Effect of Onions on Blood Fibrinolytic Activity

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Summary: Twenty-two convalescent patients at rest were given a fat-enriched breakfast with and without the addition of 60 g. of fried or boiled onions. Both forms of onions were found to prevent the expected decrease in fibrinolytic activity, and indeed the latter was actually increased.

Introduction

A casual remark by a patient that in France when a horse develops clots in the legs it is treated by a diet of garlic and onions led one of us (I. S. M.) to investigate onions as a possible source of a fibrinolytic agent. A search of the medical literature led to the discovery of an investigation of Indians residing in India which showed that fibrinolytic activity was significantly reduced after a fat-enriched meal (Gupta et al., 1966). Similar results have been obtained in other countries (Greig and Runde, 1957; Billimoria et al., 1959; Mehrotra et al., 1966; Moser and Hajjar, 1966; Menon, 1967a). Ingestion of fried onions not only prevented this reduction in fibrinolytic activity but produced a significant increase of it in the blood stream (Gupta et al., 1966). The investigation described here was undertaken in order to study the effect of onions on the fibrinolytic activity of Britons.

Subjects and Methods

Twenty-two ambulant patients aged 19 to 78 years were selected for study. They were convalescing from bleeding peptic ulcer, were free of symptoms, and were not on any drug therapy at the time of examination.

After fasting samples of blood were collected from all patients for the estimation of fibrinolytic activity, cholesterol, recalcified clotting-time, thrombotest, and fibrinogen titre. The patients were divided into two groups.

Group 1 (14 patients).—After withdrawal of the samples mentioned the first seven patients were given a breakfast containing 98 g. of fat at 9.30 a.m. and new samples of blood were collected after two and three hours. The procedure was repeated the following day, but this time 60 g. of fried onions were added to the meal. For the remaining seven patients the routine was reversed and they were given the fried onions with their breakfast on the first day.

Group 2 (8 patients).—The method was identical to that outlined above, the only difference being that four patients in this group had boiled onions with their breakfast on the first day and none on the second. The opposite was the case with the remaining four patients.

All the 22 subjects were kept in bed throughout the experiment, since a previous study had shown that moderate exercise increases fibrinolytic activity (Menon, 1966a; Menon et al., 1967). Smoking was not permitted.

Euglobulin lysis time was estimated by von Kaulla's method (1963), slightly modified as previously described (Menon and Dewar, 1967). The fibrinolytic activity has been derived from these lysis times and expressed in units by multiplying the reciprocal of these lysis times in minutes by 10,000. Fibrinogen titre was measured by the method of Schneider (1952), which gives a rough guide to the fibrinogen level inside the body. Cholesterol levels, recalcified clotting-times, and thrombotests were estimated by standard methods.

Results

Tables I, II, and III show the fibrinolytic activity of the two groups of patients. The results show that after ingestion of a fat-enriched breakfast a decrease of fibrinolytic activity

TABLE I .- Effect of Fried Onions on Fibrinolytic Activity

	Euglobulin Lysis Time (in Units)							
		Breakfast On	ly	Breakfast + Onions				
	9 a.m.	11.30 a.m.	12.30 p.m.	9 a.m.	11.30 a.m.	12.30 p.m		
1	57-1	50.0	48.7	86-9	90.9	100-0		
2 3 4 5 6 7 8	62.5	48.7	46.5	66.6	86.9	90.9		
3	41.6	27·7 35·7	28·9 37·0	33·3 86·9	41·6 90·9	41·6 95·2		
4	105·0 27·7	23.8	23.8	30·3	40-8	40.0		
6	50.0	35.7	44.4	26-3	76.9	74.0		
7	44.4	33.3	34.4	60-6	71.4	66.6		
8	20-8	20.4	21.2	12.8	41.6	38.4		
	23.8	25.0	22.7	55.5	71.4	76.9		
10	26.6	25.0	24.0	23.2	35.0	31.7		
11	71·4 83·3	76·9 66·6	54·0 62·5	33·8 31·7	74·0 64·5	83·3 66·6		
12 13	20-8	20.0	20.4	18.5	34.4	33.3		
14	23.8	25.0	22.7	27.7	34.4	33.3		
Mean	47-1	36-7	35-1	42-4	61-1	62-3		

