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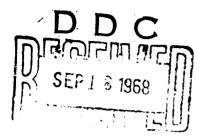
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> DEPARTMENT OF THE ARMY Fort Detrick Frederick, Maryland

THE RICHETION OF URINE CONPONENTS DURING FEVER ATTACKS

Varchow's Archiv. Vol 155, 1899, pp 1-43 W. Von Morascevski.

It is one of the most well-known facts that during fever the N-excretion is increased, whereas chlorine is retained by the body. Phosphates and potassium salts are to be added to nitrogen, and sodium salts are to be added to chlorine. The most beautiful and the best studies have led to these results; for instance, the re-sarob of Traube and Jochmann (1). Redtenbacher (2), Ringer (3), Brattler (4), Easthausen (5), Rosenstein (6) Riesenfeld (7), Unruh (8), Huppert (9), Schleich (10), E. Hallervorden (11), Salkowsky (12), Naunyn (13, 14), Zulter (15) Rohmann (16), v. Limbeck (17), Senator (18), v. Terrey (19), Ideburneister (20), Leyden (21), S. Rem Pirei and v. Calcini (22), and many others. In the oldest works by Moos, Uhle, Ranke, Wasmuth, Bartels, and in the above-mentioned works which are not complete by far, rules are formulated, in most cases, on the basis of a longer observation period, an observation running for days and not for hours (with the exception of the work by Naunyn). The experimental material which included almost all urine components provided therefore a characteristic for the stage of the fever rather than for the beginning and the end of the fever.

Only the most recent observations made in the "shortened" fever -malaria -- provided new and surprising facts. Also the present experiment has been stimulated by the work of my collesgue and friend Dr. Theodor Hitzig. The abnormal behavior of the chlorides and of phosphorus in the case of malaria which is in great contrast to the findings of the fever metabolism (v. Limbenk, Terray) drew my attention to other types of fevers and stimulated me to repeat the old observations and to carry out independent experiments. To be sure, it had also been observed by the previous workers that there is an increase in N-elimination after the fever (Naunyn) or that there is a tremendous drop in the elimination of potassium selts after the fever (Salkowsky), or that amonia is eliminated in increased amounts during the fever and is even more strongly eliminated after the fever (Hallarvorden). There was also sometimes an increase in the retention of chlorides after a fever (Kinger), as well as an increase in phosphoric acid (v. Eaxibausen) in the urime.

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This all was known and our work cannot contribute many new data. In our work, the types of fevers are compared with respect to the behavior of the salts prior to, during, and after the fever, the results are summarized and interpreted from that point of view which, on the basis of the previous observations, appears to be most appropriate.

The material comes from the clinic of Professor Dr. Mermann Eichhorst and I am indebted to him for his cooperation and continuous support; I am also grateful to his assistant Dr. Th. Hitsig in whose department the patients were located and whose friendly cooperation has contributed much to the completion of this work.

The chemical methods which I have used have been frequently described in my earlier papers and I can limit myself to refer to these papers (<u>Virchow's</u> <u>Archiv</u>, Vals 146, 150, et ceters, <u>Zeitschrift für klinische Medizin</u>, Vals 34, 35).

The determination of potassium and sodium was carried out according to Fresenius' well-known method: fussion of 50 cc urine with ammonium sulfate to transform potassium and sodium into the non-volatile sulfates, treatment of the fusion residue with barine chloride and b rium hydrate to obtain sodium and potassium chlorides, removal of the excess barium with ammonium carbonate, driving off of the excess ammonium salts, and weighing of the residue as sodium chloride and potassium chloride in a platinum crucible.

Potassium was subsequently precipitated with platinum chloride from the residue, washed with alcohol and ether, dissolved in boiling watar, the solution evaporated, and the residue weighed as potassium platinum chloride.

The amount of sodium was calculated from the found amount of potassium by subtracting the potassium chloride from the sum of both chlorides.

Chlorine was determined a cording to the Volhard method, phosphorus and calcium according to Neubauer, urea according to Mörner, uric acid according to Ludwig-Salkowsky, ameonia according to Schlösing, the alloxuric bases according to the method which I have described in <u>Virchow's Archiv</u>. Vol 150*.

The same holds for the diet and the determination of the nutrition value of the food. The food analyses which The Hitsig carried out for his work confirmed our previous studies, and they provide sufficient assurance for uniform preparation of the food, as well as for reliability of the methods.

The patients were under reliable supervision and were, if necessary, instructed by us so that the examination which has to be done with extreme

* The new work of Salkowsky in Pfglüger's Archiv 1898 was not yet known.

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care was simplified. For instance, the separation of the urine from the feces was always very carefully observed; also, punctual emptying of the bladder was emphasized. The temperature was measured by emperienced nurses.

We observed two cases of Febris continue with a gradual increase and a gradual decrease in temperature (typhus), one case of a fever involving a crisis (pneumonia), one case of malaria fever, and one case of heotic fever. We did not study other types of fevers during acute exanthemas, articular rheumatism, et cetera because it had been determined by earlier observations that such cases give less clear-out results and are subject to individual fluctuations.

To obtain as alsor a picture as possible, we have carried cut uses determinations, the determination of the total nitrogen, of unic acid, of xanthine bases, [and] of ammonia. In addition, the chlorides, phosphates, potassium, sodium, and calcium ware determined.

In the pneumonia case, the urine was collected until the orisis and after the crisis; in the case of the malaric fever, five portions were collected which corresponded 1) to the cold stage, 2) to the climax of the fever, 3) to the remission, 4) to the additional drop in the fever, [and] 5) to the apyrexia. In the case of Febris hectica, four portions corresponding to the stages 1) of the beginning fever, 2) of the climax of the fever, 3) of the remission, 4) of the apyrexia were collected. In the case of typhus I, the urine was collected as daytime urine and nightime urine during three periods: 1) the climax of the fever, 2) the paracme, [and] 3) the afebrile period. In the case of typhus II, the entire amount of -24 hours was examined, without interruption of the entire course comprising the periods 1) of the increase, 2) of the continuous fever, [and] 3) of the apyrexis. Details will be discussed in each individual case.

The compilation of the tables differed, depending on the cases. The hourly amounts were emphasized in the cases of malaria and heatic fever; on the other hand, the daily balance was less emphasized because the feces could not be considered. In the typhus and pheumonia cases, the balance was also accurately determined in addition to the hourly amounts. In cases in which the determination of the hourly amounts appeared to be without value, it has been omitted from the Tables.

The cases cited here are to be regarded as average cases of repeated observations which we have collected in the clinic and a report [on all these cases] would needlessly expand this paper

Case I. B.O., 17 years. Typhus abdominalis.

Case history: patient comes from a healthy family. He suffered from measles as a child, but claims to have been always healthy since then. Ten days ago he became sick with headaches, loss of appetite, and diarrhea.

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Prosent state: nothing abnormal discovered at the external and internal organs. Hild rescale, spleen moderately enlarged.

The patient has been put on a dist consisting of 1,200 cc milk, 800 cc broth with egg, 500-800 cc wine.

This diet contained 10,340 gram nitrogen per day 6,246 gram chlorine per day 1,260 gram phospharus per day 1,740 gram calcium per day 2,748 gram potassium per day 3,562 gram sodium per day

and has been maintained during the entire experimental series.

The foces were not considered because the examination days had been picked at random from the individual periods. The fact that the foces were not analysed introduces errors pertaining only to calcium. The amount of the other components is little affected by deforation.

The urine was collected in daytime and nighttime portions for three days. The first day was during the fever period.

Temperature during 24 hours	38.6	39.5	38.1	38.5
pulsation	106	108	104	100

The patient was bathed once a day; the temperature of the water was 30°.

The second day of urine examination was in the stage of decreasing temperature.

Temperature on the day of				
examination	37.2	36.8	38.2	38.1
pulsation	76	80	92	88

The third day belongs to the fever-free period.

Temperature	36.4	36.6	36.0	36.5
pulsation	76	63	68	76

This examination was supposed to indicate the typhus of urine during the fever period and of the urine from the afebrile periods. The musbers found were calculated for the hou-ly eliminations.

The balance was made without consideration of the feces. Table I. B.O., 17 years of age. Typhus abdominalis.

Temperature	38.6	39.5	38.1	38.5	alimax of the
-					forer
Pulsation	106	168	104	100	

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From 6 AH to 6 PH

Amount of urine 1,300 cc. specific weight 1015.

