

Extensive swelling of the vaccinated limb (ELS) and administration of the Pfizer/BioNTech vaccine (Comirnaty®).

Introduction

The Pfizer/BioNTech vaccine (Comirnaty®) is a COVID-19-mRNA-vaccine (nucleoside modified). It is indicated for *active immunisation to prevent COVID-19 caused by SARS-CoV-2 virus, in individuals 16 years of age and older*. [1] The nucleoside-modified messenger RNA in Comirnaty® is formulated in lipid nanoparticles, which enable delivery of the nonreplicating RNA into host cells to direct transient expression of the SARS-CoV-2 S antigen. The mRNA codes for membrane-anchored, full-length S with two point mutations within the central helix. [1] Comirnaty® has been registered in Europe since December 21st, 2021.

After vaccination it is possible to develop swelling and/or erythema of the injection site. In some cases this swelling and/or erythema spreads beyond the shoulder, elbow, knee or hip. Sometimes this covers more than half of the vaccinated limb. This reaction is called extensive swelling of the vaccinated limb (ELS) [2]. ELS can also include other inflammation symptoms, like warmth or pain of the injection site [3-5]. Pruritis and a lesser ability to move the swollen arm or leg are common symptoms as well [3]. Sometimes a mild fever is observed in the same period as the ELS [3]. In most cases, the reaction develops between 24 to 48 hours after vaccination. Treatment is not necessary, ELS mostly disappears spontaneously within 4 to 5 days, without remaining symptoms [2].

ELS is a known side effect of vaccines including, but not limited to, the seasonal Influenza Vaccines and the DKTP (Diphtheria, Pertussis, Tetanus and Polio) vaccination around the age of four years in the National vaccination program of the Netherlands [3, 6-7].

Reports

In the period from January 6th, 2021 until February 16th, 2021 the Netherlands Pharmacovigilance Centre Lareb received 80 reports on ELS associated with administration of Comirnaty®. Reports with the MedDRA Preferred Term (PT) “Extensive swelling of the vaccinated limb” were included in this analysis. It concerned a female in 72 reports and 8 reports concerned a male. The average age was 42.5 years (ranging from 19 years to 94 years).

The latency time was not specifically reported 14 times as “within one day” and once “within two days”. These reports were not included in calculating the average latency time. The average latency time was approximately 38 hours (so between 1-2 days). This varied from immediately after the vaccination up to 12 days.

17 times (21%) the patient had recovered at the time of reporting. The average duration of ELS in these cases was 2.6 days (varying from 1 hour to 7 days). Two cases were not included in calculating the average duration, because they were reported as “within 1 day” and “within 3 days”, and the exact timing is therefore not certain. 29 times (36%) the patient was recovering at the time of reporting. The average time between vaccination and reporting in these cases was 2.7 days (varying from 0 to 13 days). 33 times (41%) the patient was not recovered at the time of reporting. The average time between vaccination and reporting in these cases was 1.5 days (varying from 0 to 5 days) and in two cases the time between vaccination and reporting is unknown. The outcome of one report is unknown.

The diagnosis ELS was either based on the descriptions used on the reporting form following the selection of “injection site swelling”, “injection site erythema” or both. In case of one of these descriptions is selected, an additional question about the extent of the swelling is posed. An extensive swelling or erythema is defined by either an extension beyond the elbow and/or

shoulder or an extension that is visible at both the inside and the outside of the limb. In case the reporter answers this question with “yes”, the MedDRA PT “Extensive swelling of the vaccinated limb” is automatically coded next to the existing MedDRA PT “injection site swelling (29%)”, PT “injection site erythema (29%)” or both (42%).

The MedDRA PT’s that were most reported next to the MedDRA PT “Extensive swelling of the vaccinated limb” were other injection site reactions like injection site pain (65%), injection site warmth (63%) and injection site pruritus (26%). Other PT’s that were reported often included known side effects for Comirnaty® like myalgia (51%), headache (48%), malaise (45%), fatigue (43%), chills (33%), generalized joint pain (30%) and pyrexia/fever (28%).

Addendum 1 contains a table with an overview of all reported cases.