TABLE II .- Effect of Boiled Onions on Fibrinolytic Activity

	Buglobulin Lysis Time (in Units)								
	Breakfast Only			Breakfast + Onions					
	9 a.m.	11.30 a.m.	12.30 p.m.	9 a.m.	11.30 a.m.	12.30 p.m			
15 16 17 18 19 20 21	55-5 57-1 38-4 76-9 27-7 47-6 32-2 50-0	41.6 60.6 34.4 66.6 25.0 38.4 27.7 38.4	43·4 55:-5 33·8 68·9 25·0 36·3 28·5 40·0	31·2 54·0 34·4 83·3 25·6 50·0 41·6 55·5	66-6 74-0 50-0 111-1 41-6 68-9 50-0 83-3	83-3 74-0 55-5 117-6 41-6 66-6 50-0 76-9			
Mean	48.2	41.6	41.4	47:0	68.2	70-7			

TABLE III.—Fibrinolytic Activity and the Effect of Onions

	Group 1 (14 Patients)	Group 2 (8 Patients)		
·	Breakfast Only	Breakfast and Fried Onions	Breakfast Only	Breakfast and Boiled Onions	
Mean B.L.T. 9 a.m. Mean E.L.T. 11.30 a.m. Mean change 9-11.30 a.m. S.E. of mean change Significance of mean change	47·1	42·4	48·2	47·0	
	36·7	61·1	41·6	68·2	
	- 10·4	+18·7	-6·6	+21·2	
	± 4·9	±3·8	±2·0	±3·0	
	P < 0·055	P<0·001	P<0·02	P<0·001	
Mean E.L.T. 12.30 p.m. Mean change 9 a.m 12.30 p.m Significance of mean change	35·1	62·3	41·4	70·7	
	- 12·0	+19·9	-6·8	+23·7	
	P < 0·05	P<0·001	P<0·01	P<0·001	

All comparisons are made on a "within-patient" basis: individual changes g., from 9 to 11.30 ..m.) were calculated for each patient, and standard errors (e.g., from 9 to 11.3 were based on them.

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occurs. The addition of onions, whether fried or boiled, not only prevented this reduction but also caused a marked increase. Recalcified clotting-times, thrombotests, and cholesterol and fibrinogen levels were not significantly changed (Table IV).

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LE IV.—Mean Values of Recalcified Clotting-times (R.C.T.), Thrombotest, Cholesterol, and Fibrinogen With and Without Fried Onions (22 Patients)

	Ordinary Breakfast			Ordinary Breakfast Plus Onions		
	9 a.m.	11.30 a.m.	12.30 p.m.	9 a.m.	11.30 a.m.	12.30 p.m.
R.C.T. (seconds) Thrombotest (%) Cholesterol	87·8 80·1	88·4 81·2	90.2	91·9 82·2	92·3 81·8	92·1
(mg./100 ml.) D H Fibrinogen	220·7 2 6 5	3 3 9	216·7 3 2 8	212·5 2 3 7	0 5 5	219·8 2 1 7
G H	8	5 2	5 4	8 2	10 2	8

Discussion

The results confirm the findings of Gupta et al. (1966) that the addition of onions to a fat-enriched meal not only presents the expected reduction in fibrinolytic activity caused by such a diet but actually promotes significant increase of it in every patient (P<0.001). The results also indicate that the factor responsible is not only heat-stable but probably not water-

Undoubtedly there is a need for some substance which will have this effect on fibrinolytic activity in relation to fatty foods. We already have adequate agents for the prevention and treatment of excessive fibrinolytic activity-for example, aminocaproic acid, para-aminomethylbenzoic acid, and Trasylol (Okamoto and Okamoto, 1962; New Engl. J. Med., 1965; Menon, 1967b). Most of the agents potent in enhancing the fibrinolytic activity are, however, antigenic (for example, streptokinase), very expensive (for example, urokinase), or transient in effect (for example, nicotinic acid), and in any case only effective parenterally (Tillet et al., 1955; Weiner et al., 1958; Celander and Guest, 1960; Menon, 1966b). Research has been carried out in the hope of discovering a cheap nonantigenic drug (Menon, 1967c). Fearnley et al. (1967) investigated the oral antidiabetic group of drugs, sulphonylureas and

diguanides, and found metformin to be the most effective as well as the best tolerated.