N	1.001	\$	per	12	hours	10.010	rer	hour	1.084
C1	0.212	\$	*		6	2.756	•		0.229
4	0.077	\$				1.141			0.095
Ca	0.0082	\$				0.106	N	ť	0.009
<u>K</u>	0.159	\$	A		E	2.067			0.172
Dia	0.147	4				1.937			0.161
uroa	0.840	\$				10.920			0.910
uric acid	0.914	\$				0.182	•	•	0.015
allox.								•	
bases	0.018	\$ (oalor	a- *			0.234	8	q	0.019
ranthine		flate	_						
bases	0.004	SAS N	6			0.054			0.064
NH3	0.057	ち			N	0.742			0.062

From 6 PH to 6 AN

Amount of urine 590 cc. specific weight 1016

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N	1.218	\$		2 hours	7.186	per hour	0.599 g
C1	0.206	\$	- ŭ 🙀	#	1.215	* * *	1.101 g
P	0.1177	%	F H		0.694	H H	0.058 g
Ca	0.0058	5	B #		0.034	* *	0.003 g
K	0.183	\$			0.079	F	0.090 g
Na	0.116	Ś.			0.684	# #	0.057 8
ures	0+984	\$			306		0.484 g
uric acid allox	0.012	\$	* *	•	0.0708		0.006 g
bases	0.021	% Calcu			0.177	* *	0.015 g
ranthine		as h					
bases	0.009	\$		8	0.0531		0.004 g
^{NH} 3	0.075	り		8	0.466	• •	0.040 g

Result

Food	Urins	Balance
N 10.340	20.199	- 9.859 g M
C1 6.246	3.971	+ 2.275 g C1
P 1.266	1.835	- 0.575 8 P
Ca 1.740	0.140	+ 1.600 g Ca
£ 2.748	3.146	- 0.398 g K
Na 3.562	2.621	+ 0.941 g Na

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16.726 urea N

0.253 uric acid N

0.411 alloxuria bases N

0.158 xanthine bases N

1.208 amaonia N

Note: When possible losses in the feces are included, the result becomes more pronounced but it remains essentially unchanged. N, also P, may be excreted ..., still larger amounts, calcium may be retained to a lesser extent. Chlorine, potassium, and sodium remain essentially unchanged because the amount of feces is very small.

Remission

Temperature	37.2	36.8	38.2	38.1
Pulsation	76	80	92	59

From 6 AH to 6 PM

Amount of urine 510 cc. specific weight 1013

N	0.7945	۶	per	12	hours	4.052	per	hour	v. 338
Cl	0.2548	۶	• ••			1.299	•	U	0.108
P	0.0594	%				0.3029			0.025
Ca	0.0055	۶				0.0281			0.002
ĸ	0.1170	۶				0.5967			0.049
ha	0.167	\$				0.8517			0.071
uros	0.672	\$			#	3.427		8	0.289
uric acid	0.0098	\$				0.0499		8	0.004
allox.									
bases	0.0126	\$	#			0.0643			0.005
zanthine			·						
bases	0.0023	\$				0.0144			0.001
NE3	0.0434	*				0.2213	8		0.0188

From 6 FM to 6 AM

Amount of urine 380 on specific weight 1020

N	1.330	\$	per	12	hours	5.504		hour	0.421
Cl	0.309	7			-	1.295	N	•	0.108
P	0.1566	9		#		0.710			0.060
Ca	0.0075	×.	ĸ		#	0.0285			0.002
x	0.1877	\$	#		•	0.713		1 8	0.060
Na	0.2126	\$			W	0.806		•	0.065
urea	1.176	۶		F		4.474			0.373
uric acid allox	0.017	\$				0.9646	•	•	0.005
bases xanthine	0.0224	۶	•			0.851		•	0.007
bases	0.0054	\$			R.	0.0205			0.002
11H3	0.0616	\$	•	#	t:	0.2341	•	2	0.020

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	Food	Urine		Result
N	10.349	9.106		+ 1.234 or
Cl	6.246	2.594		+ 3.652
P	1.260	1.013		+ 0.247 or
Ca	1.740	0.056		+ 1.684
K	2.748	1.319		+ 1.429
Na	3.562	1.657		+ 1.905
		urez uric soid alloxuric bases xanthine bases Ammonia	7.901 0.1145 0.1494 0.0349 0.4554	calculated as N

Note: The results may be substantially changed when the losses in the feces are included. Thus, a loss in N may occur after a loss in phosphorus; calcium may be retained less. The other quantities remain the same. The

Afebrile Period

Temperature	36.4	36.6	36.0	36.5
Puisation	76	68	68	76

From 6 AN to 6 PM

Amount of urine 1,200 cc, specific weight 1013.

N	0.7945	\$	Der	12	hours	9.534	Der	hour	0.794
Cl	0.3944	Ś		_#		4.733	. .		0.394
P	0.0454	5	R		N	0.545			0.045
Ca	0.0090	%				0.108		A	0.009
K	0.1990	\$				2.388			0.1990
Ne	0.2040	<u>с</u>	2	=		2.448		#	0.2040
urea	0.680	\$		N		8.160			0.680
uric said		\$				0.108			0.009
allox. Dases	9.0136	*				0.163			0.0136
xanthine bases	0.0046	\$				0.053	£		0.0046
NH3	0.0420	\$	*			0.504	۲	•	0.042
			Fr	Citiz	6 PM to	6 AM			
N	1.155	۶	Têr	12	hours	6.468	Der	hour	0.539
C1	0.4368	\$	· •		#	2.445	•		0.204
P	0.0864	¥		-		0.484			0.401
<u>Ca</u>	0.0664	۶	ũ			0.0358		8	0.003

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K Na	0.297 0.209	****	per #	12	hours #	1.663 1.171	per H	hour	0.139 0.099
uree	0.966	Ś				5.409			0.451
uric acid allox.		۶			•	0.0784			0.006
bases Manthine	0.0189	۶	•			0.1058	*	R	0.009
bases	0.0049	۶				0.0274		۳	0.003
NH3	0.0616	\$			•	0.3449	•	•	0.029
Food				Ur	ine				Result
N 10.34				16.					- 5.662
C1 6.24					178				- 0.932
P 1.26					029				+ 0.231
Ca 1.74					144				+ 1.596
X 2.74					051				- 1.305
Na 3.56	2			3.	619				- 0.057
			urea urio eci alloxuri xanthine amnonia	o b		13.569 0.186 0.269 0.083 0.849	calcul	ated	as N

Note: The result cannot be substantially changed when the components in the foces are considered. Nitrogen would be eliminated to an increased extent, phosphorus would perhaps remain in equilibrium, calcium would be less retained. The other components were all excreted to an increased extent. Again, the behavior of K is striking because its excretion is considerably increased.

This case was supposed to provide us with information about the be-havior of the salts during a fever spell. When the results of the first day (climax of the fever) are compared to those of the last day, a rather typical picture is obtained. In the former case, chloride and sodium are retained. phosphorus is lost; in the latter case, chlorine and sodium are lost, phosphones is retained - exactly the reverse [of the former case]. The behavior of the nitrogen and the potassium salts must not lead us astray. This behavior is subject to other laws, as is known, the nature of which we have to discuss. The period of the remission exhibits a real transition between both periods when the correction for the noninclusion of the feces components is made.

As far as the nitrogen-containing components of urine are concerned, they also behaved quite typically: during the fever, there was an increased amount of urio acid, xanthine bases and alloxurio bases, in general, (as calculated sum of the santhine bases and unic acid) an increased excretion of Amonia.

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fever urea 0.83; urio acid 0.012; alloxuric bases (020; NH 3 0.000 N N

Afebrile

period " 0.85 " 0.011 " 0.017 " 0.053

Also Zülser's ratio P/N was typical

fever P/N 0.0917 fever-free stage P/N 0.064

Finally, when the ratio of N in the other components in the food and in the urins is formed, the following results for the fever stage:

N: Cl:	Pı	Ca:	K:	Na i		
1:0.62	0.12	0.17	0.27	0.35	in the food	
1:0.198	0.0917	(0.007*)	0.157	0.131	in the elimination	

For the remission period

N :	Cl:	P:	Cai	K 1	Nat
1:	0.286	0.111	0.006	0.146	0.182

For the afebrile period

N: Cl:	P:	Ca:	<u>K:</u>	Na:	on
1:0.62	0.12	0.17	0.27	0.35 in the food	
1:0.448	0.064	(0.009)	0.253	0.226 in the eliminati	
	N :	ursa	uric acid	allomuric bases	ĵ⊞۸
climax of		0.836	0.0126	0.0205	0.0604
remission		0.868	0.0125	0.0164	0.0500
afebrile p		0.878	0.0084	0.0168	0.0530

It is seen that: 1. none of the urine components are excreted to such an extent as the nitrogen components. 2. the favor-free period is characterized by an improvement in the ratio of the chlorides to the potassium salts.

