

APPENDIX

Acute radiation effects in victims of the Chernobyl nuclear power plant accident

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Introduction

1. This Appendix sets out the essential findings of the clinical observation of a group of patients suffering from acute radiation sickness following the accident at the Chernobyl nuclear power plant on 26 April 1986. The observations were conducted at the specialized treatment centre in Moscow over a period of two years.

2. An initial report on the accident was submitted by the Soviet representatives to the Post-Accident Review Meeting held at the International Atomic Energy Agency in August 1986 and was summarized in IAEA Safety Series Technical Report No. 75 [I18] and in [G33]. The proposal to present this information in its present form was endorsed at the thirty-sixth session of UNSCEAR in March 1987.

3. The basic information on the radionuclide releases and the types of exposures of the irradiated persons coincided with the expected pattern for an accident at a nuclear power plant of similar type: as much as 100% of gaseous fraction of the noble gases and nuclides may have escaped from the plant; caesium, iodine and tellurium isotopes accounted for up to 10-20% of the nuclide inventory, and other radionuclides for up to 30% [I18].

4. The plant personnel and auxiliary staff present at the industrial site in the immediate vicinity of the accident zone were subjected to the combined effect of radiation from several sources: (a) short-term external gamma/beta radiation from the gas emission cloud (in

the case of persons in the immediate area of the accident zone at the time of the explosion); (b) external gamma/beta radiation of decreasing intensity, from fragments of the damaged reactor core scattered over the industrial site; (c) inhalation of gases and aerosol dust particles containing a mixture of radionuclides; and (d) deposition of these particles on the skin and mucous membranes at the time of the intensive generation of steam or dust and the wetting of clothing (as a result of them being blown or washed off contaminated objects).

5. However, the most significant factor was the general, external and relatively uniform whole-body gamma-irradiation and the beta-irradiation of extensive body surfaces, coupled (except in two cases) with a very small intake of nuclides through inhalation, predominantly of radioiodine and caesium isotopes. Thus, the basic clinical picture was that of a distinctive acute radiation sickness caused by gamma-irradiation of the whole body and by beta-irradiation of extensive areas of the skin surface.

6. Direct and indirect dosimetry methods were used to determine the nuclide content in the body. A great many tests were carried out, both while the victims were alive and (in 28 cases) after they had died, so that it was possible to estimate the nuclide content in the body and the resultant dose levels. An example of these types of analyses is shown in Figure A.1., giving the distribution of various radionuclides in the lungs.

7. The iodine isotope content in the thyroid gland was determined repeatedly (as many as four to six

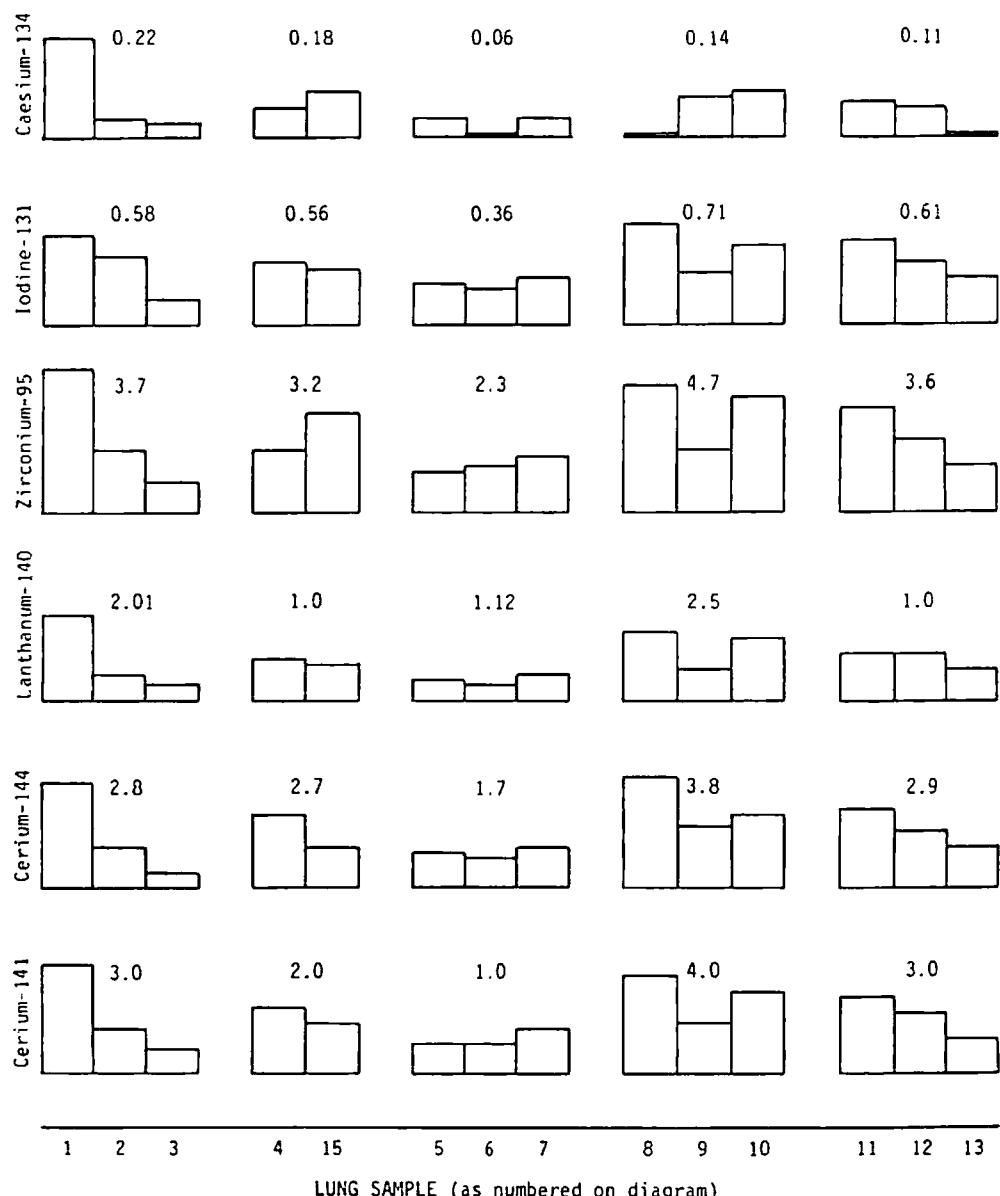
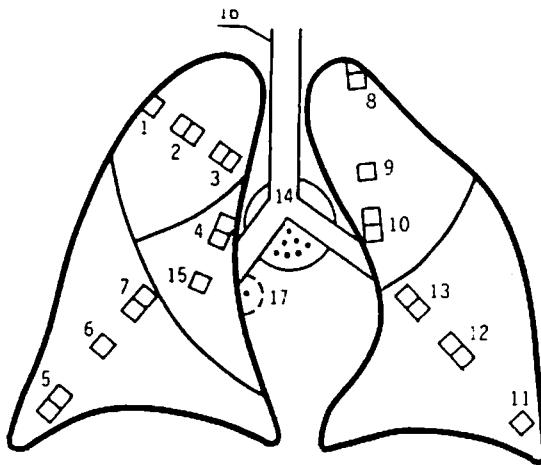


Figure A.I. Location of lung samples taken at the time of autopsy and distribution of the main radionuclides in lung samples. The number above each sample group indicates the approximate relative average value.

times) from the second day after the accident. These measurements showed that ^{131}I accounted for $80 \pm 20\%$ of the total activity of all iodine isotopes, ^{133}I for $15 \pm 10\%$, and the remaining isotopes (^{123}I , ^{124}I , ^{126}I , and ^{130}I) for not more than 2%.

8. The calculations for estimating intake quantities from the thyroid measurements were performed according to the recommendations of the International Commission on Radiological Protection [I19]. On the basis of the distribution of thyroid doses in exposed individuals (Table A.1), it may be stated that in the overwhelming majority of cases the thyroid doses were below the levels likely to cause direct injury to that organ ($< 3.7\text{ Sv}$) or of significantly influencing the clinical picture during the onset of acute radiation sickness. Low radioiodine dose levels were also suggested by the post-mortem nuclide measurements of the 28 persons who subsequently died.

9. Internal dose values according to post-mortem measurements for 6 patients are shown in Table A.2. The maximum amount of ^{137}Cs and ^{134}Cs incorporated activity was 7.4 MBq, except for two patients with extensive steam burns, which allowed intake of nuclides through the wound. The post-mortem dosimetry gave 40 and 80 MBq of ^{137}Cs plus ^{134}Cs , and 450 and 1,100 MBq of ^{131}I , for these two patients, respectively. The whole-body internal doses in these two individuals from these nuclides were estimated as approximately 1 Sv and 2 Sv during the two to three weeks before they died, which are commensurable with their external gamma doses. This fact was taken into account during the interpretation of clinical data. Internal doses for other patients did not exceed 1-3% of the external irradiation doses.

10. The transuranic elements (e.g., ^{239}Pu) were studied in urine specimens from 266 persons (635 analyses), including, in some of the cases, analyses conducted before and after the administration of pentacine. The urine activity values and a negative finding after chelation treatment confirmed the absence of a significant plutonium contamination of all the patients observed. Post-mortem tests by alpha spectrometry for transuranic elements showed their presence (74-300 Bq per organ) only in the lungs; curium accounted for as much as 90% of the specimen activity, and plutonium and americium for 10%.

11. Gamma-spectrometric analysis of the first specimens within 36-39 hours of the accident failed to reveal any sign of $^{22,24}\text{Na}$ activation, which confirmed that neutron irradiation of the victims was not significant.

12. For most of the victims, the energy peaks of more than 20 radionuclides were detectable in the spectrum of their whole-body gamma measurements; however, apart from the iodine and caesium isotopes previously mentioned, the contribution to the overall dose from the others (^{95}Nb , ^{144}Ce , ^{140}La etc.) was negligible. These measurements, performed while the victims were still alive, were also confirmed through the analysis of autopsy specimens (approximately 35 specimens from each deceased person) (Figure A.1).

