

Thesis
for
Degree of M. S.

by
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Alcohol
 Its physiological action
 and
 Specific constitutional effect—alcoholism.

There is no agent, diuretic, medicinal or poisonous, that has occupied the attention of scientific minds to the extent that alcohol has, nor is there one on whose action—beneficial or otherwise—there has been so much controversy. Its general and more visible actions on the body in health have long been recognized, and of late years the most eminent authorities in physiology and therapeutics have studied it especially, and many works the result of their experiments & experiences have been published by physicians of all nations. To attempt to add to the observations resulting from such searching inquiry would be presumption; and it is not my intention to consider ^{the question} of alcohol as a food or as a poison or to controvert its use as a stimulant in medicine. In this paper I mean merely to note the action of the alcoholic principle on



Male don

Topical action

principles on
 the healthy tissues of the body generally, and on
 the various organs individually, dwelling more
 minutely on its accumulative toxicological effect;
 and enumerating the diseases which it either directly
 induces or gradually predisposes to.

The conclusions are taken from cases not only in hospital
 and dispensary patients where from co-existing or pre-
 existing constitutional disease and debilitated condition
 of body resulting from various causes, the phenomena are
 greatly modified, but also from cases of patients in
 the higher walks of life.

In (talking of) the action of alcohol, it is unnecessary
 to state that alcohol in a very diluted state, or in
 the form of the various spirituous liquors is meant.

A very small quantity of absolute alcohol or raw proof
 or rectified spirits acts as a corrosive irritant on the parts
 it is directly brought into contact with, and taken in-
 ternally or injected directly into the venous system, it
 either kills instantaneously by paralyzing the nerve
 centres or produces coma and convulsions before death.

First let us look at the action of a small
 quantity of alcohol taken in any of its usual forms
 Locally it is irritant and astringent and remotely

Amulsi actio.

Actio of Langro dom.

and remotely
 it is stimulant; tonic sedative, diuretic and antiseptic
 [Christina class. vol. 1/2]. The topical action is decidedly
 irritant as seen by the reddening of the mucous membrane
 with which it is brought into direct contact. The astringent
 action may be demonstrated by applying it to the skin which
 it first reddens and then constricts. The naked-eye appearance
 of the lining membrane of the stomach of animals treated with
 small doses shows alternate red and white spots. Remotely
 the first effect is also stimulation, which may vary from a
 few minutes to an hour or more according to the strength and
 quantity of the dose. During this period there is often a great
 physical and intellectual excitement shown by the performance
 of most fatiguing exertions with apparently little effort; and
 by rage and witty remarks. The circulation is stimulated
 the heart's action being rendered stronger and more frequent;
 and the nervous system is also stimulated: and this is more
 easily produced when the brain has been excited before-hand.
 This is succeeded in some cases by an after period of
 depression but the general rule is that there is no subsequent
 depression or harm resulting from the greater stimulus of alcoholics.

The general effect of large doses is a more decided excitement
 of the brain and circulation, which is more transient; and is
 immediately followed by a more obvious depression of the

depression of the
 brain first and then of the circulation and consequently
 of the digestion and of every important function of the
 body. This is manifested by confusion of thought; imperfect
 or double vision; indistinct speech & tottering gait; followed
 by speechlessness, incoherence, suffused countenance, injected
 eye with dilated pupil and slow stertorous breathing, most
 of the symptoms being similar to those of narcotic poisoning.

These general phenomena however, which may be observed
 in any case of ordinary drunkenness do not interest us so
 much as the functional derangement of organs and organic
 changes produced in the tissues by the long continued use
 of alcohol, or more properly speaking cases of chronic alcohol
 poisoning. Before this condition is reached, or before there
 is any organic disease the main symptoms found in a
 patient who for a considerable period has become accustomed
 to keep up the stimulating effect of alcohol, are as follows
 loss of appetite, loaded white tongue, persistent dyspepsia
 irregular action of bowels, & flatulency, also inability to
 undergo severe physical exertion or mental ~~fatigue~~, and
 nervous excitability.

All symptoms of chronic alcohol poisoning are brought
 about by the actual presence in the blood throughout
 the entire body, of the alcoholic principle which thus?