This for the typical behavior during fever. In case II which is also a typhus case, we can study the changes with respect to the increase in body temperature.

Case II ZA., 32 years. Typhus abdominalis.

Case history does not disclose anything which would be of importance. Patient comes from a healthy family; he has been completely healthy until now. The present disease started 12 days ago. Patient lost sleep and appetite, did

* Only in urine

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not feel well and had himself admitted to the hospital to obtain better food.

Present status: No essential changes of the internal and external organs. Nothing can be discovered except for a distinct swelling of the spleen.

The patient received the same diet as Case I: 1,200 cc milk, 800 [cc] broth with egg, and 1/2 liter of wine. This diet was kept unchanged until the end of the experiment. As in all other cases, the ancunt was measured. All foodstuffs were analyzed by us.

The patient was examined by us for the components under consideration and the amount of the components per day was calculated by calculating the total amount from the percentages, and this total amount was divided by the number of days.

The urine was measured daily and examined daily. The daily loss of N. Cl. P. Ca. K. Na and NH₃ and other nitrogen compounds was determined from the urine. All other figures were calculated as Cl. Na. K. P. Ca (not as P_2O_5 , NaCl, et cetera). The nitrogen compounds were all recalculated in terms of nitrogen, obtaining, thus, the amount of nitrogen excreted as NH₃, uric soid, alloxuric bases (uric soid and xanthine), and xanthine.

At the beginning of the experiment, the temperature was ascanding, reached its maximum, and then dropped to normal. The effect of the increase and of the drop was thereby observed.

13 July	7 38.9	38.7	39-5	39.2	Temperature
	88	90	100	96	Pulsation
14 July	7 39.0	39.5	39.0	38.6	Temperature
	92	100	96	92	Pulsation
15 July	7 37.8	38.4	38.5	39.0	Temperature
-	100	96	100	92	Fulsation
16 July	7 38.5	39-5	39.9	39.6	Temperature
	92	92	96	96	Pulsation
17 July	y 38.6	39.6	38.8	37.5	Temperature
_	89	92	96	84	Pulsation
18 July	7 37.5	38.1	39.5	39.0	Temperature
-	84	96	100	92	Pulsation
19 Jul;	7 37.6	39.4	40.1	39.0	Temperature
-	92	96	100	101	Pulsation
20 Jul	r 37.6	37.2	37.2	38.2	Temperature
	1.00	100	96	3.00	Pulsation
21 Jul	y 36.5	37.2	39.3	38.9	Temperature
	96	95	96	112	Pulsation
22 Jul	y <u>3</u> 8.9	37.5	37.8	38.5	Temperature
	104	101	101	86	Pulsation
23 Jul;	y 37.0	37.1	38.3	38.5	Temperature
	92	88	100	96	Pulsation
24 Jul;		37.3	38.2	38.3	Temperature
	84	98	100	92	Pulsation

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Table II. Typhus abdominalis

14 July temperature 39.1, amount of urine 800 cc. specific weight 1026

B 2.289 \$ 18.31% per day 0.9267 in the feces 10.300 Ci 0.2467 1.1600 0.4669 in the feces 10.300 Ci 0.2467 0.1460 0.4669 in the feces 10.300 Ci 0.0035 0.1793 0.1900 1.740 1.740 Ci 0.0035 0.1793 0.1793 1.1600 1.740 Lin 0.0035 0.1792 0.1792 1.1740 1.740 Lin 0.0035 0.1792 14.4480 1.1740 1.740 Line 0.0289 0.1792 14.4480 1.1740 2.748 Line 0.0289 0.2512 14.4480 1.1740 2.748 Line 0.0289 0.2512 14.4480 1.1740 2.748 Line 0.0289 0.2512 14.4480 1.1740 2.748 Line 0.0289 0.2724 14.500 1.1790 1.1740 Line 0.009355 0.07224 14.500 1.1790 1.1279 Li 1.10750 1.1625 <th>the feces 10.340 in the food -8.499 6.246 =</th>	the feces 10.340 in the food -8.499 6.246 =
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- 11 -

16 July temperature 39.4, amount of urine 1,220 co, spec. weight 1026

•

			•					•	•					
725		20.2775	per	day	0.5267	4	tàe	feces	OHE OT	Lt.	tte	food		-10.465
6		5.3289			0.4669	e		×	6,246	Ħ	-	4	+	154.0
Эл	0.0860 \$	1.0492	a	2	5192.0	*		=	1,260	•	3		(~ 0.080
2		0.0027		8	0,1901	8	R	*	1.740	æ	•		+	L.457
		0.0150	×		0.3795	x	×	•	2.746	•	-	*	+	:353
	A COLL	1.8215			0.1683	*	8		3,562	8	*	-	+	1.572
		10.510		5										
urio acid	0.0210 \$	0.2562												
Alloxuri.o			I	I										
bases	0.0298 %	0.3635	k											
beses	0.0088 \$	0,1073												
E E M	0.0875 %	1.0075	E	t										
	17 July temperature 32.6 amount of urine 700 cc. spec. weight 1030	persture 38	n e 3.	aount	of urine	<u>8</u> 2	20	spec.	weight .	1030				
	1.967 \$	13 . 769	Der	day	0.5267	ន	٩ ٩	feces	10.340	ł	the	food	ግ	-3.956
10	0.2123 \$	1.486		•	0.4669		2	=	6"246	8	*	43	7	°292
1 2	0.1520 %	1.064	=	8	0.2912	*	2	-	1.260			r	9	-0°02
, e	0.0101 \$	0.1337	E		0.1901	=			1.740	æ	a		Ŧ	419
l ha	S the line	0.7938	æ	=	0.3795	*		10	2.748		8	#	ţ	°575
, and a second	0.0664 4	0.4649	=	*	0.1683	*	R		3.5	8	8	•	ţ.	.929
nrea	1.776 \$	12.150		¥										
urlo solà	0.02345%	1491.0	=	2										
allemnte			:											
D266	0°031748	0,2221	•	2										
zanthine		0820 0	=											
D4.5 05			: 1											
			:											

- 12 -

0.0580 0.6804

0.00829% 0.0972 \$

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Construction of the second sec

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18 July temperature 38.5, amount of urine 800 cc, spec. weight 1026

ţ.	2.037 \$	1.6.296	ted	ber day	0.5267	ų	the	in the fecas	046.01	4	the	food	-6.483
. 6		2.427	8	a	0.4669	8	**		6.246	•	¥	•	+3.353
5 A	A 000 L C	1.512		8	0.2912	-	8	•	1.260	=	•	u	-
4 C	0 0288 4	0.2300	=	8	1061-0	*	#		1.240	=	æ	•	+1.320
د د	2 0.80 C		*		0.3795	•			2.748	٠	#	¥	+1.654
4 74	0.0785 %	0.6280		-	0.1683	•	2	u	3.562	8	٠	•	+2.766
urea.	1.708 \$	13.6640	*	•			÷						
uric sold	0.0297 \$	6.2380											
alloxuric bases	0.0389 \$	0.3115		z									
xanthine bases	0.0092 \$	0.0735		*									
MR ₃	0.1127 \$	0.9016		2									
	lo July tammersturys 39.0. amount of urine 500 cc. 5pec. Weight 1030	merature 30	-0-6	6 m On I	t of urin	6 50	ů Ç	spec.	haitew .	5	ଞ		

19 July temperature 39.0, emount of urine 500 cc, spec. weight 103

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1.9915 \$	8 5 3 T 0	0.017938	0.1125 \$	0.0459 \$	1.708 \$	0.026255	\$29197\$	0.005725	\$ 6151.0
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- 13 -

20 July temperature 37.5, amount of urine 450 cc. spec. weight 1028

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- 14 -

22 July temperature 38.1 amount of urine 600 cc. spec weight 102?

9. Q	+3.560	+0.279	+1.458	+1.883	+2.840					
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When considering the second table, it is noted, above all, that the body appears to react differently toward the increase in temperature in the initial stage than in the remission stage. The increase in temperature during the first days is connected with an increased elimination of nitrogen. However, once the temperature has dropped to normal, an increase in temperature is accompanied by a decrease in N-elimination. (This rule is by no means always true and we will specially emphasize those cases in which it is true. host observations show a parallel behavior of fever height and nitrogen loss.)

