can alleviate their dystonia by lightly touching surrounding body areas (“sensory trick”), a phenomenon that illustrates the dynamic nature of dystonia and the importance of sensory influences.\textsuperscript{12} Dystonia is often accentuated within a neurological examination that includes a range of postures and tasks. A latent abnormal posture (eg, in the head and neck) can be revealed by asking the patient to close their eyes and let position drift to where it feels most comfortable. If tremor accompanies dystonia it is usually jerky, variable in amplitude, and worsened by particular positions.

\textbf{Types of dystonia}

\textbf{Isolated dystonia}

In isolated dystonia, dystonia and/or tremor are the only symptoms and signs.\textsuperscript{3} These are chronic, relatively selective disorders of posture and movement that are non-degenerative and have no effect on lifespan. Adult onset focal isolated dystonia has a typical anatomical distribution of onset. Cervical dystonia, blepharospasm, and task specific dystonia are the most common forms (fig 1). Other subtypes include oromandibular dystonia with prominent jaw opening and closing. Laryngeal dystonia (also known as “spasmodic dysphonia”) affects the voice and either adduction or abduction of the muscles responsible for phonation is seen. The true prevalence of adult onset focal dystonia remains unknown and estimates range between 16 and 730 cases per 100 000 individuals.\textsuperscript{13 14} Adult onset focal forms rarely progress to generalised dystonia. By contrast, early onset generalised isolated dystonia often begins in childhood in a limb and then generalises over months. Early onset variants of dystonia have a lower prevalence of approximately 7 per 100 000 individuals.\textsuperscript{13}
Isolated dystonias can be subtle initially and diagnosis can be delayed for many months or years. Prompt recognition is often limited by a lack of knowledge among medical professionals. Part of the difficulty is that no reliable diagnostic test exists for dystonia, and routine brain imaging is unremarkable. In the future, kinematic, 

| Young onset generalised dystonia |
|----------------------------------|
| Onset in childhood or adolescence of dystonia in the limb |
| Symptoms then generalise over months |
| Over half will have mutation in the TOR1A gene (DYT-1) which is autosomally dominant with reduced penetrance: ~30% of mutation carriers develop symptoms |

| Task specific dystonia |
|------------------------|
| Age of onset ~35 years |
| Men > women |
| Writer’s cramp is most common type: symptoms occur while writing and not during other manual activities |
| Highly skilled tasks are higher risk |
| Affects 1% of professional musicians |

| Cervical dystonia |
|-------------------|
| Also known as spasmodic torticollis |
| Age at onset ~45 years |
| Women > Men |
| Gradual onset of discomfort in neck then involuntary pulling of the head |
| Progression over 6-12 months then plateau |

| Blepharospasm |
|---------------|
| Age of onset ~60 years |
| Women > Men |
| Patients frequently report a gritty or uncomfortable feeling in the eyes before onset of spasm in orbicularis oculi muscles |

Fig 1 | Different clinical patterns of isolated dystonia. Children and adolescents typically present with a generalised dystonia whereas presentation in adults is usually with a focal dystonia |
Combined dystonia

Dystonia with another movement disorder and/or occurrence of other neurological or systemic features is referred to as combined dystonia. As there are many potential causes, syndromic associations are used to narrow down diagnostic possibilities. Deciding which movement disorders or other features dominate is key. For example, foot dystonia and mild parkinsonism in a young adult is suggestive of a genetic cause of Parkinson’s disease (such as a LRRK2 mutation). Many centres now have panels of genetic tests that are performed according to the specific syndromic clustering.

Two treatable disorders are worth emphasising: Wilson’s disease and dopa responsive dystonia. Dystonia and liver abnormalities and/or psychiatric symptoms raise the possibility of Wilson’s disease. This autosomal recessive disorder of copper metabolism has a mean age of 18 for neurological presentation, often with a mixed movement disorder. Full blood count, liver function tests, and serum caeruloplasmin are useful initial investigations. Twenty four hour urine copper and slit lamp examination (for Kayser-Fleischer rings) can be organised if serum caeruloplasmin is low, neuro-imaging is abnormal, or there is a high index of suspicion for Wilson’s disease. First line treatments include copper chelating agents that increase urinary copper excretion or zinc salts that inhibit the intestinal absorption of copper. Dopa responsive dystonia encompasses a group of genetically and clinically heterogeneous disorders that often manifest with diurnal fluctuation of dystonia (worse at end of the day). Most patients have a deficiency in an enzyme involved in the biosynthesis of dopamine, and some forms are dramatically responsive to low doses of levodopa. The typical age of onset is in early childhood with a progressive limb dystonia. However, owing to the variability of presentation and difficulties making a laboratory diagnosis, many patients with young onset dystonia (<40 years) may receive a trial of levodopa to assess for the possibility of dopa responsive dystonia.