13. The dose levels from external irradiation were reconstructed from the indication of several measurements on the basis of previous experience. Subsequently, in three cases with a lethal outcome, these findings were refined using methods earlier proposed by Kraytor for clothing fabrics [B31] and according to the electron spin resonance technique for dental enamel [T16]. These measurements agreed within $\pm 20\%$ with the dose estimates based on clinical and biological criteria.

14. The total number of affected individuals among the persons present at the reactor site in the early hours of 26 April 1986 was 203, as given in the report presented by Soviet representatives at the Post-Accident Review Meeting in August 1986 [I18]. Of these, 115 were treated, beginning on day 2, at the specialized treatment centre in Moscow; it was this group that provided most of the scientific analytical data discussed in this report. At other hospitals in Kiev there were only 12 patients with a clearly defined clinical pattern of second-degree acute radiation sickness and one person with fourth-degree acute radiation sickness, a fact that cannot in any substantial way alter the overall assessment of the data for the entire group of victims.

15. The increase in the total number of affected individuals from 203 to 237, announced in November 1986, was due solely to persons suffering from first-degree acute radiation sickness. There were 31 persons suffering from first-degree acute radiation sickness at the special treatment centre in Moscow and 109 persons in Kiev. The task of establishing a diagnosis distinguishing between first-degree acute radiation sickness and ordinary somatic diseases according to generally accepted criteria is a complex one, and one that continued throughout 1986. On the whole, a critical analysis of the data shows a decrease in the number of persons suffering from first-degree acute radiation sickness in comparison with the number given originally. At the time of writing this report, up to three quarters of these persons are for all practical purposes healthy. Their clinical signs of reaction to the accident during the first three months were neither individually significant nor typical of a reaction to irradiation. Table A.3 shows the distribution of patients with acute radiation sickness according to its degree of severity [B31] in the group selected for scientific analysis.

A. INITIAL DIAGNOSIS OF ACUTE RADIATION SICKNESS

16. The medical unit serving the plant was informed of the accident within 10-15 minutes of its occurrence. First aid to the affected individuals was provided by middle-level medical personnel and emergency teams over a time period from 30-40 minutes to 3-6 hours after the accident. First aid consisted in the evacuation of the victims from the industrial site, the simplest forms of medical attention, the administration of antiemetic and symptomatic (sedative, cardiotonic) drugs, the distribution of potassium iodide and the transportation of persons suffering from a pronounced primary reaction to the medical unit. During the first

12-24 hours after the accident, other persons who were in satisfactory condition were urged to go to the medical unit for examination; a total of 132 persons were hospitalized there during the first 12 hours. One person with severe thermal burns died during the first hour. Another, a reactor operator, could not be found; his working station was located in the collapsed high-activity zone.

17. Within 12 hours, a specialized emergency team arrived at the site and began work. Within 36 hours, this team, together with the on-site medical unit, examined more than 350 persons and carried out approximately 1,000 blood tests, each person undergoing two to three such tests. The treatment with potassium iodide was continued.

18. Within the first three days, 299 persons suspected of suffering from acute radiation sickness were sent to the specialized treatment centre in Moscow and to hospitals in Kiev, and over the subsequent days some 200 additional persons were admitted for examination.

19. The primary diagnostic criteria for assessing the priority for hospitalization were the presence, time of onset and intensity of nausea and vomiting and of primary erythema of the skin and mucosae, and a decrease of the lymphocyte count in the peripheral blood to below 10⁹/l during the first days following irradiation.

20. The diagnosis of acute radiation sickness was subsequently confirmed in 99 of the 128 persons (firemen, Unit 4 operators, turbine-room duty officer and auxiliary personnel) admitted to the specialized treatment centre in Moscow during the first two days and in six of the 74 victims hospitalized during the following three days. This is an indication of the high specificity of the screening methods used. An additional 10 cases of minor acute radiation sickness were diagnosed among persons present at the site at the time of the accident, who were later admitted to the hospital facility for a variety of reasons. In the reception area the patients were monitored again for contamination and, if necessary, subjected to decontamination measures (washing under a shower with ordinary soap and change of underwear). Blood and urine samples were taken for a quick test of the presence of radionuclides; the patients also underwent measurements (repeated a further 4-6 times during the first 6-10 days) of the radioactive iodine content in the thyroid. Measuring devices consisting of a scintillation detector or a semiconductor detection unit were used for the whole-body counting of radionuclide activity.

B. THE BONE MARROW SYNDROME AND ITS TREATMENT

21. Dosimetric data, together with an analysis of the circumstances of the accident and the presence in a considerable number of the victims of obvious primary reaction symptoms (nausea, vomiting, diarrhoea, haemorrhage of the mucosae and skin, lymphopenia), confirmed that the principal modes of irradiation had been: (a) by external, relatively uniform gamma-radiation; and (b) by deposition of beta/gamma-emitting nuclides

on the skin. Radionuclide ingestion was below the level likely to cause acute radiation injury. As already noted, two patients suffered from all three of these irradiation modalities, in combination with extensive steam burns.

22. The important diagnostic task during the first few days after the accident was the assessment of the degree of severity of the bone marrow syndrome resulting from the external gamma-irradiation dose. This was possible through the use of previously devised methods, which are based on the number of lymphocytes and on chromosome aberrations in peripheral-blood lymphocytes or on the incidence of chromosome aberrations in bone marrow cells [B31, B50, G24]. These data were later transformed into a prognosis of the overall dynamics of the blood picture. A subsequent reassessment of dose levels involving a larger sample of cells scored revealed not more than 5-10% changes in the estimated doses.

23. Dose-effect relationships for these indicators had been derived earlier through the analysis of relatively uniform accidental or therapeutic irradiations of human subjects having normal initial haematological characteristics and exposed to well established doses [P22]. Figure A.II shows the curves (and analytical expressions) for the relationships between the dose and the blood lymphocyte count for each of the first nine days and the average lymphocyte count on days 4-7 and days 1-8 after irradiation. The radiation dose received by each person was estimated according to the number of chromosome aberrations (dicentrics) in a blood-lymphocyte culture, using a dose-effect curve for 100 first-mitosis cells that had been obtained after whole-body gamma-irradiation to treat acute leukaemia patients during a period of full clinical and haematological remission [P23].

24. The formula for calculating of the dose is as follows:

$$D = (-a + \sqrt{a^2 + 4by})/2b$$

This assumes that the yield of dicentrics shows a linear-quadratic dependence on dose:

$$y = (a \pm 2.24)D + (b \pm 0.56)D^2$$

where D is the average gamma-irradiation dose in the body (Gy), y is the dicentric count per 100 cells; a = 8.36; b = 5.70.

25. Up to day 7 after the accident, the estimates of the average dose of total gamma-irradiation were refined, mostly on the basis of the peripheral-blood lymphocyte counts but also, in the more severe cases and to a lesser degree, on the chromosome aberration count. This made it possible to divide the patients into various prognostic groups [B31], according to the severity of the bone marrow syndrome as follows (see Table A.3):

(I)	slight	(1-2 Gy)
(II)	intermediate	(2-4 Gy)
(III)	severe	(4-6 Gy)
(IV)	extremely severe	(6 Gy and above)

It was also possible to separate those persons who received doses of less than 1 Gy.

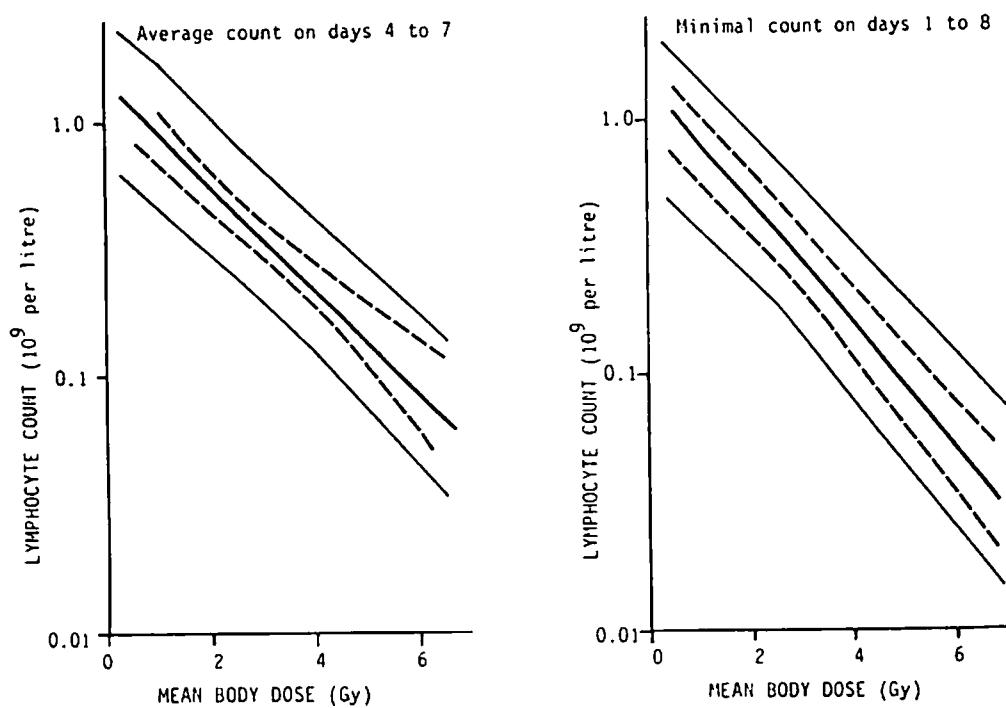
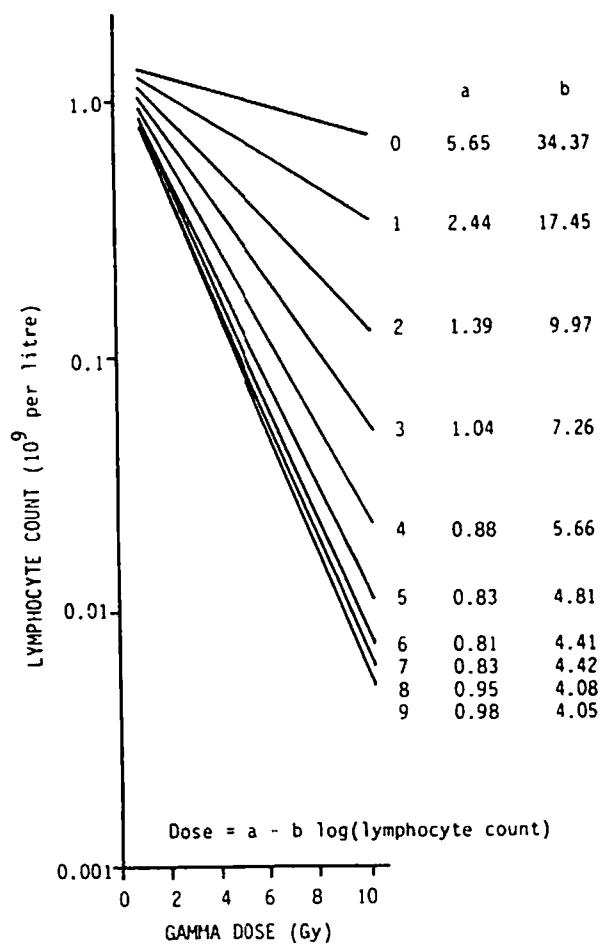