The chlorine elimination should be opposite to the nitrogen [elemination], that is, it should increase with the drop in body temperature. This is mostly so in our case; but even here an increase in temperature after the lever spell is connected with an increase in chlorine separation -- an observation which has been made by many authors (Rosenstein, Brattler), but which has not been sufficiently emphasized.

The phospholus elimination exhibits a certain sluggishness, it lasts longer than the temperature drop and decreases only when the fever is over for a long time. When the temperature drops, the elimination of phosphorus may increase; when the temperature increases the phosphorus elimination may drop. We find such behavior in our case in addition to the above-mentioned abnormalities of the nitrogen and chlorine elimination.

The elimination of calcium salts has been little observed until now; there are no data and we must limit ourselves to the more mentioning of this observation. In this case, the calcium elimination was opposite to the temperature. Any drop [in temperature] brought about an increase in the calcium elimination; any increase [in temperature] brought about a drop in calcium elimination. It cannot be decided whether this is specific for typhus.

According to Salkowsky, the potassium salts are aliminated to an increased extent during the fever; after the fever, they decrease considerably and disappear gradually. We can completely agree to this rule. In fact, it appears that this situation is not affected by an accidental drop or increase in temperature. To be sure, a drop in potassium salt elimination has been observed during a rapid ascent of the temperature, but this does not narrow down the validity of the rule. The parallal behavior, which some believed to have observed between P and K, does not exist in this case; a divergent behavior is frequently noted.

The sodium salts behave in the same way as chlorine; thus in our case they undergo the same fluctuations and abnormalities, as have been discussed in the case of the chlorine elimination.

The elimination of ammunia need not be specially discussed. According to Hallervorden (see reference cited), its elimination is supposedly increased ruing the fever. It could therefore be set equal to phosphorus and nitrogen. In our cases, the elimination of ammonia is actually parallel to

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that of phosphorus and exhibits the fluctuations discussed for the case of phosphorus. Therefore, a similarity between P and ammonia is more justified than one between potassium salt and phosphorus.

The other nitrogen-containing substances exhibit a complete dependence on the total elimination of nitrogen. I refer to the statements made in connection with the nitrogen elimination and to the Table given below.

The above-discussed case does not exhibit a continuous decrease in temperature; on the contrary, two recurrences are observed; these were welcome because we did not want to study the behavior during fever, but the behavior during changes in the course of the fever.

Table III

Temp.	<u>cı</u>	P N	Ca N	<u>N</u>	Na N	urca N	eoid N	allex. <u>basea</u> N	NE3 N
39.1	0.107	0.065	0.0076	0.043	0.0402	0.800	0.0099	0.0128	0.0401
38.5	0.241	0.070	0.0125	0.045	0.0845	0.862	0.0122	0.0164	0.0542
39.4	0.266	0.052	0.0046	0.051	0.0911	0.975	0.0128	0.0182	0.0533
38.6	0.108	0.077	0.0090	0.057	0.0337	0.882	0.0119	0.0161	0.0473
38.5	0.148	0.093	0.0141	0.044	0.0385	0.838	0.0146	0.0191	0.0553
39.0	0.152	0.071	0.0089		0.0229	0.855	0.0131	0.0160	0.0760
37.5	0.098	0.063	0.0101		0.0176	0.668	0.0111	0.0140	0.0700
38.0	0.190	0.060	0.0075		0.0343	0.800	0.0159	0.0199	0.0677
38.1	0.211	0,065	0.0093		0.0526	0.864	0.0167	0.0217	0.070

Table III has been constructed in such a way that the ratio of the daily elimination of urine components and of the nitrogen elimination was formed. Represented in this manner, the results show even more clearly the fluctuations occurring during an increase and during a drop in body temperature. The chlorides are always increased during an increase in temperature, and they are decreased during a drop in temperature. The phosphates behave exactly in a reverse way: they are increased during a temperature drop, and they are decreased during a raise in temperature. Calcium exhibits most clearly its antagonistic behavior with respect to the course of the temperature. Potassium behaves irregularly, even when these figures are considered, and it does not a gree with phosphorus in its behavior. The amonia is to be compared more likely with chlorine, however not as strictly as sodium. The nitrogen-containing compounds behave more or less in accordance with the temperature.

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This behavior of NH3 indicates an sold elimination which we have often anphasized and which, in most cases, takes place in the form of phosphoric sold.

Case III. M. G. 20 years of age. Feverish pneumonia.

Case history: father of the patient died supposedly of tuberculosis, the brothers and sisters are roportedly healthy. The patient himself was well until now. Three days ago he became sick with twinges in the sides and the appearance of fever chills.

Present state: nothing abnormal is detectable at the organs, except for an extended damping in the left lung. Cultivation tests confirm the diagnosis of pneumonia. The expectoration is of a rusty color and sparse. The appetite is highly reduced.

The nitrition consisted of milk and broth with egg. The amount was accurately measured.

The invoctigation began directly after admission to the hospital. The fever was high: 38.8 to 39.7. The urine was collected during the fever period and quantitatively analyzed for N. Cl. P. Ca. The nurse had been instructed to collect separately all urine on the day of the crisis prior to onset of sweating; therefore, the second portion of urine collected on the day of the crisis belongs to the afebrile period. On the subsequent days, the entire urine was collected and examined. After onset of the crisis, the fever rose again to a maximum, which was desirable for our studies. This day can be immediately recognized on the Table. After this rise in the fever, the temperature dropped and remained normal until the end of the experiment. The urine was quantitatively analyzed for all components during the crisis and on the four subsequent days; it was then analyzed for N, Cl. P. and Ca on two additional days.

Nitrogen, chlorine, phosphorus, calcium, potassium, and acdium ware determined in the feces, their total amounts ware calculated and then distributed over the individual days

The temperature ran as follows:

28 July	38.8 to 39.7	Pulsation	100	to	96		
29 July	38.5 to 39.3 to 39.0		<u>112</u>	to	700	te	104
30 July	38.1 to 39.5 to 39.7	8	96	to	100	to	99
31 July	37.8 to 39.7 to 38.8	H	96	to	108	to	96
	37.1 to 36.7				96		
1 August	37.5 to 39.3 to 38.0	•			104	to	108
2 August	37.7 to 36.2		- 86	to	- 84		
3 August	36		72				
4 August	36		70				
5 August	36	,	70				

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Balance		-19.333 - 0.025 - 0.181 + 0.223	-17.760 + 0.828 - 0.088 + 0.860	-18. 845
Diet		2.028 0.533 0.403 0.544	4.056 1.066 0.806 1.088	HS20.4
y Feces		0.1514 g	N per day	0.0018 g C1 0.08374 g P
Fer Day Urine in gram		21.210 0.556 0.500 0.109	21.665 0.082 0.810 0.016	9.982 0.05566 0.2576 0.2576 0.2576 0.2576 0.2576 0.12454 0.17066 0.17066
Spec. Wt.		1.025	1.026	1.025
Amt of Urine		1000 cc	1000 co	460 co
Percent	N 2.275 CI 0.0607 P 0.0500	M 2.121 CI 0.0556 P 0.0500 Ca 0.0109	N 2.1665 C1 0.0182 P 0.0810 Ca 0.0016	<pre>M 2.170 Cl 0.0123 F 0.0560 Ca 0.0018 K 0.1506 MA 0.052 MA 0.052 MA 0.052 MA 0.0249 arres 0.0371 xmuthine bases 0.0122 NH3 0.1057</pre>
Temp. and Pulsation	38.8 39.7 30.7 96	38.5 LL2 39.5 L00 39.0 L04	38.1 96 39.5 100 39.7 104	37.8 39.7 108 38.8 96 6 AK 712 712 712
Date	28.7	29.7	30.7	after the origin
			- 19 -	