Other sources of information

SmPC

The SmPC of Comirnaty® does not mention ELS as an adverse reaction [1].

Literature

ELS is not explicitly mentioned in the clinical study as a possible side effect [8]. The local reactions that occurred during this clinical study are described as follows:

“The reactogenicity subset included 8,183 participants. Overall, BNT162b2 recipients reported more local reactions than placebo recipients. Among BNT162b2 recipients, mild-to-moderate pain at the injection site within 7 days after an injection was the most commonly reported local reaction, with less than 1% of participants across all age groups reporting severe pain. Pain was reported less frequently among participants older than 55 years of age (71% reported pain after the first dose; 66% after the second dose) than among younger participants (83% after the first dose; 78% after the second dose). A noticeably lower percentage of participants reported injection-site redness or swelling. The proportion of participants reporting local reactions did not increase after the second dose.”

Databases

Table 1. Reports of ELS associated with the administration of Comirnaty® in the Lareb and WHO database [9, 10].

Database	Drug	ADR	Number of reports
Lareb	Comirnaty®	ELS	80*
WHO	Comirnaty®	ELS	115

* See Addendum 1 for all cases from Lareb.

Prescription data

On the 14th of February, approximately 600,000 and 150,000 people in the Netherlands had received their first and second dose respectively of Comirnaty® [11].

Mechanism

The mechanism of this local reaction is not well known. In the literature it is suggested that an acute allergic reaction (type I, IgE-mediated) as a cause of ELS is not probable, based on the time of onset of ELS after vaccination [3]. IgE-mediated allergic reactions usually occur within one hour after vaccination [12]. A bacterial infection is also unlikely, since this normally occurs with high fever and malaise. Both symptoms do not occur with a typical ELS [13]. Despite the fact that the mechanism of ELS after vaccination is unknown, it is generally assumed that it concerns a severe inflammatory reaction [3].

Discussion and conclusion

The Netherlands Pharmacovigilance Centre Lareb received 80 reports of ELS after administration of Comirnaty®. Comirnaty® is used for active immunisation to prevent COVID-19 caused by SARS-CoV-2 virus in individuals 16 years of age and older. ELS is a known side effect of some vaccines, including the seasonal influenza vaccine and the DKTP vaccination, and is now also observed after administration of Comirnaty®. ELS is not labelled in the SmPC of Comirnaty®. The number of reports of ELS are relatively high compared to the number of reports worldwide (table 1). As mentioned under “Reports” the PT “Extensive swelling of the vaccinated limb” is coded when the reporter confirms in an additional question that the injection site swelling or injection site erythema extends the elbow and/or shoulder or if it is visible on both the inside and outside of the arm. This additional question about the extent of the swelling in the reporting form may have led to this relatively high amount of received reports in the Netherlands.

The observed cases of ELS induced by Comirnaty® in the Netherlands have an average latency of around 38 hours, so between 1 and 2 days. In 21% of the cases, the patient was recovered at time of reporting with an average duration of 2.6 days. 36% was recovering and 41% was not recovered at time of reporting. The time between vaccination and reporting varies from 0 to 13 days in these cases. The outcome of one report is unknown.

Despite of the fact that ELS mostly disappears spontaneously and without remaining symptoms, this local reaction can appear severe and may cause concern among patients. In addition, ELS can lead to unnecessary treatment with anti-allergy drugs or antibiotics when this side effect is not correctly labeled. It is therefore necessary that people are aware of the possibility of ELS after administration of Comirnaty®.

Reference list

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- [12] Pichler WJ, Hausmann O. Classification of Drug Hypersensitivity into Allergic, p-i, and Pseudo-Allergic Forms. International archives of allergy and immunology. 2016-01-01;171(3-4):166-179
- [13] Lapphra K, Scheifele D. Vaccination site reaction or bacterial cellulitis? Paediatrics & child health. 2009-04-01;14(4):245

This signal has been raised on March 17, 2021. It is possible that in the meantime other information became available. For the latest information, including the official SmPC's, please refer to website of the MEB www.cbg-meb.nl

Addendum 1: Table with all received cases until February 16th, 2021.