Figure A.II. Estimation of the total gamma dose according to the blood lymphocyte counts. Upper panel: Dose-effect relationships for lymphocyte counts at the days post irradiation shown on the curves; analytical expression and coefficients of these relationships. Lower panels: Curves showing the dependence of average lymphocyte counts on days 4-7 and the minimum lymphocyte count on days 1-8 as a function of the irradiation dose.

26. Particular attention during the first days was directed at identifying persons with an extremely severe and irreversible degree of myelodepression, for whom an urgent decision was required regarding a bone marrow transplant. Additional signs providing further evidence that a patient belonged to this group were (a) vomiting during the first half-hour and of diarrhoea during the first 1-2 hours from the start of irradiation; (b) a swelling of the parotid glands during the first 24-36 hours; and (c) the ascertainment of an irreversible degree of myelodepression using a diagnostic table previously devised (Table A.4).

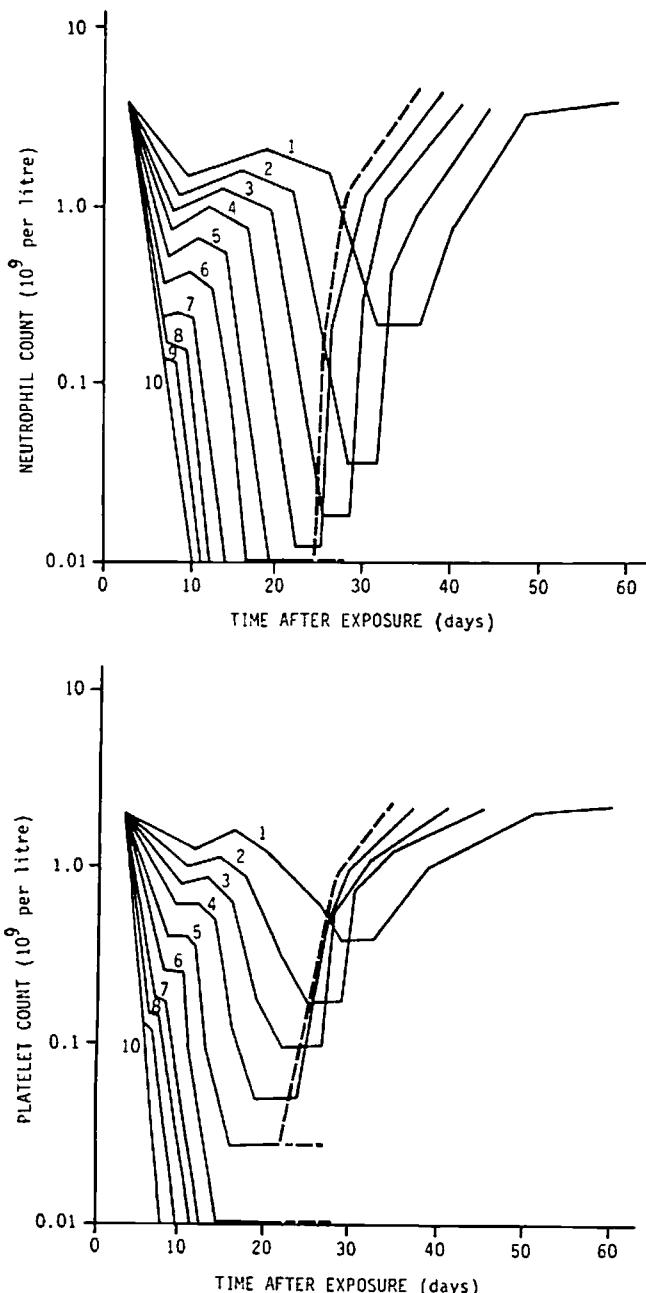


Figure A.III. Standard curves showing the changes of the neutrophil and platelet counts after various doses (numbers on the curves indicate the dose in Gy) in the case of relatively uniform whole-body gamma irradiation of human subjects. [B31]. (The broken segments of the curves at doses of 5-6 Gy indicate that recovery may not occur at these times in all patients.) Upper panel: Neutrophils. Lower panel: Platelets.

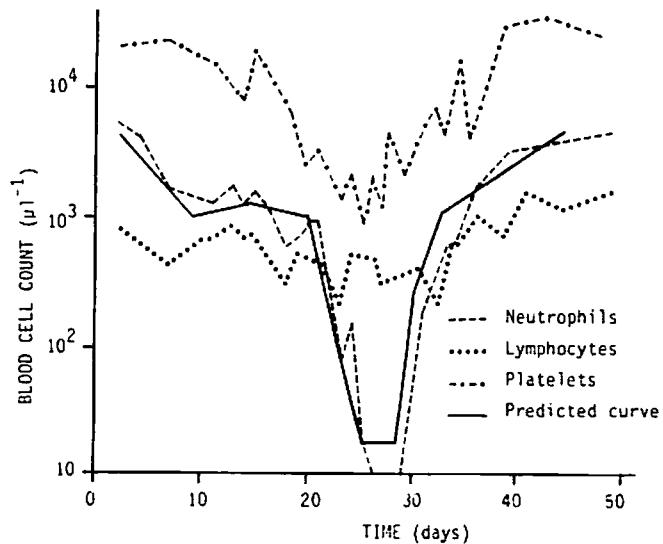


Figure A.IV. Example of the changes in neutrophils, lymphocytes and platelets observed in a patient (case 39) suffering from acute radiation sickness (estimated dose 2.4-3.3 Gy) and the predicted neutrophil curve for a total gamma dose of 3.0 Gy.

27. The results of numerous biochemical, immunological and biophysical indicators are undergoing processing and analysis at the time of writing this report. None of these indicators is as informative as the signs described above. However, it may be noted, for example, that hyperamylasemia was used as a supplementary prognostic test.

28. On the basis of the estimated dose, a prediction was made using standard curves [P22], of the overall trend with time in the neutrophil and platelet counts (Figure A.III). By way of example, Figure A.IV shows the real and predicted neutrophil curves for one patient (case 39). The total gamma-irradiation doses estimated according to the average lymphocyte count from day 4 to day 7 and according to the dicentric yield totalled 2.4 and 3.3 Gy, respectively. The patient's measured neutrophil curve almost coincided with the predicted neutrophil curve for 3.0 Gy of total gamma-irradiation.

29. The calibration curves of neutrophil counts as a function of dose were also used for a final assessment of the magnitude of the total gamma-irradiation dose. The dose calibration curve was chosen that coincided with the measured second depletion phase. Also, use was made of the dose dependency of the second depletion phase, such as the time required for the neutrophil count to decrease to $0.5 \cdot 10^9/\text{litre}$ or the time to reach the minimum of the second depletion phase (Figure A.V).

30. For relatively low doses (1.0-1.5 Gy), the diagnosis was finally established over time periods of up to three months only in those cases that showed typical post-irradiation neutrophil and/or platelet time courses with distinct second depletion and restoration phases. The latter generally developed beginning in the fourth to the fifth week after irradiation. In order to determine these changes, it was necessary to carry out blood analyses not less than two or three times a week

over a period of two to three months. Examples of these curves are shown in Figure A.VI: (a) case 48: doses, estimated according to the lymphocyte count on days 4-7 and the dicentric yield were 1.1 Gy and 1.4 Gy, respectively; and (b) case 97: doses, estimated according to the lymphocyte count on day 9 (the first blood analysis was made at this time because of late arrival) and the dicentric yield were 0.3 Gy and 0.9 Gy, respectively. It should be noted that in the case of low doses, the minimum level for neutrophils occurred later (day 30-50) than for platelets (day 20-40), and the reduction in the number of platelets and their recovery were more clearly pronounced than for neutrophils.

31. On the basis of all these data, a diagnosis of acute radiation sickness with the bone marrow syndrome of the first, second, third and fourth degree of severity was definitively established for 31, 43, 21 and 20 patients, respectively (see Table A.3). The analysis of the observations carried out on these patients is the subject of the exposition that follows.

32. Clinical manifestations of the bone marrow syndrome corresponded to the level and duration of post-irradiation pancytopenia (neutrophils $0.1-0.5 \text{ } 10^9/\text{l}$, platelets $10-20 \text{ } 10^9/\text{l}$). The main signs were fever, infectious complications and petechial haemorrhages in the skin and oral mucosa.

33. Treatment was based on the principles of supportive therapy, including isolation, antimicrobial decontamination of the intestine, administration of systemic antibiotics and replacement transfusions of

blood cells. In cases in which there was a prognosis of irreversible myelodepression, transplantations of allogeneic bone marrow and embryonic human liver cells were performed.