Acquired dystonia

Acquired dystonia occurs secondary to a precipitating event. Approximately half of children with dystonia have an acquired cause for their condition; causes include hypoxic brain injury at birth, preterm delivery, and kernicterus. Timing of the onset of dystonia can vary after a causal event. For example, in drug induced dystonia, acute reactions can be seen hours after the administration of antidopaminergic medications (eg, metoclopramide, prochlorperazine). One of the most common presentations is an oculogyric crisis, where the patient has upward, conjugate, tonic deviation of the eyes. Patients require urgent treatment, especially if laryngeal involvement compromises breathing, and supportive measures and intravenous anticholinergic usually have good effect. By contrast, tardive dystonia can occur many months or years after chronic exposure to a range of medications such as antipsychotics (eg, haloperidol). This is seen particularly in younger patients where a predominantly axial dystonia results in hyperextension of the spine and neck. Tardive dyskinesia (choreiform movements) is often seen in combination with the dystonia. Dystonia can unfortunately persist even after the causative drug is discontinued.

Management

Patients with suspected dystonia should be referred to a neurologist specialising in movement disorders for further evaluation and to access treatments. Basic blood tests, including a full blood count, urea and electrolytes, liver function tests, and caeruloplasmin can be sent at this point. It is reasonable to leave brain imaging until neurological review, as imaging protocols are often adjusted to the patient’s clinical features. Treatment is tailored to the cause of dystonia. As a general rule, the first line treatment of focal dystonia is botulinum toxin injections, whereas generalised dystonia is treated with oral medications and surgery.

Botulinum toxin injections

Regular botulinum toxin injections are the mainstay of treatment for cervical dystonia (high quality evidence) and blepharospasm (moderate quality evidence), with randomised control trials showing that dystonia and quality of life are improved by injections (fig 2). The toxin prevents the release of the neurotransmitter acetylcholine from axon endings at the neuromuscular junction, and works by causing weakness in the dystonic muscle. Commercially available types of botulinum toxin have differences in potency, and doses across types are not equivalent. Treatment is often delivered via specialist botulinum toxin clinics within neurological or ears, nose, and throat services (spasmodic dystonia).
Oral medications
Anticholinergic medications such as trihexyphenidyl can significantly reduce dystonic contractions in generalised dystonia. The dose of anticholinergics must be titrated up gradually to minimise side effects such as dry mouth, gastrointestinal upset, and confusion (particularly in older patients). Muscle relaxants, such as baclofen and benzodiazepines, are also used.

Surgery
The most common surgical approach to treat dystonia is deep brain stimulation, in which continuous stimulation is delivered to deep nuclei in the brain (mostly commonly basal ganglia and thalamic nuclei). A neurosurgical operation is required to implant permanent electrodes within the brain and these are connected via subcutaneously tunnelled leads in the neck to an implanted stimulator in the chest. The implanted stimulator can be programmed and checked by communicating hardware in clinic/at home. The precise mechanism of deep brain stimulation is debated but its long term efficacy is well documented in randomised controlled trials for isolated young onset generalised dystonia (such as TOR1A related dystonia) and refractory cervical dystonia. In the future, individually tailored surgical options will be available with novel stimulation sites and adaptive stimulation paradigms.

Neurophysiotherapy
Access to specialist neurophysiotherapy for dystonia is variable. An emerging evidence base suggests that patients can regain motor control, improve dystonic contractions, and reduce subsequent disability with neurophysiotherapy. Neurophysiotherapy is best tailored to the specific subtype of dystonia.

Non-motor symptoms
Although the treatment of dystonic muscle contractions is often given priority, depression and anxiety are common and can be just as disabling. Pain of the affected body part is also present in more than half of patients. Pain scores have been shown to improve with analgesia and neuropathic medications, effective treatment of dystonic contractions, and treatment of associated neuropsychiatric symptoms.

Education into practice
- What categories can be used to describe the clinical features of a patient presenting with dystonia?
- How would you summarise available treatments to a patient?

Resources for patients
- These organisations have a range of educational materials such as fact sheets, podcasts, and videos and work to connect people with dystonia, raise awareness of dystonia, and advance research initiatives:
  - https://www.dystonia.org.uk,
  - https://dystonia-europe.org,
  - https://dystonia-foundation.org,
  - https://www.neurosymptoms.org/en_GB/symptoms/fnd-symptoms/functionaldystonia/,
  - https://www.ern-rnd.eu/disease-knowledge-hub/dystonia/

How this article was created
We conducted a search of PubMed for evidence for dystonia until November 2021. AS wrote the first draft and then each author contributed to the manuscript until we had fully integrated our range of viewpoints. Our expertise bridges neurology, neurophysiotherapy, general practice, education, and patient advocacy.

How patients were involved in the creation of this article
- One of the authors is a patient and patients’ advocate (with Dystonia Europe, Polish Dystonia Society) and has been fully involved with the planning and writing of the article.

Contributorship and the guarantor: All authors wrote and reviewed the article and the guarantor is Anna Sadnicka.

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