Case of acute poisoning

+ Subject was a girl abt. 19 who has swallowed more than a pint of raw whiskey w/out ill. Autopsy made in Edin. police office May /72.

which thus exerts its influence on every part: and (as we shall see) from the fact of its determination to the cerebral plexus it affects in addition the function of the nerves springing from that center. The tendency is for it to be eliminated & this takes place by the lungs, skin & principally by the kidneys. A fit of intoxication is often reversed from during sleep when large quantities of alcohol are thrown out in the breath. In the urine however the greatest amount passes off and it may be got rid of more abruptly by vomiting.

In the only fatal case of acute poisoning by alcohol I remember the post-mortem phenomena usually described were observed. The lining membrane of the stomach was of a dusky red colour, this being confined to that organ alone, and more apparent towards the pyloric orifice beyond which it did not occur. When the skull-cap was removed a decided spirituous odour was felt. There was no hypertrophy of the membranes or inflammatory adhesions but the brain was much congested and there was a large amount of serous fluid below the arachnoid and in the ventricles. That in the lateral ventricles was found by the usual tests to contain a large proportion of alcohol &

Let us now consider specifically the action of alcohol on the tissues of the body and on the various organs, when introduced

Direct absorption.

When introduced
 into the stomach either in a large dose or in repeated
 small doses. It must be borne in mind that liquids
 generally and a liquid like alcohol especially, do not
 undergo the chemical changes to which solids are subjected
 when taken into the stomach. Instead of being subjected to
 the action of the saliva in mastication & the various gastric
 and intestinal juices, in short: undergoing the operation of
 chylification - they pass directly into the blood by a
 simple act of osmosis. There is an anatomical fact
 in reference to the position of the gastric absorbents upon
 which sufficient stress is not laid, and which indeed is
 often overlooked. I would wish to call attention to the relations
 of the lactals and the blood vessels in this locality.

The gastric vessels (from the divisions of the coeliac axis) run along
 the two curvatures of the stomach to supply the coats, & run between
 them. They then divide into small branches in the sub-mucous
 arcolar coat, and pass between the tubuli & ramify on the
 surface of the mucous coat. The absorbents form a network
 between the arcolar muscular coats, & between this layer of
 lactals & the internal surface over the tubuli (gastric or
 peptic according to the difference of their epithelial lining).
 On examining microscopically injected sections of these structures
 we see the capillaries forming minute loops which run up

Action on the Blood

which run up
between the tubuli, and distributed in the mucous lining close
to the surface. I have not met with a preparation de-
monstrating the existence of lactals in this situation, and I
believe no trace of them has been seen. As ^{we have seen} they lie deeper
that is with the whole thickness of the secreting structure between
them and the internal lining of the stomach, and this points
to the greater facility of the passage of alcohol directly into
the blood. This it does and that in a less altered condition
than it would be were it acted on by the secreted juices.

It is then carried by the coronary, superior mesenteric & splenic
veins by the vena porta through the liver to the heart; thence
to the lungs & what is not eliminated is carried back to
the heart from which it is distributed in the arterial blood
to all parts of the body. Thus it circulates comparatively un-
changed in every part and in every organ of the body which
are thus doubly affected by the actual presence of the poison in
them and then an irritated brain which in course of time may become
badly nourished and have its functions impaired.

The action on the blood itself is deteriorating. According to Veitch
and German pathologists generally the development of the red
corpuscles is arrested and their decay hastened. They consequently lose
their vitality & nutritive power. In confirmation of these views I have
examined the blood of persons suffering from debility then excessive

Falt de reactiva
reactiva

the excessive indulgence in alcohol, and I have found a departure from the normal condition of forming into canaliculi, with an increase of white corpuscles. In a case of cirrhosis of the liver as many as 14 colourless corpuscles were to be seen at any time in the field of the microscope, while fatty molecules were to be found ~~in~~ the liquor sanguinis & the corpuscles themselves were fatty degenerated and the alkalinity of the blood was increased. In this condition the blood is less able to take in oxygen & give out ^{carbon} & it is easy to understand that thus it arrests the removal of tissue and retards the carrying away or metamorphosis of waste tissue. Thus we may question if the exhibition of the anti-biptic action of alcohol is desirable since it is degenerate tissue which is maintained.

Its action on the tissues generally. Alcohol tends to produce fatty degeneration of the tissues and this is the most constant result. Not only do we find muscular tissue voluntary and involuntary thus metamorphosed but every tissue and organ is more or less subject to it, brain, heart, liver, kidney &c. This process as is well known is classed under the pathological head of impaired nutrition & consists in the transformation of the albuminous constituents of the tissue into fat. It is in the heart that we most frequently meet with this as caused by alcohol. In appearance a heart thus affected is uniformly paler or has pale spots throughout its substance. In the fresh

Selective & accumulative
action.

on the brain.