Onions are known to be composed of essential oils, allypropyl, disulphide, catechols, thioproprione aldehyde, protocatechuic acid, thiocyanates, as well as some calcium, phosphorus, iron and vitamins (Wealth of India, 1948; Paech and Tracey, 1955; Chopra et al., 1956). Whether the increased fibrinolytic activity in the blood is caused by one of these or by a hitherto unknown component of the onion has not yet been established, though investigations are in progress to clarify this.

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REFERENCES

Billimoria, J. D., Drysdale, J., James, D. C. O., and Maclagan, N. F. (1959). Lancet, 2, 471.
Celander, D. R., and Guest, M. M. (1960). Amer. J. Cardiol., 6, 409.
Chopra, R. N., Nayar, S. L., and Chopra, I. C. (1956). Glossary of Indian Plants, p. 11. New Delhi.
Fearnley, G. R., Chakrabarti, R., Hocking, E. D., and Evans, J. F. (1967). Lancet, 2, 1008.
Greig, H. B. W., and Runde, I. A. (1957). Lancet, 2, 461.
Gupta, N. N., Mehrotra, R. M. L., and Sircar, A. R. (1966). Indian J. med. Res., 54, 48.
Mehrotra, R. M. L., Gupta, N. N., and Mittal S. P. (1966). Indian J. Greig, H. B. W., and Runde, I. A. (1957). Lancet, 2, 461.
Gupta, N. N., Mehrotra, R. M. L., and Sircar, A. R. (1966). Indian J. med. Res., 54, 48.
Mehrotra, R. M. L., Gupta, N. N., and Mittal, S. P. (1966). Indian J. med. Res., 54, 54.
Menon, I. S. (1966a). Lancet, 2, 1365.
Menon, I. S. (1966b). Brit. J. clim. Pract., 20, 561.
Menon, I. S. (1967a). Lab. Pract., 16, 469.
Menon, I. S. (1967b). Brit. J. clim. Pract., 21, 405.
Menon, I. S., Burke, F., and Dewar, H. A. (1967). Lancet, 1, 700.
Menon, I. S., and Dewar, H. A. (1967). Brit. med. J., 2, 613.
Moser, K. M., and Hajjar, G. C. (1966). Amer. J. med. Sci., 251, 536.
Okamoto, S., and Okamoto, U. (1962). Keio J. Med., 11, 105.
Paech, K., and Tracey, M. V. (editors) (1955). Modern Methods in Plant Analysis, p. 714. Berlin.
Schneider, C. L. (1952). Amer. J. Obstet. Gynec., 64, 141.
Tillet, W. S., Johnson, A. J., and McCarty, W. R. (1955). J. clin. Invest., 34, 169.
von Kaulla, K. N. (1963). Chemistry of Thrombolysis: Human Fibrinolytic Enzymes, p. 79. Springfield, Illinois.
Wealth of India, 1948, 1, 56. New Delhi.
Weiner, M., Redisch, W., and Steele, J. M. (1958). Proc. Soc. exp. Biol. (N.Y.), 98, 755.

Experience with a Hepatitis-free Plasma Protein Solution

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Cummary: Clinical experience with a 4.3% solution of plasma protein treated to render it free of the agent of serum hepatitis is satisfactory. Sixty-seven transfusions of 400 ml. of the commercial preparation were given to 33 patients (25 with acute blood loss, 4 with severe burns, and 4 with hypoproteinaemia secondary to hepatic or renal disease).

The solution was clinically as effective as reconstituted dried plasma in expanding plasma volume and in replacing serum protein lost in burns. Adverse effects were mild pyrexial reactions in one case and facial flushing in another. No cases of serum hepatitis occurred.

The solution is available for immediate use, it can be kept at room temperature, and, as it does not cause rouleaux formation, it can be given before blood is taken for grouping and cross-matching.

Introduction

The value of reconstituted dried plasma in the restoration of blood volume by transfusion is well proved. Unfortunately such transfusion carries a slight but definite risk of transmission of disease, serum hepatitis presenting the greatest problem. The incidence of serum hepatitis after transfusion of plasma has been estimated as less than 1% in "small-pool" plasma (Medical Research Council, 1954) and as high as 18% in "large-pool" plasma (Morgan and Williamson, 1943).

It was largely because of the risk of serum hepatitis in transfusion of plasma that synthetic plasma expanders, such as the dextrans, were developed. Certain plasma fractions, particularly albumin, are also thought to be free from the agent causing serum hepatitis (Paine and Janeway, 1952).

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