Neurological control of human sexual behaviour: insights from lesion studies

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We review the human literature examining the effects of neurological insult on human sexual behaviour. We provide a synthesis of the findings to date, and identify key brain regions associated with specific aspects of human sexual behaviour. These include subcortical and cortical regions, with the mesial temporal lobe and the amygdala in particular being a crucial structure in the mediation of human sexual drive.

Despite its fundamental role in human life, there has been surprisingly little research into the neurological control of human sexual behaviour. Blumer and Walker proposed two reasons for this neglect. Firstly, physicians are not routinely trained to explore the sex life of their patients, and secondly, “there has been a stifling trend in research to investigate only what can be measured by objective methods”. Now 30 years later, human studies of the neurological control of sexual behaviour remain scarce. Although we have gained some knowledge from animal studies, the validity of comparing the sexual behaviour of rodents and humans is questionable. In this review, we focus solely on human research. Recently, there have been a number of functional neuroimaging studies of human sexual arousal. These studies have typically used visual sexual stimuli that simultaneously activate numerous brain regions associated with a range of sexual functions. Thus although these studies can identify brain areas involved in the mediation of human sexual behaviours, they cannot determine which areas are necessary for particular functions.

Studies of patients with brain lesions are unique in enabling us to examine specific brain–behaviour relationships. Current human lesion research examining the effects of neurological insult on human sexual behaviour comprises case studies, incidental observations and group studies of sexual changes associated with a particular neurological disease, cerebral injury, neurosurgery or direct cortical stimulation. The aim of this review was to synthesise this human literature, and to reflect on the insights that this provides, including identification of brain regions mediating specific aspects of human sexual behaviour. The findings highlight the critical role of the mesial temporal lobes and the amygdalae in the mediation of human sexual drive.

A crucial issue to consider in understanding the neural basis of human sexual behaviour is the distinction between disinhibited sexual behaviour as part of generalised behavioural disinhibition, and hypersexuality, or dramatically increased sexual drive and activity occurring in relative isolation. These two manifestations have been distinguished on the basis of theorised neuroanatomical substrates and effective treatments.

Disinhibited sexual behaviour is considered to be a feature of general behavioural disinhibition following paralimbic and neocortical frontotemporal lobe injury, whereas “true hypersexuality” is more commonly associated with limbic and temporal lobe injury (see “Hypersexuality after temporal lobectomy” below). Likewise, hyposexuality may occur following injury to the prefrontal convexity, as part of a general tendency towards apathy and indifference, or in relative isolation, as in temporal lobe epilepsy. We emphasise the manifestations of hypersexuality and hyposexuality in considering brain–behaviour relationships.

METHODOLOGY

Articles were ascertained using the databases “PsychInfo” and “PubMed”. Keywords included “brain”, “sexual behaviour”, “sexual function”, “hypersexuality” and “hyposexuality”, limiting the search to human studies published in English. The literature spans 68 years from 1937 to 2005. The references of articles obtained from the databases were also searched to identify other relevant studies to obtain a comprehensive listing of articles. In this paper, we cite those references that are key to building a coherent picture of the key brain regions mediating human sexual behaviour. Despite these measures, we found a paucity of research addressing this significant clinical issue. Based on the articles reviewed, we have identified particular brain areas in which there have been several reports of lesions causing alterations in sexual behaviour. In addition, we briefly summarise recent functional neuroimaging studies that complement the findings of the lesion research and provide additional evidence in support of the key brain regions identified. The literature has been divided into the broad categories of subcortical and cortical studies. Firstly, we identify neurological insult literature supporting the role of three subcortical regions, specifically the septal region, hypothalamus, and ansa lenticularis and pallidum. This is followed by a review of literature identifying the role of three cortical regions in the mediation of specific aspects of human sexual behaviour, namely the frontal, parietal and temporal lobes, and in particular the
Subcortical regions
The septal region
There have been two observations of sexual responses or altered sexual behaviour after stimulation or damage to the septal region of the limbic system. Heath conducted direct electrical and chemical stimulation of various areas of the brain in a sample of 54 patients with a range of conditions, including schizophrenia and narcolepsy. He found that a subjective "pleasurable response" and varying degrees of sexual arousal were consistently elicited by electrical stimulation of the septal region in all patients. Heath and developed a self-stimulating device that was attached to intracranial electrodes. This device was based on the technique developed by Olds and Milner to study the pleasure response in animals. Septal self-stimulation elicited orgasm and a compulsion to masturbate. Chemical stimulation of the septal region elicited similar sexual responses.