Table IV

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Balance	+ 0.949 - 0.248 + 0.858 + 0.872 + 0.256	-12.894 + 1.022 + 0.367 + 0.791 + 0.560 + 0.560	- 8.955 + 1.059 - 0.093 + 0.769
Diet	1.066 C1 0.806 P + + 0.088 Ca + + 0.920 K +	4.056 0.806 0.920 0.657	4.056 1.056 0.806 1.088
Feces	0.21238 g Calcium		0.15148 g N
Fer Day Urine in gram	12.8079 0.0589 0.0589 0.0097 1.1004 0.1014 0.1037 0.2314 0.1277 0.7182	16.7992 0.0437 0.0555 0.0355 0.0355 2.0020 0.0979 0.1541 0.1541 0.1541	12.760 0.050 0.815 0.0075
Spec.	1.026	1.025	1.028
Aut of Urine	570 aa	710 cc	500 oc
Percent	N 2.247 C1 0.0124 P 0.1250 Ca 0.0017 K 0.193% Na 0.0175 Na 0.0175 urts acid 0.0182 urts acid 0.0182 allorurts 0.0413 basse 0.0413 tantidie 0.0231 basse 0.1260	<pre>% 2.3520 Cl 0.0061 P 0.0500 Ca 0.0012 % 0.2820 % 0.2820 % 0.0138 write actid 0.0217 allonuric bares 0.0434 rantidine bases 0.0217 bases 0.0721</pre>	H 2.352 CI 0.0010 P 0.0630 CR 0.0015
Temp. and Fulsation	37.1 92 36.7 96 12 mid- 12 mid- 6 AH	33.5 39.5 38.0 108 38.0	37.7 88 36.2 84
Date	after the crisis	1.Aug.	2.Aug

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Balance	+ 0.133	-21.715 + 0.941 + 0.851 + 0.469 + 0.469	-13.777 + 0.523 - 1.001 + 0.261 + 0.058
Diet	0.920 0.657	4.056 1.066 0.806 1.088 0.920 0.657	2.028 0.533 0.540 0.460 0.460 0.460 0.460
Feces	0.15148 g N	0.00182 g Cl	0.08374 g P
Per Day Urine in gram	1.259 0.188 0.0785 0.231 0.560	25.620 0.1230 0.7850 0.1880 0.1880 0.1880 0.280 0.280	0.238 1.071 15.654 0.0083 1.3201 1.3201 1.3201 1.3201 1.3248 0.387 0.387 0.1577
Spec. Wt.	1,028	1.025	1.028
Ant of Urine	500 ca	1000 cc	830 co
Percent	K 0.2518 Na 0.0376 Wric acid 0.0157 alloxuric bases 0.0462 xathine bases 0.0305 big 0.112	<pre>g 2.562 F 0.123 F 0.123 F 0.123 F 0.125 F 0.185 K 0.2106 M 0.0188 Wallowith Allowith Allowith Allowith Allowith C.0510</pre>	bases 0.0238 ME: 0.1071 F 1.8865 C 0.001 P 0.1470 C 0.0086 MA 0.0468 MA 0.0468 WA 0.0468 WA 0.0468 WA 0.0468
Temp. and Pulsation		36.1 72	36.2 20
Date		3 Aug	t Aug

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	Temp. and	ard brad		Amt of	Spea.	Per Day Urine	Foces	Diet	Balance
Uate	Kulsai	S S	rent	Urine	Σt.	un gram			
4 Aug	36.2 70		alloouric bases 0.04450 cantiine bases 0.02530	830 cc	1.028	0.3652 0.2075	0.21238		
			11H3 0-0750			0.581	g Сл		
5 Aag 36.1		ű2	N 1.995 Cl 0.0364 F 0.1885 Ca 0.0220	830 aa	1.028	16.517 0.155 0.560 0.1830		2.028 0.533 0.544	-14.644 + 0.376 + 1.241 + 0.314
6 Aug 36.5	36.5	02	N 1.855 Cl 0.6068 F 0.121 Ca 0.00436	840 cc	1.025	15.582 5.093 1.016 0.361			、
7 Aug	36.2	70	R 0.0965 Ca	870 00	1.025	6.595 6.839 1			

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Table V

Тв	ble V r	opresent.	s the no	urly elt	mination	during e	Table V ropresents the nourly elimination during oight days.		Uric		
Date	Temp.	Z	ដ	μ,	Ca	ы	Na Ur	Urea	Ac1d	Bases N	E E
	38.9		-	0.021	0.100.0	1		•	•	ÿ	
	000		-	10.034	0.0007	ı	1	8	1	•	
	8.8°		-	0.0143	0.0005	0.0385	0.0168 0.	0.5009	0.0063	0.00% 0.0	0.0270
	36.8			0.1887	0.0016	0.1433	0.0169 1.	9285	C10.0	0.0385 0.1	175
	38.3		-	0.0148	0.0004	0.0867	0.0041	1	0°0064	0.0128 0.0	543
2 Aug	36.0		-	0460.0	0.0003	0.0525	0.0078	,	0.0032	0.0960.0)266
			-	1420-0	0.0007	0.0875	0.0078 0.	9917	0.0127	0.0216 0.0	
4 Aug	36.1		-	0.0550	0.0029	0.0775	0.0161 0.	0.552	0.0066	0.0153 P 0	7980 790
5 Aug	36.1		-	0.0650	0.0076	8	1		ł	•	
e Aug	36.1	0.649	0.2109	1640.0	0.0150	J			8	8	•
T D	hle VI	shows th	o ratio	of the e	Table xcreted	VT amount wi	Table VI shows the ratic of the excreted amount with respect to the amount of nitrogen.	to th	ງແມວແຮ ອ	: of nitrog	•ueS

ERN Allox Bases Urto Acid Urea Urea

	•	ថ	ρ.	Ca	×	2	Urea	Acid	Вазев	6 ₇₄
Dete	Tomp	Z	X	N	2	X	N	X	36 ,	R
20 Jul	38.9	0.0260	0.024	0*00*0	I	4	•	•	ł	ŧ
		0.0000	0.033	0.0008	•	İ	ł	ł	ŧ	•
	38-8	0.0260	0.026	0.0099	0.069	0.0303	0.903	0.0110	0.0158	0.049
31	36.8	0.0050	0.084	0.0007	0.067	0.0079	0.865	0.0078	0.0173	50.0
BEA	38.3	0.0030	0.021	0.00055	42C.0	0.0058	ł	0.0091	0.0183	0.0304
2 Ang	36.9	0000	0.064	0.00056	0.098	0.004	t	0,0060	0.0180	0.042
3 Aue	36.1	0,0049	690*0	0.0006	0.082	0.0073	0.929	0.01A	0.0201	140.0
44 Aug	36.1	0.0004	fi. 084	-+	0.118	0.024	0.800	0.0101	0.023	0.132
5 Aug	36.1	0.0098	0.094	1110.0	I	e	4	1	t	1
é Aug	36.1	0.322	C.063	0*0540	8	Ŧ	•	ŧ	\$	a

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It is seen from Table IV that the loss in nitrogen may be larger after the crisis than during the fever. When the amount eliminated in the urine prior to and after the crisis is considered, the percentage content and also the absolute amounts of those components, whose elimination is characteristic for the fever, are found to be increased. This is observed even more distinctly when the hourly amounts are compared. It is found (Table V) that nitregen and potassium are eliminated four times as much, chlorine and calcium three times as much, emmonia five times as much, and phosphorus eight times as much as prior to the crisis. The elimination of Na is peculiar because it is practically unchanged, whereas the nitrogen-containing components behave all similar to the other ones, that is, they are increased. It should be exphasized that the xanthine bases are eliminated twice as much [as prior to the crisis], whereas the former is eliminated four times as much. The eliminated amount surpasses by far the amount observed on fever days; also the loss in nitrogen and phosphorus, calculated from the balance, is at a maximum on the day of the crisis. When the retio of the eliminated compounds with respect to N is calculated (Table VI). P is again at the top; in fact, the other urine components behave, at times differently. For instance, ablorine, potassium, sodium, caloium, et cetera are decreased with respect to nitrogen in spice of the fact that the bourly elimination increased significantly. As is seen from Table VI, sodium, chlorine, calcium, potassium, ures, and uric acid are decreased with respect to nitrogen (sodium the most). whereas phosphorus, zanthine bases, and ammonia are increased.

So much for the characterization of the crisis. As far as the postfebrile period is concerned, each increase in body temperature is accompanied by a decrease in nitrogen and phospherus elimination, whereas the potassium and uric acid elimination increases. Chlorine, calcium, and socium behave like nitrogen; consequently, they decrease when the body temperature increases.