34. All patients suffering from a bone marrow syndrome of the second, third or fourth degree were individually accommodated in ordinary hospital rooms. These were adapted to ensure (a) barrier nursing; (b) air sterilization by means of ultraviolet lamps; (c) strict observance by the attending personnel of hand disinfection on entering and leaving the room; (d) mandatory use of individual or disposable gowns, masks, and caps; (e) antiseptic decontamination of footwear; (f) changes of underclothing for patients at least once a day; (g) use of antiseptic agents for washing the walls and floor of the room and the items of use; and (h) individual assignment of antiseptically treated nursing items in the room. This regimen made it possible to maintain the micro-organism population at less than 500 m^{-3} in the room air. Ordinary food was served, with the exclusion of raw vegetables, fruits and canned products.

35. Prophylaxis against endogenous infections was by means of the internal administration of biseptol-480 and nistatin in amounts of six tablets and five million units per day, respectively, for one and 2-3 weeks prior to the development of agranulocytosis (leucocytes $1.0 \text{ } 10^9/\text{l}$, neutrophils $0.1-0.5 \text{ } 10^9/\text{l}$).

36. With the onset of fever, intravenous administration of two or three broad-spectrum antibiotics was prescribed, one of them being from the aminoglycoside

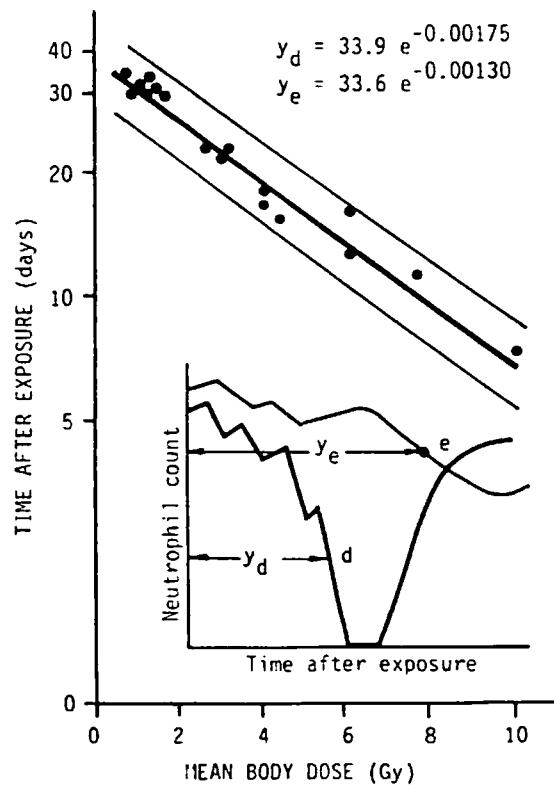
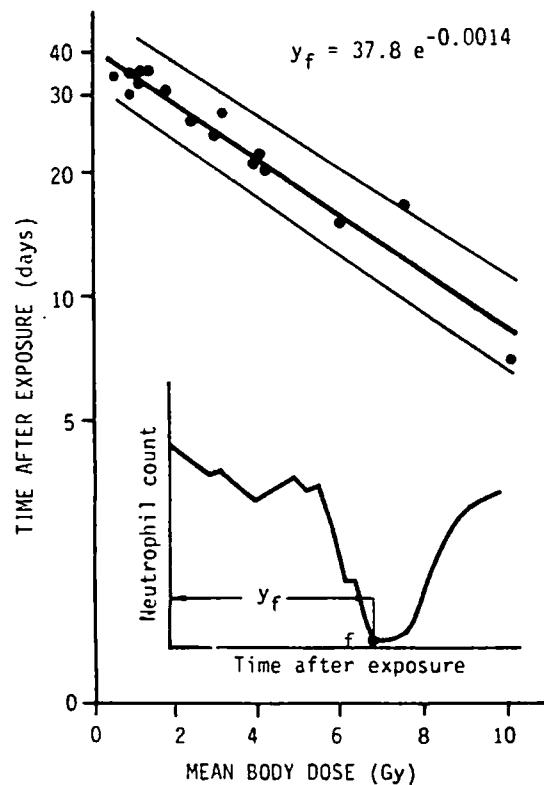


Figure A.V. Estimation of the total gamma dose according to two neutrophil counts: Left panel: time to the minimum of the second phase of depletion; Right panel: time to the "500 neutrophil day" or to the middle of the second depletion.

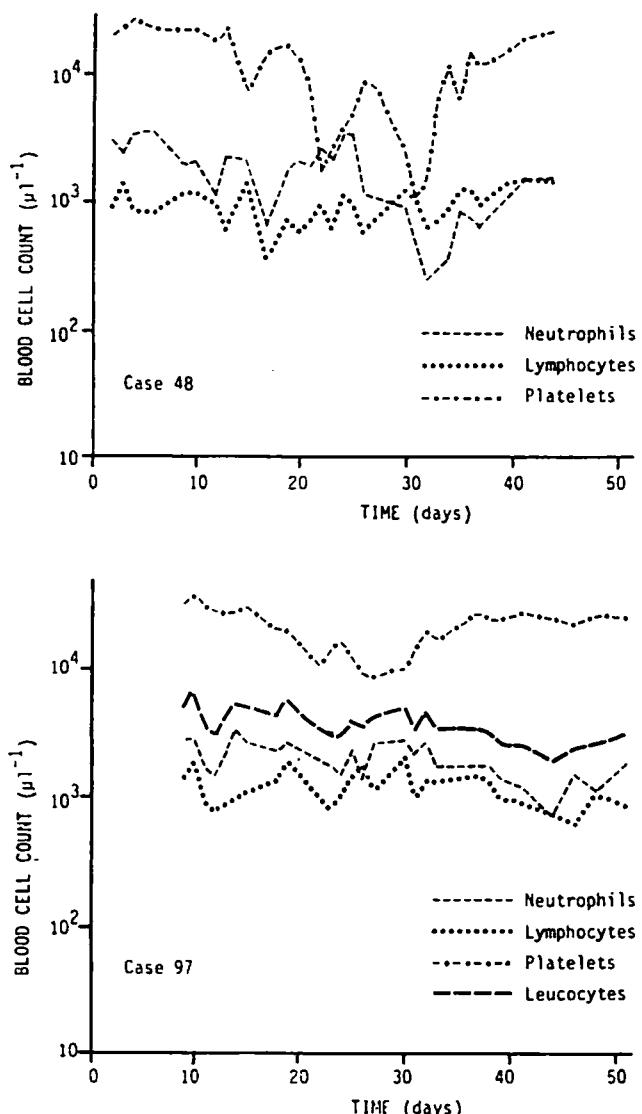


Figure A.VI. Changes in neutrophil, lymphocyte, platelet and leucocyte counts after whole-body gamma irradiation. Case 48, estimated dose 1.1-1.4 Gy. Case 97, estimated dose 0.3-0.9 Gy.

group (gentamicin or amicacin), the cephalosporins (cephzol, cephamecin, cephobide) and the semi-synthetic penicillins with activity against *pseudomonas aeruginosa* (carbenicillin, pipracil), all in maximum doses. This treatment reduced the fever in more than half of the patients. If within 24-48 hours there was no effect, extensive use was made of gamma globulin (Sandoglobulin), made available by the Sandoz Company (Switzerland). Six grams were administered intravenously every 12 hours, three or four times. The policy adopted was one of early empirical prescription of 1 mg/kg per day amphotericin-B given intravenously, if the fever had not disappeared within a week of the above mentioned antibiotics, in combination with the intravenous administration of gamma globulin.

37. In this situation, acyclovir was used for the first time, and with good effect, in the treatment of patients with acute radiation sickness suffering from a herpes simplex infection. Not less than one third of the patients with third- and fourth-degree acute radiation sickness were affected by this virus. Acyclovir was not

used prophylactically; experience has shown that this should be the case with high-dose whole-body irradiation. An ointment containing acyclovir proved effective in the treatment of skin lesions involving herpes virus.

38. The regimen described above for the empirical treatment to combat infection proved to be highly effective; there was no evidence that deaths were caused by bacterial infection alone in patients suffering from the bone marrow syndrome. This was so even among those with a severe or extremely severe form of acute radiation sickness, provided it was not complicated by burns, radiation-induced enteritis or acute secondary syndromes as a result of a bone marrow transplantation.

39. Blood screenings carried out while the patients were still alive or, in the case of those who died, posthumously, most often revealed epidermal staphylococcus. It will only be possible to evaluate the role of this form of bacteria as an agent of terminal septicaemia when these results have been compared with the histological data from analysis of various organs that have not yet been completed.

40. One of the undoubtedly successes in the treatment of the bone marrow syndrome in patients with acute radiation sickness was the rational use of fresh donor platelets for the prophylaxis and treatment of bleeding. To make these measures possible, the collection of the thrombocytes was organized on an urgent basis, using the method of fourfold thrombocytapheresis from individual donors at seven blood transfusion centres. For one transfusion, platelets obtained from a single donor (on the average $300 \cdot 10^9$ platelets in 200-250 ml of plasma) were used. Transfusions were carried out when the platelet level in the blood fell to $20 \cdot 10^9/l$ or lower, with appearance of the first signs of bleeding. The infusions were repeated every 1-3 days. As a prophylaxis against acute secondary syndrome, the thrombocytes as well as the other blood components were irradiated with 15 Gy before infusion in order to inactivate the immunocompetent cells originating from the donor.