Further evidence of the role of the septal nuclei in the mediation of sexual behaviour comes from an incidental finding reported by Gorman and Cummings. They described two male patients who manifested hypersexuality in the form of a dramatic increase in sexual interest and activity after the insertion of ventriculoperitoneal shunts for the treatment of hydrocephalus. In both patients, CT scans revealed that the tip of the shunt catheter was lodged in the septum. Gorman and Cummings concluded that the septal region was responsible for the markedly increased sexual activity and that the septal nuclei form one focus on an amygdaloïd-hypothalamic-septal circuit that mediates sexual behaviour. In particular, the septal region appears to be intrinsically involved in pleasurable response and orgasm.

The hypothalamus
Isolated hypothalamic lesions have been reported to reduce or abolish sexual drive. One 50-year-old male patient in the case series of Miller and colleagues had hypothalamic damage resulting from an infiltrating glioma of the left midbrain hypothalamic region. He developed erectile and ejaculatory difficulties and a paedophilic sexual preference.

In the 1970s, certain psychosurgical techniques were developed to target sexual behaviour. These were based on experimental work in animals that demonstrated that destruction of the hypothalamic ventromedial nuclei led to the conversion of feline post-amygdalectomy hypersexuality to hyposexualit. Roeder and colleagues subsequently developed a stereotaxic method of treating "sexual deviations" in humans. Stereotaxic lesions of the ventral medial hypothalamic nucleus were performed in 10 patients with "sexual deviation" ranging from paedophilic homosexuality to intractable exhibitionism. The results showed that sex drive was diminished or abolished in all cases.

Three case studies of patients with striking changes in sexual behaviour have been reported that were associated with lesions that included the hypothalamus but extended to other brain regions. For example, Poock and Pilleri described a female patient who displayed periodic hypersexual and aggressive behaviour after encephalitis of unknown etiology. Neuropathological examination revealed extensive lesions to limbic structures, including the hypothalamus, thalamic nuclei and the fornix. Ortego and colleagues described a woman with neuropathologically confirmed multiple sclerosis who developed hypersexuality and multiple paraphilias, including pae-dophilia, zoophilia and incest over the 2 months prior to her death. She had extensive lesions of limbic structures, including the hypothalamus, basal frontal, septal and temporal regions. Frohman and colleagues described a male patient with altered sexual behaviour characterised by an obsessive and insatiable desire to touch women's breasts. Neuroimaging showed enhancing lesions on the right side of the hypothalamus and mesencephalon. In summary, these case studies demonstrate that alterations in sexual behaviour and/or orientation can occur in association with various combinations of limbic structure lesions, including the hypothalamus. In particular, lesions of the hypothalamic region appear to mediate the neuroendocrine and autonomic aspects of sexual drive, with focal lesions resulting in a reduction or abolition of sexual drive. In contrast, lesions including but not restricted to the hypothalamus have resulted in increased sexual drive. This apparent paradox may reflect the complexity of this structure and its associations with other brain regions, with the possibility that different nuclei within the hypothalamus serve opposing functions.

Neuroimaging studies of sexual arousal elicited by visual stimuli or during ejaculation have found activation of subcortical regions, including the hypothalamus. For example, Ferretti and colleagues found peak hypothalamic activation at the initial phase of erection and suggested that the hypothalamus may serve to trigger an overt sexual response. These observations support the findings from lesions studies that the hypothalamus is a key brain region in the mediation of human sexual drive.

The ansa lenticularis and pallidum
Interrupting a bundle of myelinated fibres known as the ansa lenticularis has been found to effect sexual function. These fibres project from the globus pallidus of the basal ganglia to the thalamus. Meyers described four patients who exhibited an enduring loss of libido following bilateral surgical lesioning of the ansa lenticularis for the relief of bilateral abnormal movements. Two of these patients presented with Parkinsonian tremors and rigidity, and the other two with myoclonic movements. Following surgery all patients experienced alibido, accompanied by impotence in the males. Because of their structural proximity, however, surgical lesioning of the ansa lenticularis can affect the perifornical grey matter, posterior septal region and the dorsomedial nucleus of the hypothalamus. Thus it is not possible to ascertain the exact structure or combination of structures that were responsible for the postoperative alibido in these patients.