The reverse takes place when the body temperature drops. The nitrogen elimination is increased and so are the phosphorus and annonia eliminations. The greatest losses in nitrogen and phosphorus occur during this period, as is seen from the balance. Chlorine behaves irregularly; often, it increases during the afebrile days. However, subsequently it drops to such values which have never been observed at the climax of the fever. It follows from the balance that retention in the body is greatest after the remission and that it gradually decreases. The sodium and calcium balances behave in the same manner. On the other hand, the loss in phosphorus increases after remission and this P-loss increases further after the fever, until it begins to recede after three to four days. The potassium salts are lost by the body, to the slightest extent immediately after the fever, subsequently more distinctly.

Approximately the same follows from an analysis of the Table of the hourly elimination; only in this case the dependence on the course of the fever is more distinct.

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It is still more distinct in Table VI, where the hourly elimination is calculated with respect to nitrogen. The behavior of phosphorus exhibits the greatest regularity, chlorine is vague and so are the potassium and sodium salts.

It cannot be denied that chlorine, calcium, and potassium behave quite differently than in the typhus cases, and this may be the reason for the large difference in the results of the fever studies.

Case IV. H.K., 19 years of ege. Febris intermittens quotidiana.

Case history: patient's father died of cholera, the mother is sickly. Three brothers and sisters died when they were young. The patient comes from Hamburg, spent some time in Italy where he got sick with fever and headaches. He left Italy and went to Switzerland. During the journey by foot, he suffered from fever chills which he had not had previously during the fever sttacks in Italy.

Present state: nothing detectable at the organs, except for a slight enlargement of the spleen. Hemoglobin content 60 percent, [red] blood count 4,120,000.

The dist consisted of milk and broth and consisted per day of

10.340 g N	6.246 g C <u>1</u>
1.260 g P	1.740 g Ca
2.746 g K	3.562 g Na

The urine was collected in five portions. The first portion comprised the time period prior to the fever 10 AM to 2:30 FM. Temperature $36.4^{\circ}C_{*}$ pulsation 92. Amount of urine 125 cc.

The second portion comprised the time of the fever chills 2:30 PM to 4 PM. Temperature 41.0°C, pulsation 106. Amount of urine 235 cc.

The third portion comprised the time period of the climex of the fover 4 FA to 7 PM. Temperature 40.2°C, pulsetion 96. Amount of urine 295 cc.

The fourth portion comprised the time period after the fever attack 7 PM to 6 AM. Temperature 3919, 38.6, 37.5°C, pulsation 108, 104, 96. Amount of urine 325 co.

- 2<u>5</u> -

The patient eliminated 1165 co urine in which:

N	15.792	balance N -	5.452
Cl	4.934	C1 +	1.312
\mathbf{P}	0.917	P +	0.343
Ca	0.263	Ca +	1.477
ĸ	2.665	<u>K</u> +	0.083
Na	2.739	Na t	0.823

The food contained the above-mentioned amounts. Thus, the balance showed, as will be found on the subsequent Table, a loss for nitrogen. All other components were retained, a correction must be made for this because the feces were not considered. This concerns mainly P. Ca. K. If one remembers the known numbers, a slight loss in P and potassium may be assumed, whereas calcium, sodium, and chlorine remain probably unchanged.

In the food the ratio to the nitrogen is as follows:

N: Cl: P: Ca: K: Na: 1: 0.62 0.12 0.17 0.27 0.35 in the food 1: 0.312 0.058 0.016 0.161 0.165 in the feces

Compared to nitrogen, all components are eliminated to a lesser extent. The hourly elimination is given in Table VII. Table VII. Malaria Fever

71mo	Temp. and Fulsa- tion	Percent	Amt of Urine	Spec. Wt.	Total Elimination	Per hour Gram	Time
I.	36.4 92	1.6205 0.2912 0.0891 0.0484	125 ca	1.022	N 2.025 C1 0.364 P 0.111 Ca 0.061 K4 0.172 N4 0.172	0.455 0.081 0.025 0.014 0.014 0.038	10 Af until 2:30 PM: 4.5 bours: emount of urine per bour 27.7 cc
11.	41°0	0.7875 0.6371 0.0162 0.0171 0.2390 0.3676	235 ca	1.018	N 1.8485 C1 1.4969 P 0.0376 Ca 0.0399 K 0.5497 Na 0.8454	1.2325 0.9969 0.0266 0.3665 0.5636	2:30 PM until 4 PM: 1.5 hours; amount of urine per hour 94 co
H.	40°0	9.183 0.4854 0.0043 0.0252 0.1884 0.2584 0.0105 0.0105	295 co	1.019	~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~	1.160 0.472 0.0042 0.1849 0.2534 0.0103	4 PM until 7 PM: 3 hours; amount of urine per hour 95 cc
		0.0004 0.0266			pases 0.0021 zanthine bases 0.0018 NH ₃ 0.0784	0.0004	

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The	Temp. and Pulsa- tion	Percent	Amt of Urine	Spec. Ht.	Total Eliminetion	Per hour Gram	fur-
IY.	39.9 33.6 104 104	1.645 0.3034 0.3034 0.1458 0.1458 0.1458 0.1458 0.1798 1.120 0.0245 0.0245	325 00	1.024	ж	0.4845 0.0895 0.0041 0.0859 0.0859 0.0859 0.0872 0.0872	7 PM until 6 AM; 11 hours; amount of urine per hour 29.5 cc
*	% .8	0.0004 0.0560 1.687 0.3641 0.9234 0.02208 0.2208 0.624	185 oo	1.025	Example of the second s	0.0000 0.0169 0.01683 0.07770 0.01683 0.07770 0.05040 0.0940 0.0940 0.0940 0.0940	6 AM until 10 AM; 4 bours; amount of urine per bour 45.2 cc

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m	<u>C1</u>	P	Ca	ĸ	Na	urea	NH3	acid	DASES
Temp.	ĸ	N	N	N	N	N	N	N	N
36.4	0.178	0.056	0.0307	0.092	0.083	-	-	-	-
41.0	0.809	0.020	0.0215	0.297	0.457	0.996	0.032	-	-
40.0	0.407	0.003	0.0212	0.159	0.218	-	0.0224	0.009	0.009
38.6	0.184	0.089	0.0084	0.117	0.109	0.660	0.034	0.014	0.005
36.3	0.215	0.091	0.0139	0.137	0.121	•	-	-	•

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The results of this study do not present much new information so that it is hardly necessary to discuss them. What has been found by Limbeck, Terray, Hitzig, and others has been confirmed [here].

The elimination of nitrogen rose and declines with the temperature -however, only when taken absolutely. Relative to the water elimination, the nitrogen elimination was decreased during the fever, as may be seen from the percentage of nitrogen.

The chlorine elimination increased during the fever quite significantly when taken relatively, as well as when taken absolutely; it decreased immediately after the remission of the fever and increased during apyrexis.

On the other hand, the phosphorus elimination dropped during the fever and reached its maximum in the post-febrile period. The elimination of P was normal in the post-febrile period.

The elimination of potassium salts was completely analogous to that of chlorine, even if relative to the water elimination, at the onset of the fever, a drop could be observed which lasted during the fever.

Potassium salts take up a special position, as usual. Relative to the water elimination, they attain their maximum in the post-febrile period. Taken absolutely, they follow chlorine and have their minimum in the post-febrile period. The sodium salts are to be compared to chlorine in all aspects.

The elimination of nitrogen-containing substances could not be determined in all cases because of lack of material. It is therefore difficult to state anything about its course. It seems as if the elimination of uric acid and xanthine bases increased relatively with respect to water and N elimination after the fever, even if the absolute amount per hour is lower in the post-febrile period.

In comparing the hourly amount eliminated with nitrogan, the abovestated facts become even more pronounced.

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The daily elimination in comparison with the food consumption exhibits the course typical during the fever; a relatively diminished chlorine, calcium, and sodium elimination, and an increased nitrogen, phosphorus, and potassium elimination.

Case V. B.K., 49 years of age. Phtisis pulmonum Febris hectic ..

Case history: father, brothers, and sisters are healthy, mother suffered supposedly from cough. The (female) patient had gonitis. She got married and had two children who died both in their childhood. The present suffering started a few months ago and resulted in a general weakness.