Case	ID, sex, age, primary source, reporting date	Drug, vaccination date	Reported PTs	Latency after start of ELS	Outcome of ELS Duration	Concomitant medication
1	NL-LRB-00428979 F, 50-60 Patient 08-01-2021	Pfizer vaccine 07-01-2021	Headache Injection site swelling Injection site pain Extensive swelling of vaccinated limb	Within 1 day	Not recovered 1 day after onset.	-
2	NL-LRB-00429349 F, 40-50 Patient 11-01-2021	Pfizer vaccine 08-01-2021	Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	1 day	Not recovered 2 days after onset.	-
3	NL-LRB-00429382 F, 30-40 Patient 11-01-2021	Pfizer vaccine 06-01-2021	Dark circles under eyes Chills Headache Myalgia Malaise Fatigue Injection site erythema Injection site warmth Injection site pain Extensive swelling of vaccinated limb Asthenia Dizziness	1 day	Recovered 1 hours	-
4	NL-LRB-00429419 F, 40-50 Other health professional 11-01-2021	Pfizer vaccine 08-01-2021	Injection site swelling Myalgia Generalized joint pain Injection site erythema Extensive swelling of vaccinated limb Injection site pain Injection site pruritus	10 hours	Recovering 3 days after onset.	-
5	NL-LRB-00429633 F, 40-50 Patient 12-01-2021	Pfizer vaccine 08-01-2021	Headache Myalgia Generalized joint pain Malaise Fatigue Injection site warmth Injection site pain Injection site induration Injection site swelling Extensive swelling of vaccinated limb Pyrexia	13 hours	Recovering 4 days after onset	-
6	NL-LRB-00429638 F, 30-40 Patient 12-01-2021	Pfizer vaccine 09-01-2021	Erythema Injection site pain Injection site swelling Extensive swelling of vaccinated limb Pyrexia	1 days	Recovered 1 days	-

7	NL-LRB-00429967 F, 60-70 Patient 14-01-2021	Pfizer vaccine 12-02-2021	Erythema Chills Myalgia Malaise Injection site erythema Injection site warmth Injection site swelling Extensive swelling of vaccinated limb	Immediately*	Recovered 1 day	-
8	NL-LRB-00429977 F, 60-70 Patient 14-01-2021	Pfizer vaccine 08-01-2021	Fatigue Malaise Injection site erythema Injection site warmth Injection site pain Injection site pruritus Extensive swelling of vaccinated limb Pyrexia	1 days	Recovered 5 days	-
9	NL-LRB-00430475 F, 40-50 Patient 15-01-2021	Pfizer vaccine 10-01-2021	Chills Headache Myalgia Generalized joint pain Malaise Fatigue Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	3 hours	Not recovered 5 days after onset	Levothyroxine Omeprazol Hydroxocobalamine Colecalciferol Metformine Magnesiumcitraat
10	NL-LRB-00430658 F, 20-30 16-01-2021	Pfizer vaccine 15-01-2021	Myalgia Malaise Fatigue Injection site warmth Injection site swelling Injection site pain Extensive swelling of vaccinated limb	Immediately*	Not recovered 1 day after onset	
11	NL-LRB-00430673 F, 20-30 Patient 16-01-2021	Pfizer vaccine 12-01-2021	Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	1 day	Recovering 3 days after onset	-
12	NL-LRB-00430788 F, 30-40 Patient 16-01-2021	Pfizer vaccine 16-01-2021	Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	4 hours	Not recovered on day of onset	-
13	NL-LRB-00430863 F, 50-60 Patient 17-01-2021	Pfizer vaccine 16-01-2021	Chills Headache Myalgia Generalized joint pain Malaise	1 day	Not recovered on day of onset	-