Two cases of hypersexuality after pallidotomy for Parkinson's disease have been reported. Mendez and colleagues described a male patient who manifested hypersexuality immediately following right pallidotomy, which persisted for 1 year and then gradually decreased with a tapering of his antiparkinsonian medications. Roane and colleagues described a patient who previously underwent a right pallidotomy and developed hypersexuality after surgical implantation of a deep brain stimulator electrode in the left globus pallidus. Overall, these findings suggest that the ansa lenticularis and pallidum play a role in the mediation of sexual drive, and lesions to this area may cause hypersexuality or alibido with impotence in males.

Cortical regions
The frontal lobes
Disinhibited sexual behaviour has been reported following damage to the frontal lobes, particularly the orbitofrontal region of the limbic system. As stated in the introduction, some authors consider such behaviour to be a byproduct of
general behavioural disinhibition occurring after frontal lobe damage.\textsuperscript{7, 8}

The effect of frontal leucotomy on sexual behaviour has been assessed,\textsuperscript{26–29} with some researchers reporting cases of hypersexuality\textsuperscript{30–32} and others describing no changes after leucotomy.\textsuperscript{27} Levine and Albert\textsuperscript{33} conducted a thorough exploration of sexual changes after “frontal lobotomy” in 40 patients with various psychiatric diagnoses and found that four patients experienced sexual changes that caused social difficulties, but this was a new feature in only one patient.

Sexual automatisms characterised by writhing, thrusting, rhythmic pelvic movements and handling of the genitalia have been reported in patients with seizures arising from the frontal lobes.\textsuperscript{34} Historically, these movements were thought to indicate non-epileptic seizures, or hysteria, and the bizarre behaviour that can occur during frontal seizures can still result in the erroneous diagnosis of hysteria.\textsuperscript{35} Spencer and colleagues\textsuperscript{36} observed sexual automatisms in four of 14 patients with frontal lobe lesions. Leutmezer and colleagues\textsuperscript{37} proposed that sexual automatisms should be subdivided into two distinct groups: (1) discrete genital automatisms such as grabbing and fondling the genitals, related to temporal lobe seizure onset and (2) hypermotoric sexual automatisms involving pelvic or truncal thrusting accompanied by genital manipulation, representing a frontal onset. These observations suggest that sexual seizures arising from the frontal lobes activate motor components of sexual behaviour. In other words, it appears that the frontal lobes mediate the motor components of sexual behaviour in addition to the control of sexual response that may be disinhibited after frontal lobe damage in the context of general behavioural disinhibition.

Activation of numerous frontal regions, including the right prefrontal cortex,\textsuperscript{38–40} anterior cingulate cortex and gyrus\textsuperscript{19, 20, 34} and orbitofrontal region\textsuperscript{39} has been observed during sexual arousal involving masturbation induced orgasm.\textsuperscript{40} Orbitofrontal activation has been interpreted as being related to the representation of pleasurable bodily sensations,\textsuperscript{39, 41} while dorsal anterior cingulate activation has been attributed to the modulation of skeletomotor activities that characterise sexual arousal\textsuperscript{42} and the perceived urge to act.\textsuperscript{43} Ferretti and colleagues\textsuperscript{44} found that during the initial phase of erection anterior cingulate activation peaked and was then maintained while the erection was sustained. They suggested that the activation may reflect the continual correspondence between the affective value of the stimulus and the sexual response.\textsuperscript{45} These studies complement those of lesion studies in identifying a critical role of the frontal lobes in human sexual response.

The parietal lobes

The parietal cortex, specifically the “paracentral lobule”, has been implicated in tactually related sexual phenomena. Sexual seizures arising from the parietal cortex have certain distinguishable features.\textsuperscript{35–38} They may involve heightened sexual arousal or sensations in the genitalia or other perigenital areas. The individual is typically conscious and alert during the episodes and can recognise their sexual nature without necessarily finding them pleasurable or erotic.\textsuperscript{35–36} The “ego alien” quality of some parietal sexual seizures has been highlighted by several case studies which noted that focal genital sensations may be described by patients as irritating, painful or frightening.\textsuperscript{37–38}