Present state: patient exhibits in both lungs the phenomena of a decomposing infiltration; she expectorates a lot. The other organs do not axhibit any abnormalities. Patient has hectic typhus. The difference in body temperature amounts to 2.5°C.

The rationt was given the same diet for several days. This diet consisted of milk and broth. The urine was analyzed on two days; no differences were observed. The daily elimination was collected in four portions of which

the first portion was collected from 12 noon to 4 PM, corresponding to the prae-febrile period. The body temperature was 37.0, pulsation 100;

the second partian was callected in the febrile period from 4 PM to 10 PM. Temperature 38.0 to 39.0;

the third period corresponds to the remission from 10 PM to 6 AM. Temperature 37.0:

the fourth and final period lested from 6 AM to 12 noon and corresponded to the apyrexia. Temperature 36.0, pulsation 90.

Also in this case, the foces were not examined; however, this does not affect the problem under consideration. The results were:

<u>Flimination</u>	Food	Balance
5.6006 N	10.340	+4.7394 gram N
3.7664 01	6.246	+2.4796 - Cl
0.2749 P	1.260	+0.9851 - P
0.0952 Ca	1.740	+1.6/48 - Co
1.0642 K	2.748	+1.6840 -
1.1028 Na	3-562	+2.4592 - Na

The ratio of N to Cl. P. Ca. K. Na in the food was:

		Cl: 0.62	 Ca: 0.17	K: 0.27	Na: 0,35
in the eli- mination:	-		0.016		

The difference concerns Ca and P; both are ample in the foces. We may assume that no special retention takes place, only a general apathy, an insufficient assimilation.

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	Temp. and	4			Total Elimina	Per bour	Water
Time	Fulsa- ticn	Nrtha	vpec.	Percent	tion	Gram	Per hour
C f	000			212-2 N	0.9548	0.2387	і.
				CI 0.1214	0.5342	0.1335	210 cc
111 - 00 4 4				P 0.0235	420.1.0	0.0259	
		40	1	Ca 0.0049	0.0215	500°0	
		0	.0	K 0.0401	0.1764	11110°0	
	وعند	cu	06	Na 0.0515	0.2860	0.0715	
				urea 0.196	0.8624	0.2156	
				alloxiric			
					0.0430	c.0108	
				NH3 0.0056	0.0246	0.0061	
					0000	0.2808	E
4 PM to	138-39			N 0.3/3			
10 PM	106			C1 0.315	1.138	6ThZ*0	1 70.0 00
6 hours				P 0.0290	0.1335	0.0223	
				Ca 0.0068	0.0312	0.0052	
		46	ì	K 0.0629	0.2893	0.0482	
		0	•Û	Na 0.0852	0.3919	0.0653	
		c0	10	urea 0.341	0.5685	0.2614	
		1	,	uric acid 0.0077	1550.0	0,0060	
				ellomric			
	_			bases 0.0109	0.0502	0.0084	
				9	0,20	dena a	
					0150 0	0.0085	
				ATTON CON	0T(0.0	()))···	
Of NG OF			_	N 0.392	1.764	0.2205	H.
AM.	18	4		C1 0.2669	1.201	0.1501	<u>5</u> 6 80
f hours	<u>}</u>	-50	1.	P 0.0160	0.072	0.0090	
e mon 0) 0	00	Ca 0.0078	0.0351	0.0043	
		0	9	K 0.0829	1676.0	Thto 0	
				Ma 0.0590	0.2655	0.0332	
				urea 0.322		1191.0	-

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Water Per hour	IV. 40 cc	r 8
Per bour Gram	0.0045 0.0063 0.0055 0.0055 0.0070 0.0070 0.0070 0.0070 0.0070 0.00356 0.0033	0.269 0.179 0.0266 0.0056 0.0026 0.0074 0.1710 0.0062 0.0062
Total Elimina- tion	0.0365 0.0522 0.0158 0.0437 0.05822 0.05822 0.05822 0.1594 0.1594	1.0780 0.7140 0.1064 0.0225 0.0329 1.0260 0.0392 0.0392 0.0392
Percent	urio acid 0.0081 Alloxuric bases 0.0116 xanthine bases 0.0035 bases 0.0035 bases 0.0035 ku, 476 cl 0.2427 P 0.0275 ca 0.0031 K 0.0939 Ma 0.664 Mk3 0.0084 Second Day	<pre>M 0.385 CI 0.255 P 0.039 uric ucid 0.0081 alloxuric bases 0.0115 xanthine bases 0.0015 MH3 0.0105 MH3 0.0105 MH3 0.0105 MH3 0.0105 MH3 0.0105 alloxuric bases 0.0109</pre>
Spec. Wt.	1.011	1
Amt of Urine	240 co	280 cc 360 cc
Temp. and Pulsa tion	36.0	33 °
T1me	6 AM to 12 noon; 6 hours	12 noon to 4 PM; 4 hours 4 PM to 10 PM; 6 hours
	- 32 -	

Water Per hour	III. 31 cc		. 0 9 00	
Per hour Gram	0.0080 0.0080 0.0161 0.016 0.010 0.010	0,0039 6000,0 5203,0	0.0011 0.0011 0.0017 0.0006	
Total Elimina- tion	0.0022 0.0479 1.286 0.637 0.080	0.0395 0.0025 0.0420	0.1183 0.0065 0.0102 0.0036	
Percent	xanthine bases 0.0006 NH3 0.0133 B 0.5145 C1 0.2548 P 0.0320 uric acid 0.0116	alloxuric bases 0.0126 xanthine bases 0.0010 NHg 0.0168	N Cl 0.2366 uric actd 0.0131 allowuric bases 0.0205 xanthine bases 0.0073	
Spec. Wt.	I		ł	
Amt of Urine	250) co	50 cc	
Temp. and Pulsa. tion	380		36	
T 1me	10 PM to 6 AM 8 hours		6 AK to 12 noon; 6 hours	
		-	33 -	

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Table X

Temp.	<u></u>	<u>P</u>	Ca	K	Na	urea	2518	allox. bases	NH3
• • • • • •	N	N	N	1	N	N	N	N	N
37°a	0.557	0.109	0.022	0.180	0.299	0.903	-	0.041	0.025
ъ	0.667	0.100	-	4	-	-	0.021	0.030	0.027
39°a	0.833	0.077	0.014	0.1(4	0.285	0.902	0.021	0.029	0.029
b	0.557	0.069	-	-		-	0.019	0.021	0.025
38°2	0.681	0.041	0.020	0.200	0.151	0.823	0.021	C.029	0.025
b	0.494	0.062	•	-	-	•	0.022	0.024	•
د ائ	0.510	0.058	0.006	0.197	0.140	-	-	•	0.030

The results of the observation of elimination during hectic fever confirm the results observed in the malaris case. Also in this case, the chlorine elimination is lower during the febrile period than during the affebrile period. The minimum is found during the apyrexia. The phosphorus end tion drops with increasing body temperature, reaches a minimum durin and post-febrile period, and subsequently increases during the period of the apyrexia up to the prae-febrile period during which it reaches its maximum.

Calcium drops with the onset of the fover, decreases somewhat more, and drops to its minimum in the prac-febrile period.

Strangely enough, sodium does not behave in parallel with chlorine. It drops at the climax of the fever, subsequently it drops still more, does not reach the minimum in the post-febrile period (in agreement with chlorine), and has its maximum in the prac-febrile period. Polassium increases with the fever, drops steadily up to the apyrexia during which it reaches its minimum, whereas the maximum occurs at the climax of the fever. Urea is eliminated to the largest degree at the climax of the fever; the alloxuric bases have their minimum in the prac-febrile period, whereas their maximum occurs during the apyrexia.

Amnonia, like chlorine, is eliminated to a larger extent at the climax of the fever; subsequently it drops, reaches its minimum during the apyrexia, and ascends in the prae-febrile period.

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When the elimination is compared with nitrogen, it appears to be somewhat different, but no essential differences are noticed. Thus, chlorine has its maximum at the climax of the fever, its minimum during the apyrexia; phosphorus has its maximum during the prae-febrile period, its minimum during the post-febrile period. Calcium has its minimum during the apyrexia, its maximum during the prae-febrile period. Potassium has its minimum at the climax of the fever, its maximum during the post-febrile period. Sodium reaches its minimum during the apyrexia, its maximum during the prae-febrile period. The alloxuric bases are eliminated most during the prae-febrile period, least during the apyrexia. Ammonia behaves exactly in the reverse way.

