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Comparison of cancer risk between neutrophilic dermatoses and plaque psoriasis patients: a cancer registry-based study

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An increased risk of developing tumors is recognized for patients affected with neutrophilic dermatoses [2–4], while this risk is less explicit for plaque psoriasis [5]. Our objective was to assess and compare, for the first time, the cancer risk between patients with neutrophilic dermatoses and those with plaque psoriasis.

This population-based, retrospective cohort study included all the patients who had received a histological diagnosis of either neutrophilic dermatoses or plaque psoriasis between 1995 and 2015 in the province of Ferrara, northern Italy. It was intended to assess the risk of cancer development based on the Ferrara Cancer Registry dataset.

Patients with prevalent cancers (diagnosed before the diagnosis of plaque psoriasis or neutrophilic dermatoses) and those without a histological confirmation of either plaque psoriasis or neutrophilic dermatosis were excluded. Multiple tumors following the first, both in cohort and in general population subjects, were not considered in risk calculation. The following indices were assessed: (i) standardized incidence ratio (SIR), (ii) individual attributable risk (AR), and (iii) attributable risk among the exposed (ARe).

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The study cohort included 278 patients (51.8% males, age at diagnosis ranging from 9 to 93 years, median 54.5), of whom 52 (18.7%) were affected with neutrophilic dermatosis and 226 (81.3%) with plaque psoriasis. The group of neutrophilic dermatoses was represented by Sweet's syndrome (7 cases), pyoderma gangrenosum (14), hidradenitis suppurativa (4), subcorneal pustulosis (7) and pustular psoriasis (20). We included pustular psoriasis within the cohort of neutrophilic dermatoses due to its immune pathway and inflammatory pattern, which are closer to those observed in these disorders than in plaque psoriasis [1].

During the follow-up (median 10.7 years, total 3005 person*years), 49 patients (17.6%) received a diagnosis of cancer (row incidence rate 1630.6×100,000, CI 95% 1206–2156), occurring from 11 months to 20 years from the diagnosis of their skin disease (median 7.6 years) (Table 1).

A first remarkable finding was that the whole cohort experienced a 31% global excess of cancer risk with respect to the age-matched general population living in the same area (Table 2).

With respect to the main study objective, the increased cancer risk among patients with neutrophilic dermatoses (SIR 1.35, CI 95% 0.7–2.4), when compared with the general population, did not differ significantly from that of patients with plaque psoriasis (SIR 1.30, CI 95% 0.9–1.8) both in univariate and multivariate Cox model.

This novel result merits a few considerations. First, it is conceivable that the systemic chronic inflammatory state present in these patients could represent a cofactor of tumor risk in both groups. If chronic inflammation can be considered a plausible background that may lead to carcinogenesis, it should also be emphasized that the inflammatory pathways of neutrophilic dermatoses are considerably different from those of plaque psoriasis. Further studies are expected both to deepen the understanding of the relationship between these inflammatory skin diseases



Table 1 Tumors diagnosed in the entire study cohort

Cancer sites	Cases n	%	Male cases, n	Female cases, n	Neutrophilic dermatoses Cases, <i>n</i>	Plaque psoria- sis Cases, n
No cancer	229	82.4	112	117	41	188
Oral cavity	1	0.4	_	1	1	_
Stomach	4	1.4	4	_	3	1
Colon	3	1.1	1	2	1	2
Liver, biliary tract	3	1.1	3	_	1	2
Lung	5	1.8	4	1	1	4
Skin (keratinocytic)	7	2.5	4	3	1	6
Breast	8	2.9	1	7	1	7
Uterus (body)	1	0.4	_	1	_	1
Prostate	2	0.7	2	_	_	2
Kidney	3	1.1	3	_	_	3
Urothelial ^a	6	2.2	5	1	1	5
Central nervous system	3	1.1	_	3	-	3
Non-Hodgkin lymphomas	1	0.4	1	_	_	1
Myelodysplastic syndromes	1	0.4	1	_	_	1
Non-specified site	1	0.4	1	_	1	_
Total	49	17.6	30	19	11	38

Bold represents the amount of these percentages

Italic represents the percentage of cases described

Table 2 Cancer incidence risk in the cohort patients

	Incident cancers, n	Person years ¹	SIR ²	CI95%	AR^3	Are (%) ⁴
Total	49	3005	1.31	0.97–1.73	9.2	57
Males	30	1423	1.48	1.0018-2.12	12.5	59
Females	19	1582	1.10	0.66-1.72	60.0	50
Diagnosis ⁵ 0–54 years	9	1162	2.14	0.98-4.07	5.5	72
55 + years	40	1843	1.21	0.86-1.64	3.1	14
Neutrophilic dermatoses	11	571	1.35	0.68-2.42	10.7	55
Plaque psoriasis	38	2435	1.30	0.92-1.79	9.6	61
Cutaneous carcinoma ⁶	7	3005	1.03	0.42-2.13	0.7	32
Female breast cancer	7	1582	1.64	0.66-3.39	2.5	58

Significant values are reported in bold

Italic is used only to distinguish percentage from absolute values

and cancers and to underline any similarities and differences between the two types of inflammatory disorders in the carcinogenic process.

From a practical perspective, a more intense cancer surveillance may be warranted in these patients. This is a somewhat consolidated practice in patients with



^aRenal pelvis, ureter, bladder, urethra (uncertain malignant potential, in situ, invasive)

¹From Ferrara Unit of Emilia-Romagna Cancer Registry 1995–2017 (70,181 incident cases; 8,113,698 p*y)

²Standardized Incidence Ratio (observed/expected*)

 $^{^{3}}$ Attributable risk × 1000

⁴Attributable risk in exposed (cohort)

⁵Age at histological diagnosis of neutrophilic dermatosis/plaque psoriasis

⁶Keratinocytic forms

neutrophilic dermatoses and should ideally be activated for psoriatic patients as well.

Author contributions All authors contributed to the study conception and design. Material preparation, data collection and analysis were performed by AB, SF and NS. The first draft of the manuscript was written by AB and SF and all authors commented on previous versions of the manuscript. All authors read and approved the final manuscript.

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Data availability The data that support the findings of this study are available from the corresponding author, N.S., upon reasonable request.

Declarations

Conflict of interest The authors declare that they have no conflict of interest.

Ethical statement Data were collected by the local Cancer Registry according to the rules of the International Agency for Research on Cancer (IARC).

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