			Injection site erythema Injection site warmth Injection site pain Extensive swelling of vaccinated limb Body temperature increased			
14	NL-LRB-00430965 F, 20-30 Patient 17-01-2021	Pfizer vaccine 16-01-2021	Myalgia Injection site erythema Injection site pain Injection site warmth Injection site swelling Extensive swelling of vaccinated limb	30 minutes	Not recovered 1 day after onset	-
15	NL-LRB-00430990 F, 40-50 Patient 17-01-2021	Pfizer vaccine 15-01-2021	Extensive swelling of vaccinated limb Chills Headache Nausea Myalgia Malaise Fatigue Injection site erythema Injection site warmth Injection site pain Injection site swelling	1 hours	Not recovered 2 days after onset	Levothyroxine Ethinylestradiol
16	NL-LRB-00431007 F, 30-40 Patient 18-01-2021	Pfizer vaccine 17-01-2021	Headache Myalgia Fatigue Injection site pruritus Injection site warmth Injection site erythema Injection site swelling Extensive swelling of vaccinated limb	10 minutes	Recovering 1 day after onset	Ethinylestradiol/ Levonorgestrel Multivitaminen + mineralen tablet
17	NL-LRB-00430421 F, 60-70 Patient 15-01-2021	Pfizer vaccine 12-01-2021	Myalgia Fatigue Headache Injection site erythema Injection site pruritus Injection site swelling Extensive swelling of vaccinated limb Pyrexia	1 day	Recovering 2 days after onset	-
18	NL-LRB-00431048 F, 50-60 Patient 18-01-2021	Pfizer vaccine 16-01-2021	Dry mouth Fatigue Injection site erythema Injection site warmth Injection site pruritus Injection site swelling Injection site papule Extensive swelling of vaccinated limb Malaise	1 day	Not recovered 1 day after onset	Sertraline

19	NL-LRB-00431065 F, 20-30 Patient 18-01-2021	Pfizer vaccine 15-01-2021	Eye pruritus Headache Nausea Myalgia Generalized joint pain Malaise Injection site warmth Injection site erythema Injection site pain Extensive swelling of vaccinated limb	Within 1 day	Not recovered 3 days after onset	-
20	NL-LRB-00431097 F, 50-60 Patient 18-01-2021	Pfizer vaccine 13-01-2021	Headache Myalgia Generalized joint pain Malaise Fatigue Injection site erythema Injection site warmth Injection site pain Injection site pruritus Injection site swelling Extensive swelling of vaccinated limb	Within 1 day	Not recovered 5 days after onset	-
21	NL-LRB-00431470 F, 50-60 Patient 19-01-2021	Pfizer vaccine 07-01-2021	Headache Myalgia Injection site erythema Injection site pain Injection site swelling Extensive swelling of vaccinated limb Pyrexia	1 day	Recovered 7 days	-
22	NL-LRB-00431481 F, 50-60 Patient 19-01-2021	Pfizer vaccine 15-01-2021	Chills Headache Myalgia Malaise Fatigue Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	16 hours	Recovering 4 days after onset	-
23	NL-LRB-00431569 F, 20-30 Patient 19-01-2021	Pfizer vaccine 19-01-2021	Chills Headache Nausea Myalgia Generalized joint pain Malaise Fatigue Injection site hematoma Injection site erythema Injection site warmth Extensive swelling of vaccinated limb	Within 1 day	Recovering on the day of onset	-

24	NL-LRB-00431605 F, 20-30 Physician 19-01-2021	Pfizer vaccine 19-01-2021	Myalgia Malaise Injection site erythema Injection site warmth Injection site pruritus Extensive swelling of vaccinated limb	10 minutes	Recovering on the day of onset	-
25	NL-LRB-00431632 F, 50-60 Patient 19-01-2021	Pfizer vaccine 17-01-2021	Headache Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	2 days	Not recovered on the day of onset	-
26	NL-LRB-0043731 F, 10-20 Patient 20-01-2021	Pfizer vaccine 18-01-2021	Rash erythematous Headache Nausea Myalgia Generalized joint pain Malaise Fatigue Injection site erythema Injection site warmth Injection site pain Injection site pruritus Injection site haematoma Injection site swelling Injection site rash Extensive swelling of vaccinated limb Pyrexia	4 hours	Not recovered 2 days after onset	Magnesiumcitraat
27	NL-LRB-00431853 F, 20-30 Patient 20-01-2021	Pfizer vaccine 06-01-2021	Lymphadenopathy Headache Myalgia Generalized joint pain Injection site erythema Injection site warmth Injection site pain Injection site haematoma Injection site swelling Injection site induration Extensive swelling of vaccinated limb Pyrexia	1 day	Recovering 13 days after onset	Desloratadine
28	NL-LRB-00431962 F, 50-60 Patient 20-01-2021	Pfizer vaccine 17-01-2021	Injection site burning Chills Injection site warmth Injection site pruritus Injection site swelling Erythematous rash Extensive swelling of vaccinated limb Pruritus	10 hours	Recovering 3 days after onset	Clonidine Oxycodone Pantoprazole