Bancaud and colleagues\textsuperscript{39} described 10 patients with seizures involving sexual manifestations due to involvement of the paracentral lobule and 26 cases of temporal lobe sexual seizures. The authors reported that seizures arising from paracentral lobule lesions involved lateralised parasthesias of the genital areas and occasionally the chest, which were experienced without sexual resonance. In contrast, sexual seizures arising from the temporal lobe involved sexual emotions. This distinction was contradicted by Erickson\textsuperscript{40} who described a female patient with a haemangioma in the right paracentral lobule who experienced episodic “hot spells” and genital sensations, as if she were having sexual intercourse. These symptoms were mislabelled “erotomania” or “nymphomania”.\textsuperscript{41} Bates\textsuperscript{42} observed anal retraction on stimulation of the paracentral lobule, while central fissure stimulation elicited sensation on the opposite side of the penis,\textsuperscript{43} and the spreading of paraesthesia from the abdominogenital area to the isilateral nipple.\textsuperscript{44} Orgasm and associated autonomic changes may result from the spreading of seizure activity from the parietal or temporal region to the hypothalamus\textsuperscript{45} but the specific neuroanatomical localising value of orgasm is unknown.\textsuperscript{46} In summary, these observations suggest that the medial surface of the parietal lobe is specifically involved in the mediation of genital sensations.\textsuperscript{35–36}

The temporal lobes

The area of the limbic system most frequently implicated in the mediation of human sexual behaviour is the temporal lobes. Accumulating evidence of changes in sexual behaviour following temporal lobe dysfunction has led to the phrase the “libidinous temporal lobe”.\textsuperscript{47}

Klüver Bucy syndrome

The integral role of the temporal lobes in the mediation of sexual behaviour was demonstrated in the manifestation of hypersexuality following bilateral temporal lobectomy in rhesus monkeys and subsequently in humans.\textsuperscript{48} Bilateral temporal lobectomy produces a constellation of symptoms that constitute Klüver Bucy syndrome (KBS), including: (1) visual agnosia, an inability to recognise objects by sight, (2) hyperorality, a tendency to examine all objects by mouth, (3) “hypermetamorphosis”, an irresistible impulse to react and attend to visual stimuli, (4) emotional changes, particularly an absence of fear or expressions of anger, often termed the “taming effect”, (5) changes in dietary habits and (6) hypersexuality.

There have been 27 reports of full blown or partial human KBS. Terzian and Dalle Ore\textsuperscript{49} first reported a case of KBS in a man who underwent bilateral temporal lobectomy for intractable epilepsy. With the exception of hyperorality, all other KBS features were present. Hypersexuality manifested as frequent exhibitionism, homosexual and heterosexual advances toward doctors, and masturbation several times per day. The first full blown case of human KBS was described by Marlowe and colleagues\textsuperscript{50} in a male with herpes encephalitis and associated bilateral temporal lobe damage. Hypersexuality manifested as frequent inappropriate sexual references in conversation and numerous homosexual advances. In their review of human KBS, Lilly and colleagues\textsuperscript{51} noted that alterations of sexual behaviour are common and typically manifest as sexual advances and comments whereas increased copulation or masturbation and changes in sexual orientation are less frequent. Poock\textsuperscript{52} argued that the chief manifestation of KBS is hyperorality, and proposed that only patients with this feature alone or in combination with other features should be described as manifesting KBS. This notion has implications for the conceptualisation of hypersexuality as a Klüver Bucy feature after temporal resection and suggests that the exclusive manifestation of hypersexuality without other KBS features should not be labelled “KBS”.

In almost all cases displaying partial or full blown KBS there has been bitemporal damage. There has been no attempt to further delineate the structures within the temporal lobe that may be responsible for this syndrome, although a number of authors have postulated involvement of the amygdala on the basis of findings from animal studies. Bilateral lesions limited
to the amygdala in humans, however, do not consistently produce hypersexuality or other features of KBS (see discussion below).

Paraphilias and sexual orientation
There have been 11 reports of paraphilias, such as fetishism, associated with temporal lobe dysfunction in the literature.50–53 Kolarsky and colleagues54 examined the relationships between “sexual deviation”, age of lesion onset and localisation of lesion (temporal vs extratemporal). The authors defined two diagnostic categories: (1) “sexual deviation”, involving a deviation of sexual object (for example, paedophilia). Homosexuality was included in this category, which would now be considered inappropriate, and (2) “sexual disturbances other than deviations”, including orgasm in response to stimuli unrelated to the subject’s sexual preference, hypersexuality and hyposexuality.55 The results showed that “sexual deviation” was associated with early onset lesions (prior to age 3 years). There was an insufficient number of participants with extra-temporal lesions of known duration to determine whether the presence of an early lesion per se or an early temporal lesion was the effective variable. Nevertheless, the finding that temporal lesions were significantly more common in patients with “sexual disturbances other than deviations” compared with those without sexual disorder led the authors to claim an association between temporal lobe lesions and “sexual disturbance”. They carefully noted, however, that brain lesions are not necessarily present in all cases of sexual disorder.54