41. Platelet transfusion prevented life-threatening bleeding, even in patients with protracted (more than 2-4 weeks) and severe thrombocytopenia. The majority showed no signs of bleeding at all, although autopsies disclosed microcirculatory failures and porosity of the capillaries in a number of organs. In this situation, successful use was made of cryo-preserved allogeneic and, what is particularly important, autologous thrombocytes. The latter were obtained from patients with second- or third-degree bone marrow syndrome during the first days following irradiation (1-2 sessions) — this had no effect on the post-irradiation behaviour of their platelet counts — and were used with great effectiveness when the patients developed critical thrombocytopenia. No cases of refractoriness to thrombocytes transfusion were observed. On the average, from three to eight transfusions of standard amounts of thrombocytes ($300 \cdot 10^9$ cells) were required for the treatment of a single patient with second- and third-degree acute radiation syndrome. Leucocytes

were not used for the prophylaxis or treatment of agranulocytic infections. Requirements for erythrocytes turned out to be considerably greater than expected, even for patients with second- and third-degree acute radiation sickness uncomplicated by severe radiation burns.

42. Figure A.VII illustrates the clinical record of one patient (case 21), showing the typical extent and duration of substitutive and supportive therapy for the bone marrow syndrome. In the case of this patient there were no life-threatening acute radiation injuries to the other tissues.

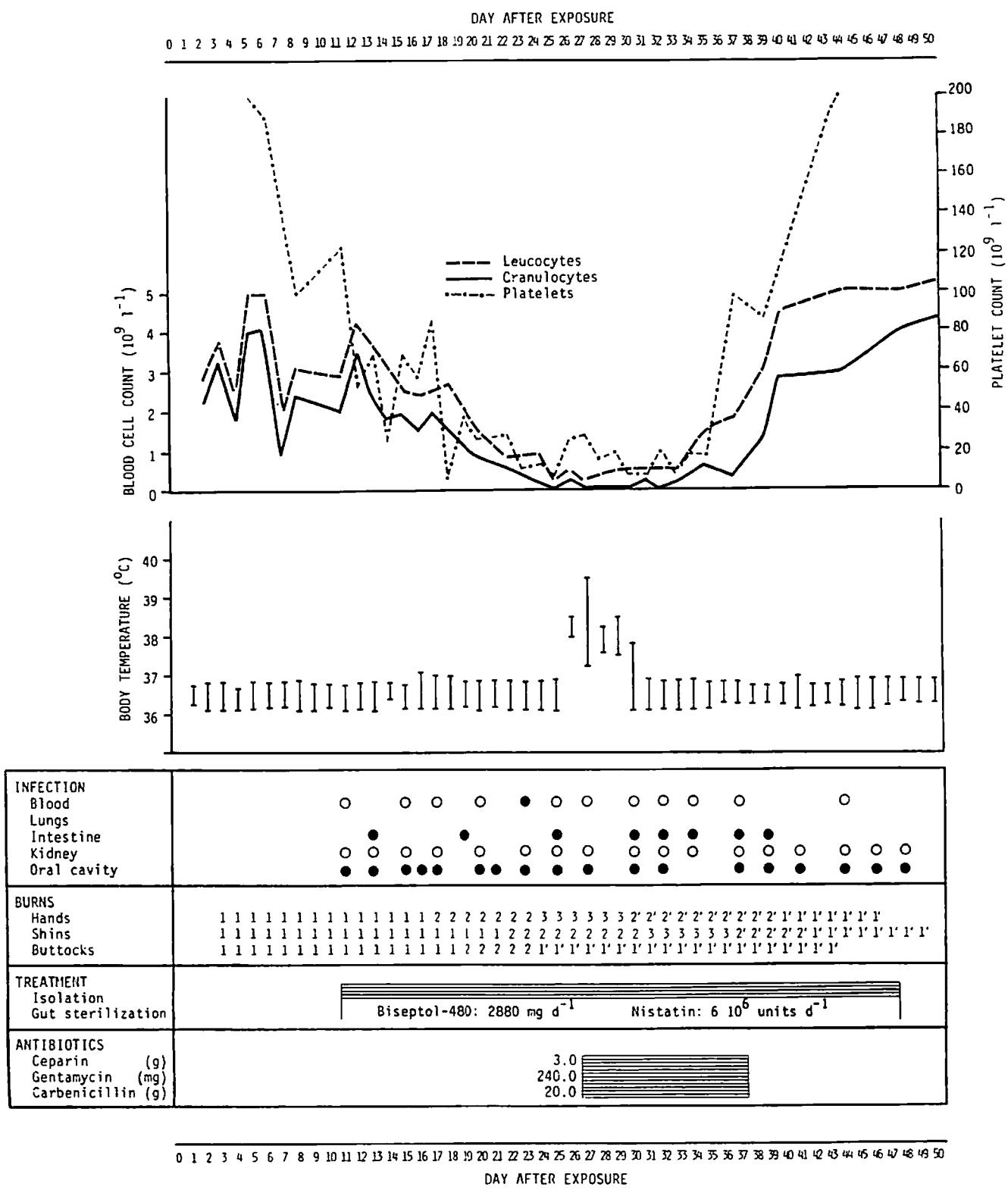


Figure A.VII. Clinical record of a patient (case 21) suffering from acute radiation sickness with bone marrow syndrome of an average degree of severity; whole-body gamma dose of 3.9 Gy, estimated according to the chromosome aberrations in blood lymphocytes. The circles in the infection section indicate the results of bacteriological tests of biological samples (blood, mucus, faeces, etc.); empty circles indicate absence of growth; solid circles indicate the presence of bacteria or fungi. The numbers in the burn segment designate the three degrees of radiation-induced burns.

43. The indication for an allogeneic bone marrow transplantation or an embryonic liver cell transplantation was the whole-body gamma-irradiation dose, estimated according to the peripheral-blood lymphocyte count and the chromosome aberrations at about 6.0 Gy and above. At these dose levels the prognosis expected was for irreversible or extremely protracted severe myelodepression.

44. A total of 13 allogeneic bone marrow transplants and six embryonic liver cell transplants were performed. The latter contains haemopoietic stem cells and a minimum of immunocompetent cells, which sharply lowers the risk of an acute secondary syndrome.

45. Seven patients who received allogeneic bone marrow transplant died between two and 19 days (15-25 days after irradiation) from acute radiation injuries to the skin, intestines and lungs.

46. Of six patients who did not suffer fatal skin burns and intestinal injuries and whose total doses had been estimated at between 4.4 and 10.2 Gy, two survived allogeneic bone marrow transplants (gamma-ray doses of 5.6 and 8.7 Gy). Both had haplo-identical female donors (sisters), rejected the partially functioning transplant (at days 32 and 35) and experienced a restoration of their own myelopoiesis, beginning at day 28.

47. Four patients who received allogeneic bone marrow transplantation died between 27 and 79 days (34-91 days after irradiation) from mixed viral-bacterial infections. Two of them had effectively functioning HLA-identical transplants (cases 6 and 28: total gamma-ray doses 5.2 Gy and 6.4 Gy, respectively), and two had early rejection (day 16 and 42) of "haplo + 1" and haplo-identical transplants, but during times when their own myelopoiesis was restored (cases 5 and 16: gamma-ray doses 4.4 and 10.2 Gy).

48. Similarly, all the patients who received embryonic liver cell transplants died from skin and intestinal injuries within a brief period (14-18 days after irradiation), with the exception of one woman of 63 years (case 8, embryonic-liver cell transplant from an 18-week male donor), who lived 30 days, having received a dose of 8-10 Gy. At death (17 days after transplant), numerous mitoses were discovered, against a background of severe marrow pancytopenia, and all the cells had a female karyotype, i.e., regeneration of the host bone marrow had begun.

49. This experience confirms that in an emergency situation like the one described, the group of persons for whom bone marrow transplants would be indicated with a reasonable prospect of success is extremely limited. Seven of 13 patients died as a result of skin and intestinal injuries before bone marrow engraftment could be expected. In Table A.5 a comparison is given of survival or cause of death of patients receiving bone marrow transplantations and of patients in a control group.

C. OTHER INJURIES AND THEIR TREATMENT

50. The extensive skin lesions caused by beta-radiation represented a distinctive feature of the injuries suffered in this emergency situation. Radiation-induced skin burns in firemen and personnel from the plant were observed only in combination with radiation injury to haemopoiesis and were therefore an integral part of the general acute radiation sickness.

51. This situation may be regarded as one in which there is an extremely non-uniform distribution of dose as a function of the depth of penetration within the body; the skin doses are estimated to be 10-20 times greater than the bone marrow doses. There was a definite correlation in the severity of the injuries in both tissues.

52. Table A.6 and Figure A.VIII show the distribution of cases involving radiation-induced skin burns of various degrees in patients with acute bone marrow syndrome of different severity. Skin injuries were observed in more than one half of the patients and in virtually every patient suffering from third- or fourth-degree bone marrow syndrome.

53. The aggravating contribution of radiation-induced skin injuries to the overall clinical picture arose not only from their severity, but also from the duration of the injuries, characterized as they are by recurrences of the pathological process. As a rule, the burns occurred at different times on various parts of the body. The most frequent locations during the early period were the wrists, the face, the neck and the feet; later, lesions appeared also on the chest and back, and still later on the knees, hips and buttocks. Exceptions to this sequence were encountered in individual cases.