29	NL-LRB-00429430 F, 20-30 Patient 11-01-2021	Pfizer vaccine 08-01-2021	Erythema Sensation of warmth Taste metallic Malaise Fatigue Injection site erythema Injection site warmth Extensive swelling of vaccinated limb	5 minutes	Recovered 2 hours	Clemastine
30	NL-LRB-00432075 F, 30-40 Patient 20-01-2021	Pfizer vaccine 17-01-2021	Injection site pruritus Nausea Injection site erythema Injection site swelling Extensive swelling of vaccinated limb	6 hours	Recovering 3 days after onset	Ethinylestradiol/ levonorgestrel
31	NL-LRB-00432126 M, 30-40 Patient 20-01-2021	Pfizer vaccine 19-01-2021	Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	1 day	Not recovered on day of onset	-
32	NL-LRB-00432325 F, 50-60 Patient 21-01-2021	Pfizer vaccine 19-01-2021	Chest pain Headache Nausea Myalgia Generalized joint pain Fatigue Injection site pain Injection site erythema Injection site swelling Extensive swelling of vaccinated limb Pyrexia Dyspnea Lung pain Throat pain	Within 1 day	Not recovered 2 days after onset	-
33	NL-LRB-00432384 F, 50-60 Patient 21-01-2021	Pfizer vaccine 18-01-2021	Chills Headache Nausea Myalgia Fatigue Generalized joint pain Malaise Injection site swelling Extensive swelling of vaccinated limb Pyrexia	1 hours	Recovered Within 1 day	-
34	NL-LRB-00432949 F, 20-30 Patient 23-01-2021	Pfizer vaccine 21-01-2021	Headache Nausea Fatigue Injection site erythema Injection site pain Extensive swelling of vaccinated limb	40 hours	Not recovered on day of onset	Multivitamins

35	NL-LRB-00432976 F, 60-70 Patient 23-01-2021	Pfizer vaccine 17-01-2021	Chills Headache Nausea Myalgia Generalized joint pain Malaise Fatigue Injection site erythema Injection site warmth Injection site pain Injection site pruritus Injection site swelling Extensive swelling of vaccinated limb Fever Pruritus	8 hours	Recovering 5 days after onset	Ezetimib Amlodipine Enalapril Rosuvastatine Pantoprazole
36	NL-LRB-00433013 F, 40-50 Patient 23-01-2021	Pfizer vaccine 20-01-2021	Lymphadenopathy Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	1 day	Not recovered 2 days after onset	-
37	NL-LRB-00433154 F, 40-50 Patient 23-01-2021	Pfizer vaccine 18-01-2021	Chills Headache Fatigue Injection site erythema Injection site pain Injection site pruritus Injection site haematoma Extensive swelling of vaccinated limb Body temperature increased	1 day	Recovered Within 3 days	Esomeprazole Fluticasone nose spray Rosuvastatine Beclometason/ formeterol aerosol Magnesiumcitraat Colecalciferol
38	NL-LRB-00433257 F, 50-60 Patient 24-01-2021	Pfizer vaccine 07-01-2021	Injection site erythema Injection site swelling Extensive swelling of vaccinated limb	1 day	Recovered 7 days	-
39	NL-LRB-00433654 F, 50-60 Patient 25-01-2021	Pfizer vaccine 22-01-2021	Fever Malaise Fatigue Injection site swelling Injection site pain Injection site erythema Extensive swelling of vaccinated limb	3 days	Not recovered on day of onset	Levothyroxine Omeprazol
40	NL-LRB-00433656 F, 50-60 Patient 25-01-2021	Pfizer vaccine 19-01-2021	Headache Nausea Malaise Injection site erythema Injection site warmth Injection site pain Extensive swelling of vaccinated limb	1 day	Not recovered 5 days after onset	-