An association between temporal lobe abnormalities and paedophilia has been reported by Mendez and colleagues.56 They described two cases of homosexual paedophilia occurring late in life associated with frontotemporal dementia and bilateral hippocampal sclerosis. Both male patients showed right temporal lobe hypometabolism on positron emission tomography. The authors concluded that bitemporal dysfunction, with a right-sided predominance, results in hypersexuality. They emphasised, however, that there is currently no evidence for a “specific ‘paedophilia lesion’ in the brain”.55

Temporal lobe epilepsy
Interictal sexual behaviour
Alterations in interictal sexual behaviour have been frequently reported in patients with epilepsy, particularly temporal lobe epilepsy (TLE). The most common interictal sexual dysfunction associated with TLE is hypersexuality.54–56 Hypersexuality is typically defined as sexual activity less than once per month, and may or may not be accompanied by erectile or orgasmic dysfunction.58

Early studies describing a relationship between hypersexuality and TLE have received methodological criticisms.60 Control groups and female patients were rarely included, and diagnostic criteria for epilepsy were often poorly defined. Many studies included heterogeneous or biased samples without consideration of comorbid conditions. Nevertheless, the majority of studies have supported an association between TLE and hypersexuality,51–52 with few exceptions.63–64 This provides further evidence of the critical role of the temporal lobes in the sexual neural network.

Ictal sexual behaviour
In general, temporal lobe sexual seizures are thought to involve sexual emotion, which can be contrasted with the isolated genital sensations typically associated with parietal lobe sexual seizures. The manifestation of sexual seizures arising from the temporal lobe is varied. Such seizures can involve erotic feelings that may or may not be accompanied by genital sensations and/or orgasm. Remillard and colleagues59 described 12 female patients with ictal sexual phenomena associated with temporal lobe seizures. The sexual component was the initial event of the complex partial seizure in eight patients. This typically involved an erotic and pleasurable sensation in the genitalia or other erogenous areas, occasionally proceeding to orgasm in four patients. Alternatively, a feeling of sexual arousal or pleasure preceded a brief loss of consciousness.60 Warneke61 described a male patient with TLE who experienced an orgasmic-like sensation which spread across his face. Sexual automatism involving sexual statements, simulated intercourse, masturbation and exhibitionism for which the patient is subsequently amnesic have also been reported.62–64 These automatisms may or may not be preceded by a sexual aura.

Finally, postictal sexual arousal has been described in patients with TLE.65 It typically involves the patient seeking out a “customary sexual partner” in an appropriate setting such as a sexually responsive partner. It is recalled by the patient and is not considered part of the immediate postictal phase. It was noted that postictal sexual arousal was only observed among patients who were not markedly hyposexual interictally.1

A number of studies have proposed right hemisphere dominance in the mediation of sexual function. An increased incidence of hypersexuality has been reported in patients with right-sided TLE.59–70–71 There is also evidence that ictal sexual manifestations, such as orgasmic auras, typically originate from the right hemisphere.65 Toone67 has claimed that the proposed laterality effect of seizure related sexual phenomena is supported by the published literature which shows a bias in favour of right-sided pathology over left in a ratio of approximately 2:1. Finally, in a recent review of case studies of patients with hypersexuality and hypersexuality and unilateral lesions, Braun and colleagues67 found that hypersexuality was associated with left-sided lesions and hypersexuality with right-sided lesions.

Sexual outcome after temporal lobectomy
Ten studies have examined the effect of epilepsy surgery on sexual behaviour. In general, the majority of patients in each study reported no postoperative change. If a change was described, it was typically an improvement or increase in sexual behaviour. Bladin70 found that the majority of patients described improved sexuality, while Blumer and Walker77 reported that an equal number of patients described no change or improved sexual function.

Surgical removal of epileptogenic tissue by unilateral temporal lobectomy has been reported to abolish pre-existing sexual paraphilias,60–63 cause a decline in libido,74–75 restore normal sexual functioning or occasionally result in hypersexuality.77,55,56,62,68,69,70,74,76–79 The current literature has focused on temporal lobectomy only, with two exceptions.67,77,78 Christianson and colleagues77 targeted both temporal and extratemporal surgery patients (surgery to areas outside of the temporal lobe) but exact details of the final sample were not given, precluding a group comparison. Baird and colleagues78 found that postoperative sexual change was significantly more likely in patients undergoing temporal resection (64%) than extratemporal resection (25%). Temporal patients typically reported a sexual increase to a perceived level of “normal” functioning while a small number described an increase or decrease to a level above or below patient perceived normality.78 In keeping with the right-sided laterality effect, Baird and colleagues78 also found that patients who had undergone right temporal resection were more likely to describe changes in sex drive after surgery than those who had undergone left resection.