To the free?

it is soft and friable & flabby. Microscopically the fibres are seen to have lost their striated appearance, and contain fat granules. (In advanced stages the sarcolemma is free of large molecules of fat and oil globules). In the kidney & liver it first makes its appearance as small granules in the cells, then the nucleus and cell-wall are destroyed. In the brain it is met with in the condition commonly called "Edema of the Brain" & in the arteries it is also met with as a rule beginning in the connective tissue cells of the intimal coat.

Like many medicinal remedies which not only act thru' one particular channel but are determined to one particular locale alcohol has an inherent power of selecting a particular organ and tissue for the seat of its special operation and it is now well known that it fixes on cerebral matter by this peculiar selective power. This has been found by French pathologists to be twice as great for the brain as that for the liver where next in order it is found to accumulate. A post-mortem examination of the brain of persons used to a long continued over-indulgence in alcoholics shows a congested state of the capillaries of the cerebral substance & of the vessels of the meninges. These vessels when separatedly brought into this condition not being able to rid themselves of the engorgement in the usual way, are relieved by serous exudation into the areolar spaces below the arachnoid membrane and into the cerebral ventricles. This naturally sets up

Alcoholism

sets up

a great amount of cerebral irritation & consequently all the important functions of the body are interfered with. In this congestive stage apoplexy and epilepsy may occur & if it passes on to the inflammatory stage, delirium tremens, convulsions, shaking palsy, paralysis are likely to result.

As small doses long continued, of certain agents e.g Lead, Arsenic or Mercury produce at first no noxious but even a beneficial result & ultimately a condition of plumbism, arsenicism or mercurialism, so oft-repeated indulgence in alcoholic drinks produces an action which because it is brought on by nothing else is specific, and the resulting constitutional condition is called alcoholism.

The actual presence of alcohol in the brain has been referred to in a case of acute poisoning. Now precisely the same localization takes place when it is used in small doses, continuously and repeatedly. As in acute cases it selects brain, so does it in chronic. It is not entirely eliminated from the system but a certain amount remains behind & that chiefly in the cerebral pulp where each fresh dose produces a like accumulation. This poisoning of the grey matter of the brain develops a constitutional condition invariably similar in every fresh case and attended by phenomena of uniform character & course. There is a fact confirmatory to the view of its being accumulative. There is no drug whose action is so much modified by habit. A person habituated to its use soon becomes very much less susceptible to its action so that we would

Alcoholism

we would

expect him to become in time insensible to its stimulating or rather exciting effect; and this is the case. He may go on for a long period gradually increasing his daily allowance with no proportionate effect - no apparent evil result; until a certain stage is reached. This is when it has accumulated to the extent that his whole system is charged with it - until in fact he is alcoholized. When this stage is reached, a very much less quantity will produce an equal or even greater result than the larger amount previously indulged in. That the whole system is as it were saturated with the poison is manifest from the familiar example of the obstinacy to heal of the most trifling wound or even abrasion of the skin where this exists in a person thus alcoholized & the irritable nature of any pre-existing sore or ulcer in what ever part of the body. Every house surgeon knows how liable scalp wounds in drunkards are to take on suppurative action. Proof may also be taken from the dangerous & acute character which inflammatory diseases assume & the great liability such patients have to succumb to attacks of fever &c. Having then determined that alcohol by acting on and accumulating in the cerebral tissue, and by disturbing the functions of the sensory, produces such a condition of system that a smaller quantity will produce a more intense and lasting result, let us consider the more important diseases induced by this condition and first in rank comes Delirium Tremens.

Belgium truncus

This disease may be defined as one of the manifestations of the specific toxicological effect produced by long continued and oft repeated doses of alcohol. It is brought on solely by over indulgence in alcoholic drinks which have been shown by Drs Craigie & Priddy to be not only the pre-disposing but the directly exciting cause. After the able writings of such authors little remains to be added to their observations except to confirm from cases in the pathological theatre that the primary lesion is in the cerebrum itself, & that this organ is not sympathetically affected thru' the solar plexus by the gastric disorder.