			Pyrexia			
41	NL-LRB-00433665 M, 50-60 Patient 25-01-2021	Pfizer vaccine 09-01-2021	Generalized joint pain Malaise Fatigue Chills Myalgia Injection site pain Injection site warmth Injection site erythema Extensive swelling of vaccinated limb	1 hours	Recovering 3 days	-
42	NL-LRB-00433819 F, 50-60 Patient 25-01-2021	Pfizer vaccine 20-01-2021	Headache Malaise Injection site erythema Injection site pain Injection site swelling Extensive swelling of vaccinated limb	3 days	Recovering 2 days after onset	Levothyroxine Colecalciferol
43	NL-LRB-00433875 F, 30-40 Patient 26-01-2021	Pfizer vaccine 16-01-2021	Injection site erythema Injection site warmth Injection site pruritus Injection site induration Extensive swelling of vaccinated limb	8 days	Not recovered 2 days after onset	Tioguanine Levothyroxine Amitriptyline
44	NL-LRB-00434458 F, 50-60 Patient 27-01-2021	Pfizer vaccine 15-01-2021	Injection site erythema Injection site haematoma Extensive swelling of vaccinated limb	12 days	Not recovered 1 day after onset	Ramipril Colecalciferol
45	NL-LRB-00434479 F, 20-30 Patient 27-01-2021	Pfizer vaccine 18-01-2021	Erythema Injection site erythema Injection site pruritus Extensive swelling of vaccinated limb Pruritus	Within 1 day	Recovered 1 day	-
46	NL-LRB-00434538 F, 60-70 Patient 27-01-2021	Pfizer vaccine 16-01-2021	Injection site erythema Injection site warmth Injection site induration Extensive swelling of vaccinated limb	8 days	Recovering 3 days after onset	Magnesium Red yeast rice Colecalciferol Japanese q10 Non-specified blood pressure medication
47	NL-LRB-00434636 F, 40-50 Patient 27-01-2021	Pfizer vaccine 26-01-2021	Injection site warmth Myalgia Injection site swelling Injection site warmth Injection site pain Extensive swelling of vaccinated limb	Immediately*	Recovering 1 day after onset	-
48	NL-LRB-00434701 F, 40-50 Patient 27-01-2021	Pfizer vaccine 23-01-2021	Myalgia Malaise Headache Injection site erythema Injection site warmth Injection site pain	Within 1 day	Recovering 4 days after onset	-

			Injection site pruritus Injection site swelling Extensive swelling of vaccinated limb			
49	NL-LRB-00434712 F, 40-50 Patient 27-01-2021	Pfizer vaccine 16-01-2021	Injection site erythema Injection site warmth Injection site pruritus Injection site induration Extensive swelling of vaccinated limb	10 days	Not recovered 1 day after onset	-
50	NL-LRB-00434798 F, 40-50 Patient 28-01-2021	Pfizer vaccine 26-01-2021	Chills Headache Myalgia Fatigue Injection site swelling Extensive swelling of vaccinated limb Pyrexia	Unknown (within 2 days)	Not recovered after unknown duration	Levothyroxine Budesonide Fluticasone nose spray
51	NL-LRB-00434977 F, 50-60 Patient 28-01-2021	Pfizer vaccine 18-01-2021	Chills Headache Nausea Myalgia Generalized joint pain Malaise Fatigue Injection site erythema Injection site warmth Injection site swelling Extensive swelling of vaccinated limb Hyperpyrexia	Within 1 day	Recovered 4 days	-
52	NL-LRB-00435023 F, 40-50 Patient 28-01-2021	Pfizer vaccine 26-01-2021	Limb discomfort Injection site swelling Extensive swelling of vaccinated limb Injection site discomfort	1 day	Recovering 1 day after onset	-
53	NL-LRB-00435046 F, 40-50 Physician 28-01-2021	Pfizer vaccine 26-01-2021	Lymphadenopathy Injection site pain Injection site swelling Extensive swelling of vaccinated limb	2 days	Not recovered on day of onset	-
54	NL-LRB-00435049 F, 10-20 Patient 28-01-2021	Pfizer vaccine 23-01-2021	Myalgia Headache Injection site swelling Injection site pain Extensive swelling of vaccinated limb	2 days	Recovered 3 days	-
55	NL-LRB-00435073 F, 30-40 Patient 28-01-2021	Pfizer vaccine 22-01-2021	Lymphadenopathy Throat pain Myalgia Generalized joint pain Injection site erythema Injection site warmth Injection site pain Injection site swelling	1 days	Recovered 1 day	-