54. The development of the injury was similar to that described by Cronkite et al. [C16], but in a more severe form. The diffuse hyperaemia in the first few days (primary erythema) was followed after 3-4 days by a period of latency. Secondary erythema in the more severe cases developed after 5-6 days and in the majority of patients from day 8 to day 21. Depending on the degree of the injury, it reached a level of dry (first-degree radiation burn) or moist desquamation with the development of blisters (second-degree burn), or the formation of vesicular-ulcerated and ulcerated-necrotic dermatitis (third- to fourth-degree burn). The re-epithelialization of the desquamated surfaces continued for two or three weeks from the occurrence of the visible injury to the skin. In six patients, the healing of the burns over skin areas involving deep necrosis did not begin until the end of the second month. A characteristic feature in the time course of the burns, and one which could be monitored throughout in this group of victims, was the appearance of recurrent waves of erythema, beginning by the end of the fourth week and continuing up to days 45-60. These changes were characterized by hyperaemia on the previously unaffected skin areas or by the increase in clinical signs of injury at the foci of the primary lesions then in the process of healing. For example, late secondary erythema appeared in the area of the

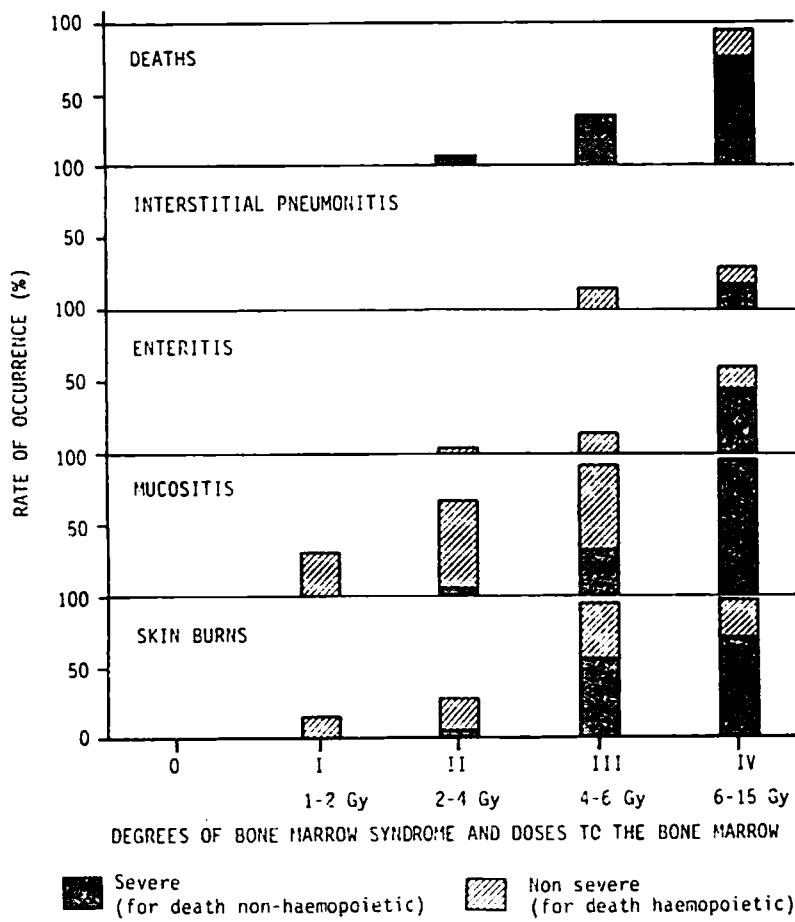


Figure A.VIII. Frequency of deaths and non-bone marrow syndromes for various degrees of bone marrow syndrome in accident victims suffering from acute radiation sickness.

ankles and feet, or on the hips and buttocks, of those patients who during the first three weeks displayed "flowering" burns on their knees. By the time of appearance of this late erythema, the lesions that had occurred earlier in many instances had already been repaired. As a rule, late erythema was accompanied by oedema of the subcutaneous tissues, which was particularly noticeable when located on the knees: pain was experienced in walking; palpation of the skin and underlying tissues (muscles, tendons) caused discomfort. The most severe cases involved fever and a general worsening of the patient's condition. Late secondary erythema was successfully resolved within two weeks by purely topical treatment, although in the more severe cases it was necessary to resort to additional therapeutic means, such as the prescription of glucocorticoids, a form of treatment that fairly rapidly eliminated all manifestations of epidermatitis and subcutaneous oedema, both general and local.

55. As may be seen in Table A.6, the burns suffered by the patients with acute radiation sickness covered from 1% to 100% of the body surface. It may be noted, in this connection, that if there were relatively early (from day 5-6) second- or third-degree burns over an area of even 30-40% of the body, followed by the spread of hyperaemia, these burns were life-threatening. In 19 of the 56 patients suffering from burns, the burns proved fatal (Figure A.VIII). It was found that patients with early secondary erythema over a body area of more

than 40% first developed a febrile-toxic syndrome, followed by renal-hepatic insufficiency and encephalopathic coma with cerebral oedema, resulting in death at 14-48 days after irradiation. A causal link connecting the fatal renal-hepatic insufficiency and the encephalopathic coma to the skin injuries is confirmed by the fact that a similar development of such fatal syndromes was observed in several patients who had neither severe bone marrow syndromes nor intestinal syndromes. However, in the majority of cases, burns were combined with an extremely severe bone marrow syndrome and severe acute enteritis, and in some cases the burns may have been the primary cause of death.

1. Intestinal syndrome

56. The intestinal syndrome was one of the more threatening manifestations of acute radiation sickness. In 10 patients, diarrhoea was observed from day 4 to day 8. This suggested that these persons had received total gamma doses of about 10 Gy or above; all these patients died during the first three weeks following irradiation. The occurrence of diarrhoea after eight days in seven other persons was an indication that they had received lower doses. The presence of radiation-induced enteritis lasting from day 10 to days 18-25 in spite of intensive water-electrolyte-protein supportive treatment suggests that the intestinal syndrome was not the main cause of death.

2. Oropharyngeal reactions

57. Acute radiation-induced inflammation of the oral and pharyngeal mucosa was observed in 82 patients. Its more benign manifestations (first and second degree of severity) were characterized by desquamation and oedema of the mucosa in the area of the cheeks and tongue and by tenderness of the gums. These were observed in 42 persons (dose range 1.7-4.0 Gy) from days 8-9 to days 20-25. The basic signs of a more acute oropharyngeal reaction were observed in 40 patients with third and fourth degree acute radiation sickness (dose range 4.5-16.0 Gy), and these were erosions and ulcers of the oral mucosa, sharp pain, and a large production of rubber-like mucus occasionally blocking the throat and causing breathing problems. The first signs appeared as early as days 3-4, attained their maximum intensity by day 10 and then subsided after days 18-20, when there was also granulocytopenia. The process involved no selective localization, as is characteristic of the ulcerated lesions in the area of the tonsils and gums when there are infectious complications. However, in a significant number of cases, the radiation-induced inflammation of the mucous membranes was complicated by secondary microbial and viral infection, which prolonged its course.

58. Another typical finding was the early (days 3-4) appearance of herpes-like rashes forming massive crusts on the lips and facial skin; this was observed in nearly 30% of the patients with severe bone marrow syndrome. Within this group of patients, primarily those suffering from fourth-degree acute radiation sickness, a pronounced radiation-induced parotitis was observed, coupled with an inability to salivate and a high level of amylase in the blood from days 1-4. The swelling of the parotid glands disappeared without special treatment, whereas recovery of salivary gland secretion was slower.

3. Lung reactions

59. Lung reactions were observed in seven patients suffering from third- and fourth-degree acute radiation sickness. Its characteristic signs were a rapidly intensifying dyspnoea together with respiratory insufficiency progressing over a period of two to three days culminating in death. Autopsies revealed large, blue lungs with pronounced interstitial oedema, without destruction of the mucous membranes of the trachea and bronchi. As a rule, interstitial pneumonitis developed several days before death, generally in combination with extremely severe lesions of the skin and the intestine. The times to death were 14-30 days after irradiation.

4. Causes of death

60. The frequency of non-haemopoietic injury increased as a function of total dose (Figure A.VIII). Clinical observations indicated the essential role of skin injuries in pathological processes prior to death. Among the patients that died, in two thirds of them there was extensive and severe radiation and thermal skin burns, which were considered life-threatening. In

five cases skin injuries were the sole cause of death, because there was neither radiation enteritis or irreversible myelodepression. Deaths were observed between 10 and 96 days after exposure. The clinical picture in all fatal cases was characterized as following a difficult course, because in every case two or three radiation syndromes had occurred with complex toxicity, infection and circulation disorders. A summary listing of patient identification and causes of death is given in Table A.7.

61. A detailed clinico-morphological analysis made it possible to identify the predominance, within specified time periods, of particular lethal syndromes. Up to day 24, a total of 19 patients (65%) died. In one half of these patients the competing causes of death were skin and intestinal reactions (cases 3, 4, 10, 14, 15, 17, 20, 23, 26, 2097; in all the cases the gamma-radiation dose in the bone marrow was estimated to be greater than 10 Gy). Four patients showed acute radiation injury in the lung (cases 2, 9, 12 and 27; the doses were, respectively, 9.2, 9.7, 9.3 and 8.3 Gy), and, of these, two (cases 2 and 9) suffered also severe injuries to the skin. Two patients died of combined thermo-radiation burns (cases 24 and 25; the gamma-radiation doses in the bone marrow were estimated to be 3.7 and 5.7 Gy, respectively, in combination with internal irradiation doses. Within this time-frame, one patient (case 62, dose about 6 Gy) died almost exclusively from severe radiation burns at a time when haemopoiesis had begun to be restored. Three cases (cases 17, 26, 62) had involvement of mycobacterial sepsis. One patient (case 30, dose about 5.5 Gy) died of bleeding caused by mechanical injury to the sub-clavicular vein during catheterization, and another (case 7, dose about 4.7 Gy), suffering from severe radiation injury to the skin, died of post-transfusion shock. Characteristics of the deaths on days 11-24 were marked circulatory problems during the terminal period. This was shown by the relatively high frequency of signs indicating cerebral oedema and focal haemorrhaging into the brain and spinal cord.

62. Six patients died during the period from days 25-48. All six cases were characterized by extremely severe complications of a toxic or infectious nature. Two patients (cases 31 and 34) involved subtotal skin injuries (bone marrow doses about 6.7 and 5.8 Gy, respectively, with death occurring on days 32 and 48, respectively), coupled with practically restored haemopoiesis. The immediate cause of death in these cases was severe respiratory insufficiency and cerebral oedema. In one patient (case 28, dose about 6.4 Gy; death on day 48) the cause of death was severe, graft-versus-host disease and fungal and viral infections. One additional patient (case 5, dose about 4.4 Gy) died on day 34 of severe pulmonary and renal insufficiency, caused, most likely, by the transplantation of HLA-non-identical bone marrow and by post-transplant immunosuppression using cyclosporin and methotrexate. Two patients (cases 1 and 8, doses about 6.6 and 8.3 Gy) died on days 25 and 30, respectively, with symptoms of severe toxicity and pulmonary insufficiency. In nearly all six cases there were marked circulatory disorders in the lungs, intestines, brain and myocardium.