The preliminary symptoms are digestive derangement - foul tongue, & constipation - moist skin, scanty urine, & pulse high & small. There is also restlessness & a nervous irritability of manner, with a cessation of the desire to drink. The symptoms which follow are well known and are sleeplessness, muscular tremors of the hands & tongue, and twitching of the facial muscles, and the delirium which is characteristic being not violent but busy & abrupt & apprehensive. The hallucinations of sight & hearing are also constant and the anxious expression of face is marked. Dr Craigie considers that these phenomena are due to the condition of the blood sent to ^{the brain} & directed in the meningeal & cerebral vessels. There as we have seen are full of alcoholic particles & the blood is imperfectly aerated. This is no doubt the pathological condition during the preliminary & first stage but an opportunity of examination at this time is seldom afforded. In post-mortem examination of

Pathology of delirium tremens

Delirium tremens

examination of fatal cases where the characteristic delirium & hallucinations have supervened we naturally expect to find something more than signs of pure cerebral irritation. In such I have observed hypertrophy of the meninges especially of the arachnoid with inflammatory adhesion to the other membranes of the dura mater to the skull & of the pia mater to the brain-substance. In some cases in attempting to strip this last from the convolutions, portions of the cerebral pulp came away. The cerebrum itself is usually soft & friable & the convolutions are shrunken. These conditions along with opacity and fibrous character of the membranes we would only expect to notice in cases where history showed excessive drinking extending over a long period. The effusion is not only sub-arachnoid and into the sulci but infiltrates into the cerebral substance itself.

The theory that the sudden cessation from the usual stimulus was the directly exciting cause of delirium tremens has exploded. It arose from the fact that one of the preliminary symptoms is the gastric derangement and an aversion to drink is produced. On account of this the subject lies for some time previous to the exhibition of the more apparent symptoms, entirely given up his usual diet.

Delirium ebriosum is another form of disease to which the abuse of alcohol may lead. It is caused by heavy drinking for a short period until a sort of mania is produced. When this is manifested the desire for more strong drink is so strong as to lead the patient

Dilecium traumaticum

And the patient
 to perform dangerous & violent acts in order to obtain it.
 The delirium is wild and raging, muscular tremors are usually
 present. The skin is dry & hot & the pulse is strong & high. These
 symptoms are popularly supposed to be those of delirium tremens
 but they are quite characteristic of this form of mania as we
 may call it which also differs in pathology & treatment from delirium
 tremens. In this the patient must be put under restraint while
 in delirium tremens restraint must not be employed as it increases
 the irritation & may induce a tendency to violence. The expression
 of face in each is distinctive. In the one it is anxious with a
 look of expectant dread, while in the other it is sullen & fierce.
 The pathological condition can only be guessed at - the symptoms
 point to a great amount of vascular excitement - but the cerebrum
 is not so full of alcohol as in delirium tremens, in ~~fact~~ ^{fact} ~~the~~
 the state of alcoholism is not developed. In this form as
 in delirium ^{tremens} nature tends to produce recovery. The strong craving
 for more drink when yielded to produces nausea & vomiting
 after which rapid recovery takes place. In delirium tremens as we
 have seen there is no craving for drink for more would
 aggravate the symptoms & prolong recovery, thus we are justified
 to treat without stimulants.

A febrile attack with delirium of a low type is sometimes in-
 duced as the result of ^{Surgical} injury to a patient who has been

Action in the stomach

who has been

in a state of alcoholism at the time. This is the delirium
traumaticum of Dupuytren. I have not met with a well defined case.

It would be a tedious task to enter into the symptoms
and pathology of the many diseases resulting from the action
on the nervous centre of alcohol. Suffice it to say that among
the most frequent are apoplexy, epilepsy, the various cerebral
softenings (giving rise to paralysis), convulsions, shaking palsy & insanity.

Space does not admit of the consideration of the diseases
in the various organs induced by the abuse of alcohol, for each
is a study in itself. I shall only mention some of the most
important met with, and first those of the stomach.

From the irritant action of alcohol and from the fact that
drum-drinkers frequently indulge on an empty stomach it is easily
seen that a certain amount of inflammation will be set
up after each dose. Dr. Braumont's experiments on St. Martin
showed that the contact of alcohol with the walls of the
stomach was followed by an increased secretion of fluid
which he found was not gastric juice but the ordinary
mucus. This direct irritation with congestion and stimulation
to increased secretion results in chronic gastritis which
soon produces thickening and induration of the coats.