			Extensive swelling of vaccinated limb Pyrexia			
56	NL-LRB-00435133 M, 30-40 Patient 28-01-2021	Pfizer vaccine 25-01-2021	Chills Myalgia Generalized joint pain Malaise Injection site erythema Injection site pain Injection site swelling Extensive swelling of vaccinated limb Body temperature increased	Within 1 day	Not recovered 3 days after onset	Valaciclovir
57	NL-LRB-00435271 F, 50-60 Patient 29-01-2021	Pfizer vaccine 28-01-2021	Breath shortness Fatigue Injection site pain Injection site swelling Extensive swelling of vaccinated limb	Within 1 day	Not recovered 1 day after onset	Methylprednisolone
58	NL-LRB-00435275 F, 20-30 Patient 29-01-2021	Pfizer vaccine 28-01-2021	Chills Headache Nausea Generalized joint pain Malaise Fatigue Injection site pain Injection site haematoma Injection site swelling Extensive swelling of vaccinated limb	1 hours	Recovering 1 day after onset	
59	NL-LRB-00435491 F, 40-50 Patient 29-01-2021	Pfizer vaccine 27-01-2021	Feeling cold Headache Myalgia Malaise Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	10 hours	Recovering 1 day after onset	Paracetamol Ibuprofen
60	NL-LRB-00435511 F, 20-30 Patient 29-01-2021	Pfizer vaccine 19-01-2021	Injection site erythema Injection site warmth Injection site pruritus Injection site induration Extensive swelling of vaccinated limb	9 days	Not recovered 1 day after onset	-
61	NL-LRB-00435516 F, 20-30 Patient 29-01-2021	Pfizer vaccine 28-01-2021	Headache Myalgia Malaise Generalized joint pain Injection site erythema Extensive swelling of vaccinated limb Pyrexia	1 day	Not recovered on the day of onset	-
62	NL-LRB-00435625	Pfizer vaccine	Headache	1 days	Recovered	-

	F, 50-60 Patient 29-01-2021	25-01-2021	Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb Pyrexia		2 days	
63	NL-LRB-00435708 F, 40-50 Patient 30-01-2021	Pfizer vaccine 27-01-2021	Chills Injection site erythema Injection site haematoma Injection site pain Injection site swelling Extensive swelling of vaccinated limb	Within 1 day	Recovering 3 days after onset	-
64	NL-LRB-00435852 F, 20-30 Patient 30-01-2021	Pfizer vaccine 29-01-2021	Chills Generalized joint pain Myalgia Nausea Headache Injection site swelling Injection site pain Extensive swelling of vaccinated limb Pyrexia	Within 1 day	Recovering 1 day after onset	-
65	NL-LRB-00435883 M, 70 years and older Patient 30-01-2021	Pfizer vaccine 26-01-2021	Myalgia Generalized joint pain Malaise Fatigue Injection site erythema Injection site haematoma Injection site swelling Injection site pain Extensive swelling of vaccinated limb	1 day	Recovering 3 days after onset	-
66	NL-LRB-00435999 F, 40-50 Patient 30-01-2021	Pfizer vaccine 27-01-2021	Chills Malaise Fatigue Headache Myalgia Injection site erythema Injection site warmth Extensive swelling of vaccinated limb Pyrexia	1 day	Not recovered 2 days after onset	Losartan
67	NL-LRB-00436185 F, 50-60 Patient 31-01-2021	Pfizer vaccine 28-01-2021	Chills Headache Nausea Myalgia Malaise Fatigue Injection site erythema Injection site warmth Injection site pain Extensive swelling of vaccinated limb	1 days	Recovered 1 day	Levothyroxine