The rise in body temperature is accordent to the second s

temperature is accompanied by an increase in phosphorus and nitrogen, and by a decrease in all other components. Calcium, sodium, and potassium exhibited an increase only in the second fever period.

The study of the elimination in the various types of fevers has shown with certainty only one thing: elimination during remission may possess the characteristics of an elimination during a fever in a pronounced way. Elimination during rise of the fever can exhibit the same characteristics to a lesser extent.

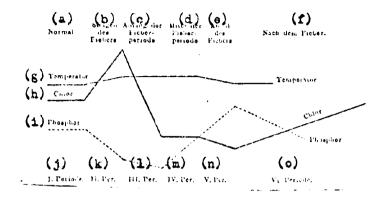
When I am talking about elimination during the fever, I mean the known retention of chlorine and the increase in phosphorus and nitrogen. This can also occur during remission and can disappear when the fever increases.

In more dotail, this means that after the drop in body temperature, chlorine is especially retained, and phosphorus and nitrogen are being especially [strongly] eliminated. If another increase in body temperature follows, chlorine is increased, although it should be retained; phosphorus is eliminated in smaller quantities instead of in increased quantities.

It appears that the body reacts differently at the onset and at the end of the fever than during the fever. In the long run, the fever causes the known changes; if, however, a temperature increase occurs also during the fever, then a situation arises which corresponds to the corvalescence pariod (the afebrile period). If a decrease in body temperature takes place, increased fever phenomena occur. Rise and drop in body temperature shall be a measure for the fever. The question why this is justified, will not be discussed here.

Therefore, when we wish to devise a scheme according to what has been said, it will be of the following form: the first rise in temperature re-sults in an increase in chlorine (I will omit sodium, potassium, calcium, and annonia as not to complicate the discussion), in a decrease of nitrogen (or in an increase), and in a diminishing of phosphorus. When the fever persists unchanged, the second period exhibits a gradual decrease in chlorine elimination, an increase in nitrogen (or nitrogen stays at the provious height), a decrease in phosphorus. The third period -- still at the same temperature -- exhibits a decrease in chlorine, and an increase in phosphorus and nitrogen; to be sure, chlorine has dropped below normal, and phosphorus and nitrogen have risen above normal. At this point, we have the type of elimination which is known as fever urine. This situation may last when the body temperature remains unchanged. When the body temperature drops (perhaps due to dilution of the blood), the fourth stage set in: chlorine is retained still more, phophorus and nitrogen are eliminated still more strongly. When the drop in temperature lasts, chlorine attains its minimum, phosphorus and nitrogen attain their maximum, and the situation changes gradually. The fifth stage begins, the chlorine elimination rises up to normal, 2 and N recede to normal.

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Key to Figure: a--Normal; b--rise of fever; c--onset of fever period; d--middle of fever period; e--remission; f--after the fever; g--temperature; h--chlorine; i--phosphorus; j-- I. period; k--II period; 1--III period; m--IV period; n--V period; o--VI period.

This would be the situation of the fever as it occurs in malaria and as we have found it in other types of fever.

If an explanation is attempted, it must be presupposed that the blood vessels are the ones, or rather that it is the blood which undergoes the same changes as the urine. The blood contains more water and chlorine than the tissues, but it contains less phosphorus and nitrogen.

Thus, when we find the urine to be enriched in chlorine and poor in phosphorus at the onset of the fever (I period), then this implies that the flow from the tissues into the blood is interrupted; the blood supplies a small amount of water and a small amount of sodium chloride to the tissues, it takes up a small amount of phosphorus and a small amount of nitrogen. This corresponds to a circulation which makes the fiffusion flow little effective either by narrowing of the vessels or by slowing down the blood flow.

This type of circulation does not last long; it is followed by a stage of enlargement of the vessels and fast circulation. The II and III stage occur. The blood which is richer in water supplies its whiler and sodium chloride to the organs; it retains the externally added water and carries the nitrogen-containing substances along. The decomposition of the nitrogencontaining substances occurs within a short period of time, the phosphoruscontaining substances are more stable. For this reason, the minimum of phosphorus occurs at this point and the N in the urine increases at a faster rate than the phosphorus. Thus, we are at the climax of the fever and the decomposing large albumin molecule yields increasingly more substances. The small molecule of the latter increases the osnotic pressure

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and water and salts^{*} are increasingly retained. The albumin decomposition is, therefore, caused and maintained by the increase in water content of the body. When the fever starts to recede, that is, when the circulation is improved and the organs give off water, the fever phenomena are enhanced; the flow is reversed and the water with its salts is transferred from the organs into the blood instead of from the blood to the organs; the salts are poor in chlorine and rich in phosphorus. The elimination of water is increased, however the elimination of sodium chloride is not increased; phosphorus salts and nitrogen are eliminated with the water until the equilibrium is being reestablished and everything goes back to normal. If we imagine during the remission stage a sudden increase in the body temperature then the phosphorus elimination is inhibited due to the disturbance in the circulation, [and] the chlorine is increased; the reverse takes place when we think of a drop in temperature during the fever. It is seen from the studies by Rosenthal (23), E. Aronsohn (24), Geigel (25), Maragliano (26) that a decrease in heat evolution (contraction of the vessels) takes place in acute fever cases when the body temperature increases (Rosenthal); [it is also found] that after the temperature remission and during the remission, the volume of the vessels increases (pletismograph -- experiments by Maragliano).

The increased nitrogen elimination in the stage of the latent fever period, which has been observed by Naunyn, does not contradict the abovestated facts; because firstly, we observed also an increase in N with temperature, and secondly, the experiments of Naunyn involve dogs who had been subcutaneously injected with samies; thus, the fever started with the decomposition in the blood, as it is also the case with malaria.

Röhmann's explanation of the chlorine retention by the tissue albumin still holds. I have expressed a similar opinion when I studied the metabolism in anaguia cases; however, I believe that, even without this assumption, there are enough reasons for the chlorine retention.

Von Terray explained the increased chlorine elimination with the decomposition of the blood corpuscles. The blood corpuscles contain such a small amount of chlorine that they would all have to decompose to provide the necessary amount. When, however, a chronic decomposition of blood corpuscles takes place, the chlorine is by no means increased; on the contrary, it is decreased, as it is the case with all chronic anaemias.

There are also theories about the post-febrile nitrogen elimination. For instance, Naunyn and others believe that the increased N-elimination is due to an accumulation of nitrogen in the body. The kidneys are overloaded with decomposition products and give off the entire load at once. This opinion would be difficult to contradict, were it not for the fact that in

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^{*} Cf. E. Schulze, Zeitschrift für physikalische Chemie Vol 23. page 18. "On the metabolism of albumens in the living plant"

the post-fobrile period a decrease in chlorine elimination is observed in addition to the accumulation of nitrogen. This leads us to believe that a change in the circulation is the cause of all phenomena. After the fever, the exchange between the tissues and the blood corpuscles must become more vigorous; this way, an elimination occurs which represents a more acute picture of the fever elimination.

This post-febrile and prae-febrile change in the vessels has a special effect, particularly on soluble salts; only sodium and potassium salts -probably the soluble chlorides -- exhibit the peculiarity of a swift rise, as well as that of a rapid drop. Calcium phosphate does not follow the fluctuations as rapidly, and its typical behavior remains: it decreases with a temperature increase and it increases with a temperature drop. Phosphorus and ammonium salts belong also to the sluggish substances. The phosphorus elimination must be preceded by a nucleic acid decomposition. For this reason, phosphorus reaches its maximum only in the second stage and lasts longer than the fever. The formation of the ammonium salts requires and "acidification"* of the fluids. Therefore, ammonia behaves similar to phosphorus.

Potassium exhibits, no doubt, the most peculiar behavior, and when we explain the increased elimination during the temperature rise with the fast diffusion, there remains no explanation whatsoever for the drop in the postfebrile period which, by the way, is not a rule, except that we would have to assume that potassium obeys here the same laws as those obeyed by sodium.

The explanation, whether elegant or not, is not the essential point of concern and it does not change anything in the observations. In the old studies of Langley and Huppert we have found figures which had been neglected or which had been considered useless by them, [and] which become even more significant in view of our explanation. We hope that we have been able to supply with our observations reliable data also for other theories.

 "Acidification", in the sense used previously, caused by a decrease in the alkalinity which decrease coincides with the increase in ammonia.

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