63. At a relatively late stage, days 86-96, three patients died. One patient (case 6, dose about 7.5 Gy) died on day 86 of graft-versus-host disease complicated by cytomegalovirus (CMV) infection. Cytomegalovirus infection was also the cause of death for another patient (case 16, dose about 10.1 Gy) on day 91. A female patient (case 33, dose about 4.1 Gy) died on day 96 displaying marked disruptions of cerebral blood circulation against a background of renal-hepatic insufficiency and foci of mycococcal infection (pneumonia). This patient suffered also skin injuries from beta-radiation which extended over one third of her skin surface and underwent a severe recurrent wave of erythema with oedema of the subcutaneous tissue.

5. Eye damage

64. Eye injuries were characterized by the early and subsequent involvement of all eye tissues in the pathological process (Table A.8). In this group of patients, damage to the skin and eyelid conjunctiva was caused, to a considerable degree, by beta-radiation.

65. At doses not exceeding 1 Gy there were no visible alterations in the structure of the eyes. In the case of patients suffering from first-degree acute radiation sickness, changes were noted only in the front segment of the eye: there was in individual cases a slight erythema in eyelid skin during the first two to four days and an intensification of the vascular pattern in the lid and conjunctiva of the eyeball. In 40% and 100% of the patients suffering from second- and third-degree acute radiation sickness, respectively, the eyelid skin showed a first wave of erythema within 6-12 hours of irradiation, and within 2-3 weeks there was a second wave. These cutaneous alterations disappeared without trace, leaving hyperpigmentation and scaling. In all patients suffering from fourth-degree acute radiation sickness, the times to the appearance of the first and second wave of erythema were 1-2 hours and 8-10 days, respectively.

66. Microscopy of the bulbar conjunctiva revealed a number of alterations in the microcirculation: there was a dilation of the venules and capillaries (more rarely the arterioles), and an increase in the number of functioning vessels coupled with a reduced blood flow.

67. Two patients suffering from combined radiation and thermal second-degree lesions on the lid skin and conjunctiva experienced ulcerations on the skin around the eye that did not re-epithelialize for a long time. Epilation of the eyebrows was noted at days 15-17 in 16% of the persons with second-degree acute radiation sickness, and in 67% and 100% of those with third- and fourth-degree acute radiation sickness, respectively. The epilation was partial and transient. Hair growth on the head was fully restored. All patients retained their eyelashes.

68. Corneal damage was manifested in an early reduction in corneal sensitivity coinciding with the first wave of erythema, although first-degree patients

did not show such an effect. At later times (days 35-55), superficial radiation-induced keratitis was observed in patients suffering from second-, third- and fourth-degree radiation sickness in 5%, 52% and 100% of the cases, respectively. Also noted were focal defects on the superficial epithelium of the cornea; these defects, which often merged, stained with fluorescein. The radiation keratitis regressed over a period of 1-1.5 months, leaving no opacification of the cornea.

69. Signs of disturbances in the haemodynamics of the retina were related to the dose and the degree of severity of radiation sickness. From a few days after irradiation, a reduction was observed in the level of diastolic pressure in the central retinal artery, followed later by signs of hypotonic angiopathy of the retina. Coinciding in time with the peak of the sickness, other injuries appeared, e.g., retinal oedema along the vessels and increased permeability of the retinal vessels (plasma discharge and haemorrhaging). The low diastolic pressure in the central retinal artery persisted over the entire acute phase.

70. In one severely ill patient (case 29, dose about 8.7 Gy) with fourth-degree acute radiation sickness, who survived the acute phase, the symptoms of angioretinopathy with haemorrhaging and plasma discharge recurred within 4.5 months, accompanied by a persistently low diastolic pressure in the central retinal artery (up to 5-10 mm Hg).

71. In the acute period, the treatment consisted in the topical application of ointments to the scaling surface of the eyelid skin and the instillation of 20% albucid, sopradex and vitamin solutions as eyedrops into the conjunctival cavity.

72. Within observation periods of up to one year, no obvious radiation-induced alterations of the lens were noted.

6. Treatment of radiation burns and other injuries

73. The treatment of radiation burns and other non-bone-marrow syndromes and their complications posed complex and multifaceted problems [J18]. From day 2 through day 8, 15 haemosorption sessions (purification using activated charcoal) were conducted for 13 patients suffering from the most severe skin lesions. Three patients who had been exposed to a total dose range of 2.0-4.6 Gy survived; they underwent haemosorption on a single occasion at days 5-8, i.e., considerably later than the time at which this might have affected the treatment of the bone marrow syndrome. This method of treatment did not change the outcome of the illness by modifying the haemocytopenia.

74. During the haemosorption process, and particularly towards the end of the session, many patients experienced a short-term improvement (lasting from a few hours to a single day), a reduction or disappearance of the pain in the extremities, and also a decrease of the oedema in their tissues. In this connection, contributory effects from the medication accompanying the procedure cannot be totally excluded.

75. A more widely used technique to combat the development of renal-hepatic insufficiency and fatal encephalopathic coma was plasmapheresis. Lesions induced by beta-irradiation over 30-40% and more of the body surface served as an indication for the application of this procedure. Plasmapheresis sessions were conducted for 17 patients from days 18-37. For a number of patients, daily sessions were conducted, up to six times.

76. The positive effect of repeated plasmapheresis was shown by a reduction of bilirubinemia and transaminasemia and a lowering of the nitrate level in patients suffering from renal-hepatic insufficiency caused by skin burns. On occasion, the plasmapheresis sessions were accompanied by reactions of minor severity such as chills and fever; there were no fatal complications. Another method used to treat toxicosis due to skin injuries was the injection of 1,000 ml of freshly-frozen plasma, accompanied by round-the-clock administration of heparin (1,000 active units/hour) with a liquid load (2-6 litres/day) and forced diuresis adequate to the intake volume. A precondition for this treatment was the presumption of disseminated intravascular clotting (DIC) syndrome (no typical anomalies in respect of coagulation were present) as a possible cause of encephalopathy and renal-hepatic syndrome. In its most strictly applied form, the heparin treatment method was used with two patients over a period of 7-15 days. The impression was that these patients survived longer than did patients whose condition was similar in terms of severity and extent of their burns. Their renal-hepatic insufficiency was less pronounced; however, a death due to encephalopathic coma was not averted.

77. The topical treatment of the burns required the involvement of a group of surgeons and nurses. A broad range of preparations and agents having an anti-inflammatory, bacteriostatic and regeneration-stimulating effect was used. Good results were achieved with lioxanol aerosol, an anti-burn ointment based on hydrocortisone with locally acting antibiotics, as well as BALIZ-2 solution and collagenous coatings. In each individual case the treatment varied in accordance with the stage of the lesions. Experience gained in the use of bactericidal fabric, both as a dressing material and for supplementary bedding, for patients with extensive burns deserves a particularly favourable comment in this connection [Z2].

78. Treatment of pain, as is typical of radiation injuries, was rather ineffective. At present, there are clearly no suitably effective local anaesthetics.

79. In patients suffering from severe radiation-induced inflammation of the oral mucosa, and enteritis, total parenteral nutrition had a positive effect; this was based on alvesin hydrolysate or an aminoacid mixture, aminone and a 40% glucose solution as the energy material. The treatment was carried out according to the principles and rules described by Dudrick et al. [D18]. This method was tested over a number of years with good results in patients receiving whole-body therapeutic gamma-irradiation at a dose level of 10 Gy for allogeneic bone marrow transplantation.

The danger, which has possibly not been fully evaluated, is the probability that certain severely injured, comatose patients may enter a state of hyperosmolarity. Data on plasma osmolarity that would appear to be necessary in a programme of total parenteral nutrition were not provided for all patients.

80. For the majority of patients suffering from first- and second-degree bone marrow syndrome, the period of clinical convalescence was completed by the third or fourth month. A longer period of treatment was required by persons suffering from severe radiation burns and the sequelae of third- and fourth-degree bone marrow syndrome. At the present time, the bulk of the patients have resumed work with the exclusion of any contact with radiation sources.

81. Over the period from the fourth month to one year after the accident, the specialized treatment centre was periodically visited by patients with skin lesions (dystrophic and ulcerated areas and also oedema of the subcutaneous tissues, mainly on the knees and feet). These patients are being treated with agents designed to improve local blood circulation and tissue trophism. Five patients with deep and extensive ulcers on their arms and other areas of the body underwent repeated plastic surgery, and a number of them will require more extended treatment.

82. Immunological examination data, acquired 0.5-1.5 years after the accident, have shown that in the peripheral blood of the patient groups with a history of acute radiation sickness of the second, third and fourth degrees a decline was observed in the number of T-lymphocytes with helper activity along with an increase in the number of T-lymphocytes with suppressor activity. This led to a considerable reduction in the normal ratio between these immunoregulatory lymphocyte sub-populations. At the same time, there was no reduction in the general lymphocyte level or in their T- and B-sub-populations. As an average for the groups, the level of class A, M and G immunoglobulins in the patients' blood serum corresponded to the physiological norm. Similar changes were not observed in the case of patients with a history of acute radiation sickness of the first degree. During this time they experienced no severe or life-threatening infections. In a number of cases an effort was made at immuno-corrective therapy using T- and B-activin.

83. Within these same patient groups, an estimate of the number of respiratory illnesses over the same period of time was conducted retrospectively. It was found that the incidence of illness in the group of 19 patients with a history of first-degree acute radiation sickness did not differ from the incidence of illness for the group of persons for whom no acute radiation sickness diagnosis had been established, and that it averaged 0.3 cases per person per year. During the same period, this indicator approached 1 for 22 patients who had experienced second-degree acute radiation sickness, and 3 for 8 persons with a history of third- to fourth-degree acute radiation sickness.