			Fever			
68	NL-LRB-00436239 F, 30-40 Patient 31-01-2021	Pfizer vaccine 30-01-2021	Feeling hot Injection site haematoma Injection site erythema Injection site pruritus Extensive swelling of vaccinated limb Facial swelling Pruritus facial Erythema facial Injection site discomfort Facial rash	1 day	Unknown	Levothyroxine
69	NL-LRB-00436280 M, 20-30 Patient 31-01-2021	Pfizer vaccine Unknown	Injected limb mobility decreased Headache Generalized joint pain Injection site erythema Injection site warmth Injection site pain Injection site haematoma Injection site swelling Extensive swelling of vaccinated limb	2 days	Not recovered after unknown time	-
70	NL-LRB-00436312 F, 30-40 Patient 31-01-2021	Pfizer vaccine 17-01-2021	Injection site erythema Injection site warmth Injection site pruritus Injection site induration Extensive swelling of vaccinated limb	8 days	Recovering 6 days after onset	-
71	NL-LRB-00436381 F, 40-50 Patient 31-01-2021	Pfizer vaccine 30-01-2021	Dyspnoea Chills Headache Nausea Myalgia Generalized joint pain Malaise Fatigue Injection site erythema Injection site warmth Injection site pain Injection site pruritus Extensive swelling of vaccinated limb Epistaxis	1 day	Recovering on day of onset	-
72	NL-LRB-00436483 F, 20-30 Patient 01-02-2021	Pfizer vaccine 30-01-2021	Lymphadenopathy Injection site pain Injection site swelling Extensive swelling of vaccinated limb	1 day	Recovered 3 days	-
73	NL-LRB-00436499 F, 40-50 Patient 01-02-2021	Pfizer vaccine 29-01-2021	Chills Headache Myalgia Generalized joint pain Fatigue	2 days	Not recovered 1 day after onset	-

			Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb Pyrexia			
74	NL-LRB-00432087 F, 50-60 Patient 20-01-2021	Pfizer vaccine 16-01-2021	Malaise Fatigue Injection site pruritus Injection site warmth Injection site erythema Injection site swelling Injection site rash Extensive swelling of vaccinated limb Tachycardia Blood pressure increased Itchy rash Feeling jittery Burning sensation skin Palpitations	1 hours	Recovering 3 days after onset	Sertraline
75	NL-LRB-00436660 F, 20-30 Patient 01-02-2021	Pfizer vaccine 30-01-2021	Malaise Fatigue Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb Body temperature increased	Within 1 day	Not recovered 2 days after onset	-
76	NL-LRB-00436678 F, 30-40 Patient 01-02-2021	Pfizer vaccine 30-01-2021	Chills Myalgia Malaise Injection site erythema Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb	12 hours	Recovering 2 days after onset	-
77	NL-LRB-00436720 F, 40-50 Patient 01-02-2021	Pfizer vaccine 30-01-2021	Lymphadenopathy Chills Malaise Generalized joint pain Fatigue Injection site warmth Injection site pain Injection site swelling Extensive swelling of vaccinated limb Pyrexia	1 day	Recovering 1 day after onset	-
78	NL-LRB-00437235 M, 50-60 Patient	Pfizer vaccine 01-02-2021	Erectile dysfunction Chills Headache	Within 1 day	Not recovered 1 day after onset	-

	02-02-2021		Myalgia Malaise Injection site swelling Injection site pain Extensive swelling of vaccinated limb Pyrexia			
79	NL-LRB-00433631 M, 30-40 Patient 25-01-2021	Pfizer vaccine 24-01-2021	Syncope Injection site swelling Extensive swelling of vaccinated limb Visual accommodation disturbance of Lightheadedness	8 hours	Recovering 1 day after onset	-
80	NL-LRB-00435284 M, 20-30 Patient 29-01-2021	Pfizer vaccine 27-01-2021	Chills Myalgia Fatigue Injection site swelling Extensive swelling of vaccinated limb	2 days	Recovered 3 days	Paracetamol Paroxetine

* Originally reported as 1 minute.