Hypersexuality after temporal lobectomy
Hypersexuality, or a dramatic increase in sexual drive, has been reported in some patients after unilateral temporal
lobectomy. Blumer defined hypersexuality as “sexual arousal and response that is clearly abnormal in frequency and intensity for a given individual.”

As previously noted, a number of authors have conceptualised hypersexuality after unilateral temporal lobectomy as a feature of KBS. For example, Blumer identified common traits in three patients, including (1) the manifestation of hypersexuality after a “postoperative silent period” of 3–6 weeks, (2) persistent sexual arousal, (3) homosexual behaviour, (4) accompanying dietary changes and (5) loss of anger or “tameness”. Blumer proposed that temporal lobe epileptic discharges result in an excited limbic system, causing episodic hyperaggressiveness and hypersexuality, whereas efficacious surgery results in depression of the limbic system, associated with decreased irritability or tame behaviour and hypersexuality. He noted that the latter state corresponds to KBS. Walker and Blumer argued that such changes may be due to bitemporal damage. Cogen and colleagues suggested that hypersexuality after unilateral temporal lobectomy represented a “partial KBS” because of the underlying pathology in the contralateral temporal lobe. No other features of KBS were noted in their patient and the time course of this development was much longer than that observed in non-human primates and other human KBS cases which typically manifested within the first year post surgery. Finally, Anson and Kuhlman proposed that postictal dysfunction of the remaining temporal lobes produced a transient functional bilateral temporal lobectomy in their patient. While these two case studies assumed the presence of bitemporal pathology, one case study raised the possibility that a unilateral temporal lesion could produce the presence of bitemporal pathology, one case study raised the possibility that a unilateral temporal lesion could produce a transient functional bilateral temporal lobectomy.

The amygdalae

Findings from animal studies have clearly demonstrated that within the temporal lobe it is the amygdala that is the pivotal structure in mediating sexual behaviour. A recent study has directly examined the role of the amygdala in human sexual functioning. Baird and colleagues compared amygdalar volumes in groups of patients with or without sexual changes following temporal lobe resection for epilepsy and in age matched controls. On average, patients who reported a postoperative sexual increase had a significantly larger amygdalar volume contralateral to the site of their resective surgery than patients with a sexual decrease, no change or controls. There was a significant positive relationship between contralateral amygdalar volume and the maximum degree of sexual change. This suggests that a larger contralateral amygdala may contribute to the expression of increased or improved sexuality after temporal lobe resection. The mechanism underlying this relationship is unclear. One hypothesised mechanism emphasises the role of the amygdala in the processing of emotional stimuli. It has been proposed that the amygdala regulates the attachment of appropriate emotional significance to sensory stimuli and determines reinforcing or discriminative properties of stimuli (see Zald for a review). A larger amygdala may function better in its role as a processor of emotional stimuli, specifically social/sexual cues, and in the attachment of significance to such stimuli. This would increase the likelihood of sexual response, resulting in a sexual increase. Irrespective of the mechanism, this finding points to an important role of the amygdala in regulating human sexual behaviour, specifically sex drive.

Further evidence for the role of the amygdala in human sexual functioning is derived from the observation that stimulation of the amygdala can elicit human sexual responses. Gloor described a “sexual experiential response” during electrical stimulation of the right amygdala in a female patient with epilepsy. On stimulation, the patient described a pleasant feeling in her vulva and inner surface of her thighs as though she were having sexual intercourse. She reported that she experienced the same sexual feeling at the onset of her

### Table 1: Summary of studies reporting postoperative sexual changes and/or symptoms of Klüver Bucy syndrome after amygdalotomy