84. This comparison underlines the importance of the immune system in maintaining anti-infection

resistance in radiation convalescents and raises the question as to the usefulness of conducting supportive immunomodulating therapy courses, long after the incident, for persons who have undergone severe forms of radiation sickness.

85. The experience of the specialized treatment centres in Moscow and Kiev in the organization of medical care of persons exposed in this nuclear reactor accident has been described [N16]. For the survivors, a plan of scheduled follow-up observation is in effect, and decisions as how best to arrange their living and working conditions are being taken.

D. CONCLUSIONS

86. The analytical data presented in this Appendix and derived from clinical observations of the victims of the accident at the Chernobyl nuclear power plant are in agreement with the data in Annex G.

87. However, the fact that such a large group of 115 patients, who had all received uniform whole-body irradiation, was treated simultaneously for acute radiation sickness of varying degrees of severity, represents a unique event that makes it possible to clarify numerous aspects of early effects in man. A complicating factor was the presence of severe and extensive beta-radiation skin injuries in 58 patients which aggravated the course of the sickness in 19 of the 28 who died. Two more patients died during the first days as a result of severe combined injuries (trauma plus thermal burns plus irradiation).

88. The analysis provides a basis for describing the principal clinical syndrome, the bone marrow syndrome, with various degrees of severity in all 115 patients. In the case of some of them the bone marrow syndrome was combined with intestinal and oropharyngeal injuries and radiation damage to the skin, the forward segment of the eye (keratitis), and the lungs.

89. The treatment provided was in accordance with international practice and proved highly effective for the patient group exposed to doses of 2-4 Gy and for two thirds of the patients who received doses of 4-6 Gy. In the group of patients receiving 6-16 Gy, two patients who received doses of 8-9 Gy survived past 60 days.

90. The average bone marrow dose and the prognosis regarding the further course of the illness were determined on the basis of biological criteria. During the early period, most information was obtained from the karyological analyses, the lymphocyte counts and the primary reaction periods; later, from the granulocyte counts. The remaining indications were of an auxiliary nature. In three cases, the dose value coincided with the electron spin resonance study of dental enamel after death.

91. There is a need for further analysis of the time course of the early effects for a more accurate understanding of the nature of lung and neurological injuries, and for more detailed data on the relevance of biological dose indicators and the reasons for disparities between them. It is hoped that these data will be of use in the preparedness to respond in the event of an accident of a similar type in the provision of medical treatment.

Table A.1

Range of thyroid doses (Sv)	Number of persons
0 - 1.2	173
1.2 - 3.7	18
3.7 - 6.1	4
6.1 - 8.6	4
8.6-11.0	2
11.0-13.4	2
13.4-15.9	0
15.9-18.3	2
18.3-20.8	0
20.8-23.2	1

Table A.2

Case number	Thyroid dose a/ (Gy)	Lung dose a/ (Gy)	Whole-body dose (Sv)	
			Internal	External
24	30	2.5	2.0	1.7
25	6	2.0	1.0	4.7
17	1	0.4	0.2	10.0
3	0.3	0.3	0.2	12.0
4	1.2	0.4	0.1	11.0
26	0.5	0.3	0.1	12.0

a/ Doses accumulated until time of death.

Table A.3

Degree of severity	Number of patients	Bone marrow dose range (Gy)	Number of deaths	Time to death (days)
I	31	0.8-2.1	-	-
II	43	2.2-4.1	1	96
III	21	4.2-6.4	7	16,18,21,23,34,48,48
IV	20	6.1- 16	20	10,14,14,15,15,17,17, 18,18,18,20,21,23,24, 24,25,30,32,86,91

a/ In addition to the patients who died of acute radiation sickness, one person died at the plant site and another within the first 12 hours following the accident, as a result of thermal burns, at the in-patient clinic in Pripyat where he had been given first aid.

Table A.4

Assessment of irreversible myelodepression
according to diagnostic scores
in cases of acute radiation sickness

Sign		Diagnostic score a/
Time to the onset of vomiting	(hours)	
	0.00- 0.4	+ 8
	0.41- 0.8	+ 4
	0.81- 1.2	+ 2
	1.21- 1.6	- 2
	1.61- 2.0	- 6
	> 2.01	-10
Lymphocyte count on the second day	(10 ⁹ /l)	
	0.00- 0.2	+ 6
	0.21- 0.4	+ 2
	0.41- 0.6	- 2
	0.61- 0.8	- 8
	> 0.81	-15
Lymphocyte count on the third day	(10 ⁹ /l)	
	0.00- 0.1	+ 8
	0.11- 0.2	+ 2
	0.21- 0.3	- 2
	0.31- 0.4	- 9
	> 0.41	-10
Lymphocyte count on the fourth day	(10 ⁹ /l)	
	0.00- 0.1	+ 4
	0.11- 0.2	+ 2
	0.21- 0.3	0
	0.31- 0.7	- 2
	0.71- 0.8	- 3
	0.81- 0.9	- 8
Lymphocyte count from day 4 to day 7	(10 ⁹ /l)	
	0.00- 0.1	+ 5
	0.11- 0.2	+ 2
	0.21- 0.3	- 1
	0.31- 0.4	- 5
	0.41- 0.5	-13
	> 0.51	-15
Average reticulocyte count from day 3 to day 5	(10 ⁹ /l)	
	0.0 - 8.0	+ 2
	0.1 -10.0	0
	10.1 -14.0	- 4
	14.1 -18.0	- 6
	18.1 -20.0	-10
Minimum neutrophil count for day 6 to day 7	(10 ⁹ /l)	
	0.00- 0.3	+12
	0.31- 0.6	+ 5
	0.61- 0.9	0
	0.91- 1.2	- 3
	1.21- 2.4	- 6
	2.41- 3.0	- 8

a/ The diagnostic signs are used to determine the diagnostic scores, which are then added together. A sum of +10 is the basis for a prognosis of irreversible myelodepression; a sum of -10 for a prognosis of no irreversible myelodepression. If after the diagnostic coefficients of all the available signs have been added no positive value has been reached, the answer is indeterminate (the available information is insufficient for a differential diagnosis, with an error probability of not more than $\pm 10\%$).

T a b l e A.5

Survival or cause of death of patients receiving bone marrow transplantations and of patients in control group

Dose range (Gy)	Bone marrow transplant patients				Control patients		
	Number of patients	Deaths a/	Deaths b/	Number of survivors	Number of patients	Deaths a/	Number of survivors
< 6.5	4	0	3	1	5	0	5
6.5-9	3	2	c/	0	4	3	1
> 9	6	5	1	0	5	5	0
Total	13	7	4	2	14	8	6

a/ Skin and intestinal injuries.

b/ Bone marrow rejection (graft-versus-host disease) plus infection.

c/ Positive graft-versus-host disease post-mortem histology.

T a b l e A.6

Distribution of cases of radiation burns of different degree in the presence of acute bone marrow syndrome

Degree of severity of bone marrow syndrome	Total number of patients	Number of patients with radiation burns to various percentages of the body surface		
		0-10%	10-50%	50-100%
I	31	2	1	0
II	43	2	9	1
III	21	3	15	3
IV	20	1	10	9
Total	115		56	

T a b l e A.7
Patient identification, estimated dose, cause and day of death

Degree of severity of ARS	Case number	Bone marrow dose (Gy)	Treatment a/	Day of death b/	Cause of death
II	33	4.1		96	Infection, renal-hepatic insufficiency and skin injuries
III	5	4.4	BMT	34	Infection, post-transplantation immunosuppression
	7	4.7		18	Skin injuries, post-transfusion shock
	24	3.7		23	Thermal and radiation burns
	25	5.7		16	Thermal and radiation burns
	28	6.4	BMT	48	Infection, graft-versus-host disease
	30	5.5	BMT	21	Bleeding from mechanical injury during catheterization
	34	5.8		48	Respiratory insufficiency, cerebral oedema
IV	1	6.6	BMT	25	Toxicity, respiratory insufficiency
	2	9.2	BMT	15	Skin and lung injuries
	3	12	BMT	17	Skin and intestinal injuries
	4	11.8	BMT	18	Skin and intestinal injuries
	6	7.5	BMT	86	Infection, graft-versus-host disease
	8	8.3	LCT	30	Toxicity, respiratory insufficiency
	9	9.7		23	Skin and lung injuries
	10	11.1	LCT	14	Skin and intestinal injuries
	12	9.3		24	Lung injuries
	14	10.9	LCT	18	Skin and intestinal injuries
	15	>10	LCT	14	Skin and intestinal injuries
	16	10.1	BMT	91	Infection, graft-versus-host disease
	17	10	BMT	18	Skin and intestinal injuries
	20	12.4	LCT	17	Skin and intestinal injuries
	23	13.7	LCT	15	Skin and intestinal injuries
	26	12.5		20	Skin and intestinal injuries
	27	8.3	BMT	24	Lung injuries
	31	6.7		32	Respiratory insufficiency, cerebral oedema
	62	6.1		21	Radiation burns (skin injuries)
	2097	10.2		10	Skin and intestinal injuries
	(Kiev)				

a/ BMT = bone marrow transplantation; LCT = liver cell transplantation.

T a b l e A.8
Type of eye changes and per cent incidence
in the victims of the accident

Nature of the changes	Degree of acute radiation sickness			
	I	II	III	IV
First wave of erythema	6.1	39.5	100	100
Second wave of erythema		20.9	80.9	100
Reduction in cornea sensitivity		18.6	100	100
Epilation of the eyebrows		16.3	66.7	100
Keratitis		4.6	52.4	100
Fundus				
Dilation of blood vessels		32.6	74.4	100
Decreased diastolic pressure of the central retinal artery		48.8	95.2	100
Retinal oedema		4.6	—	80
Haemorrhaging		13.9	23.8	80
Plasmorrhaging		4.6	23.8	80

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