This I have noticed with narrowing of the
pylorus but never to the extent of ulceration and

Action on the liver

Cirrhosis.

Pathology of Cirrhosis.

ulceration & perforation as is sometimes described, nor have I seen a case of gastric scirrhus as resulting from this directly exciting cause.

The liver as we have noticed may, in common with the tissues of other organs, become fatally degenerated one of the results of impaired nutrition. But there is an affection of a chronic nature, met with in this organ, which must not be passed over. I refer to Cirrhosis.

This is, in short, a chronic inflammation of the areolar tissue in the interstices of and covering the organ. We have seen that alcohol is directly absorbed and is carried by the portal vein thro' the liver. This I am inclined to agree with those who hold that this affection is due to the direct irritation of alcohol in the liver. The pathology is interesting.

At first the organ is congested & consequently enlarged. This congestion occurring intermittently, alternately compresses and relaxes the hepatic lobules. The interlobular connective tissue being thus subjected to non-continuous irritation and pressure becomes hypertrophied & consequently contracts drawing the capsular ligament with it so that the surface is irregular, and hence the term "hob-nailed" liver. At this stage the whole

Case of cirrhosis of liver

the whole

organ becomes contracted and smaller. Many important secondary symptoms result from the contracted condition & obstruction to the circulation, such as dropsy of the limbs, ascites, enlargement of the spleen &c.

A well-marked case in Hospital was for some time under my care. Patient was an Englishman in a brewery and was in the daily habit of imbibing large quantities of sweet ale and that generally when there was no food on the stomach and so he was braced and propped up so that there was every facility for the alcohol being rapidly absorbed and exerting its influence on the tissues & on the liver especially. His first symptom was failing eye-sight immediately followed by yellowness of the conjunctivae and other indications of jaundice. Some days after this he complained of pain in the right hypochondrium. On admission the liver was found to be enlarged, & painful on palpation. The deep dullness in the mamillary line was six & a half in. & there was considerable ascites, the circumference of the abdomen at the umbilicus being forty-two inches; & the patient also suffered from nausea & vomiting. The interesting features of the case were the secondary affections, as the dropsy of the abdomen & the limbs and the enlargement of

enlargement of the spleen. Some weeks after admission when the congestion of the liver had been subdued and the organ resumed its normal dimensions was about $5\frac{7}{8}$ in. by the external application of iodine and other means, the ascites increased considerably. This was undoubtedly owing to the circulation being more obstructed from hypertrophy of the connective tissue & compression of the vessels and cells. The spleen also was found to be enlarged to about $5\frac{3}{4}$ in. from the upper to the lower end, and there was a marked increase in the number of the pale Flavel corpuscles. As many as from 14 to 16 of these columnar discs were to be seen at a time in the field of the microscope.

The remaining disorders affecting the digestive system whose origin may be traced to the abuse of alcohol are numerous. At first dyspepsia & constipation result, the latter owing to diminished secretion and increased alkalinity of the blood, but later if ulceration of the stomach or intestine is present purging is the result. In the kidney we find fatty degeneration, & as a rule this begins in the capillaries of the Malpighian bodies. In the urine albumen & phosphates exist to a great extent and.

great extent; and
diabetes is not uncommonly met with.

In short, by the long continued use of alcohol the nutrition of the whole system is impaired and the vitality is consequently lowered, so that before any organic lesion exists there is great functional derangement.

In this state of things is as it were any demand for increased exertion on the part of an organ it is unable to perform its function or the performance of it is followed by organic disease.

I shall conclude with a quotation from "Prædicæ" on the treatment of Delirium Tremens without stimulants or opiates.
"The habitual & excessive use of alcoholic liquors however does not affect all individuals alike. Some drinkers are rendered by disease of the heart-liver or other organ to which they may have a hereditary or constitutional liability; others in fever or inflammation which they have no stamina to contend against; others by apoplexy or paralysis from the direct effects of a drunkard; & some from hereditary predisposition or otherwise are rendered to spend the remaining years of a miserable existence in mania, idioty or in a general paralytic condition of the system. Some few drinkers again by reason of extra ordinary constitutional vigour escape all these ills and live on to old age."

but the greater number who are not-rarely removed from society by the diseases transmitted suffer more or less from attacks of delirium tremens."

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University of Edinburgh 9. 1878.

Edinburgh 1 April 1878.