<table>
<thead>
<tr>
<th>Reference (author, year)</th>
<th>Sample size (n)</th>
<th>Postoperative sexual change (operation type)</th>
<th>Description of sexual change</th>
<th>Postoperative KBS symptoms</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sawa (1954)</td>
<td>11</td>
<td>n = 2 (bilateral)</td>
<td>“Acceleration of sexual impulses” in the form of increased sexual innuendo in conversation</td>
<td>Erroneous recognition of people and objects. Abnormal oral behaviour, changes in emotional behaviour including rage reactions and placidity</td>
</tr>
<tr>
<td>Narabayashi (1963)</td>
<td>60</td>
<td>n = 1 (not reported)</td>
<td>“Slight hypersexuality” lasting only a few days</td>
<td>Nil</td>
</tr>
<tr>
<td>Mark (1972)</td>
<td>10</td>
<td>n = 1 (bilateral)</td>
<td>Impotent for 2 y after surgery</td>
<td>Hyperphagia lasting 3 months</td>
</tr>
<tr>
<td>Kim (1972)</td>
<td>63</td>
<td>Not reported</td>
<td>“Elevation of sexual libido was generally observed”</td>
<td>Elated mood and increased appetite were generally observed</td>
</tr>
<tr>
<td>Heimburger (1978)</td>
<td>58</td>
<td>n = 3 (not reported)</td>
<td>Increased sex drive which lasted from a few days to a few weeks and subsided</td>
<td>Nil</td>
</tr>
<tr>
<td>Jacobson (1986)</td>
<td>1</td>
<td>n = 1 (bilateral amygdalotomy and subcaudate tractotomy)</td>
<td>“A complete absence of sexual contact on the ward or in her personal life contrasted with her description of ‘sexual arousal’ in my presence”</td>
<td>Disorder of facial recognition, emotional changes including placidity, hyperalertness to visual stimuli, motor stereotypes and oral behaviour</td>
</tr>
</tbody>
</table>

ATL, anterior temporal lobectomy; TLE, temporal lobe epilepsy.
spontaneous seizures. Following stimulation, an after discharge involving the amygdala and other limbic structures of the right temporal lobe was observed with subsequent spread to the temporal neocortex.

Stereotaxic human amygdalotomy, or destruction of part of the amygdala, has been performed for the treatment of severe behavioural disturbances such as intractable aggression or hyperactivity. Methodological differences such as patient selection criteria, length of follow up and lateralisation of lesion make cross study comparisons difficult. The surgical procedures used in humans differ from the “amygdalotomy” typically performed in monkeys. Animal studies have used an anteromedial temporal approach and removed the entire temporal lobe structures. Few experimental studies with monkeys have used a stereotaxic approach or aimed to produce amygdalotomies. Patients who have undergone amygdalotomy allow the postoperative effects of precise amygdala destruction to be examined. In particular, the manifestation of hypersexuality and other symptoms of KBS has been reported in several studies. Table 1 gives a summary of these studies. Other studies have found no evidence of altered sexual behaviour or KBS symptoms after amygdalotomy. It is unclear, however, whether researchers specifically questioned patients about sexual changes or relied on self-report or obvious manifestation. Overall, it appears that hypersexuality and other features of KBS are relatively uncommon after selective amygdalotomy in humans. When sexual changes occur, they are typically more subtle than those observed in animals, possibly reflecting the greater complexity of human sexual behaviour.

Further evidence for the integral role of the amygdala in mediating human sexual drive is gained from functional neuroimaging studies which have demonstrated amygdalar activation in response to sexual stimuli. Interestingly, two positron emission tomography studies of sexual arousal and ejaculation found deactivation of temporal areas, including the amygdala. This deactivation was interpreted as an alleviated inhibition allowing for the development of sexual arousal. Hamann and colleagues proposed a differential role for the amygdala in male appetitive versus consummatory sexual responses, as highlighted in previous animal studies. Specifically, consummatory sexual behaviour (erection and orgasm) corresponds with decreased amygdala activity whereas appetitive sexual behaviour (viewing of sexual stimuli) is associated with amygdala activation. Further research is required to examine this proposal.

### BRAIN REGIONS MEDIATING HUMAN SEXUAL BEHAVIOUR

Table 2 shows the key brain regions mediating human sexual behaviour based on a review of the human literature examining the effect of neurological insult on sexual behaviour, and recent functional neuroimaging findings that complement this research. Six main regions, and their associated sexual functions, are identified, including three subcortical and three cortical regions. The subcortical regions comprise: (1) the septal region, involved in the mediation of pleasurable response and orgasm, (2) the hypothalamus, mediating neuroendocrine and autonomic aspects of sexual drive with possible involvement in sexual orientation and (3) the ansa lenticularis and pallidus, implicated in the mediation of sexual drive. The cortical regions comprise: (4) the frontal lobes, mediating the motor components of sexual behaviour and the control of sexual response, (5) the parietal lobes, specifically the paracentral lobule, involved in genital sensation and (6) the temporal lobes, in particular the amygdala, implicated in sexual orientation, sexual drive and disorders of sexual function (e.g. paraphilias).

The neurobiology of sexual behaviour can be conceptualised in terms of the various facets of the human sexual response cycle defined by Kaplan who proposed a triphasic model: (1) sexual desire, (2) excitement and (3) orgasm. Kaplan argued that the three phases are interconnected but governed by separate neuropsychological systems. Associations between the key brain regions we have identified and phases of the sexual response cycle can be made. The current evidence from lesion

<table>
<thead>
<tr>
<th>Key brain regions</th>
<th>Sexual function</th>
<th>Psychosurgery/ neurosurgery/ stimulation</th>
<th>Seizure related</th>
<th>Other: case studies</th>
<th>Neuroimaging</th>
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<tr>
<td>Subcortical</td>
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<tr>
<td>Parietal lobes: paracentral lobule</td>
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studies to date suggests that the initial phase, sexual desire, is mediated by subcortical structures, specifically the hypothalamus, ansa lenticularis and pallidum. The temporal lobes, specifically the amygdala, also play a significant role in this initial phase. Sexual excitement (phase 2) is mediated by cortical structures, namely the parietal and frontal lobes, controlling genital sensation and the motor aspects of sexual response, respectively, sustaining sexual activity until progression to orgasm (phase 3) for which the septal region has been implicated. Rather than a linear progression between phases and underlying mediating neural structures, it is likely that there is an interconnected neural network where various brain regions and associated phases of sexual response are simultaneously activated, producing the physical and psychological manifestations of human sexuality. Further research is required to delineate this complex network.

LIMITATIONS

Our identification of the key brain regions involved in the mediation of human sexual behaviour is primarily based on the findings of single case studies, and the occasional incidental observation. Case studies have a number of methodological constraints, including the validity of extrapolating from single patient observations. Nevertheless, a case study often provides the first insight into a specific phenomenon, and can stimulate further research. A limitation in the study of the behavioural effects of human brain lesions is that with the exception of neurosurgical cases and brain injuries, we must rely on naturally occurring neurological events. Therefore, observations at the single case level often constitute a unique opportunity to investigate a phenomenon, and consequently provide an important step in contributing to our understanding of human brain–behaviour relationships.

There have been very few group studies of changes in sexual behaviour associated with focal cerebral injury or neurosurgery. With the exception of one small scale study of sexual outcome after frontal lobotomy, these studies are entirely limited to temporal lobe conditions, primarily involving patients with temporal lobe epilepsy or patients with psychiatric conditions who have undergone amygdalopectun. Thus there is substantial evidence supporting the significant role of the temporal lobes in mediating human sexual behaviour. In contrast, support for the role of other cortical and subcortical regions relies heavily on case studies. In particular, there have been no group studies of sexual outcome after subcortical lesions. The scarce number of studies to date indicates that large scale and systematic research into the neurological control of human sexual behaviour is long overdue.

There is a paucity of neuroimaging research examining the neural correlates of human sexual behaviour. With the exception of the recent study by Ferretti and colleagues, all studies to date have used visual sexual stimuli that simultaneously activate various regions of the sexual neural network and correspondingly numerous phases of the sexual response cycle. Future neuroimaging research needs to clarify the relationship between specific components of the human sexual response cycle and corresponding neural substrates.

CONCLUSION

We have provided the first synthesis of the literature to date examining the effects of neurological insult on human sexual behaviour, and complementary functional neuroimaging findings. Our understanding of this issue is limited and primarily based on case and small scaled studies. There is, however, considerable convergence among the available research that has enabled the identification of six key brain regions, each mediating specific aspects of human sexual behaviour. Accumulating evidence suggests that the temporal lobes are a critical region in the mediation of human sexual behaviour, with the amygdala playing an integral role in regulating human sexual drive. This review highlights the need for further examination of the neural correlates of this fundamental and universal aspect of human life. It is hoped that this review will stimulate further research, particularly into the key brain regions we have identified to date.

REFERENCES


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53 Mendez MF, Kolarsky A.

54 Hoenig J, Ruff RL, Toone B.

59 Herzog AG, Blumer D.

60 Herzog AG.

64 Hutton EL, Levine J.


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