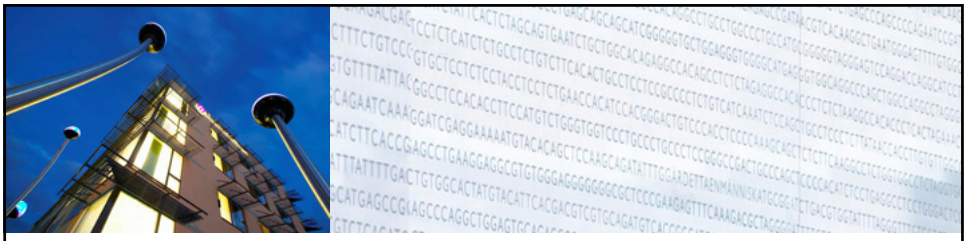

The Nordic Authority Expectations of a Successful Clinical Trial Application

Elin Karlberg
Regulatory Assessor, Medical Device
Department of Clinical Trials and Special Permissions
Medical Products Agency (Sweden)

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Council Directives

93/42/EEC on Medical Devices (MDD)
90/385/EEC on Active Implantable Medical Devices(AIMDD)
98/79/EC on In Vitro Diagnostic Medical Devices (IVDD)

Common foundation, implemented in national legislations

→ National differences

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MEDDEV guidance on clinical investigations

MEDDEV 2.7/4 Guidelines on Clinical investigations: a guide for manufacturers and notified bodies

MEDDEV 2.7/3 rev. 3 Clinical investigations: serious adverse reporting under directives 90/385/EEC and 93/42/EC

MEDDEV 2.7/2 rev. 2 Guidelines for Competent Authorities for making a validation/assessment of a clinical investigation application under directives 90/385/EEC and 93/42/EC

http://ec.europa.eu/growth/sectors/medical-devices/guidance/#meddevs

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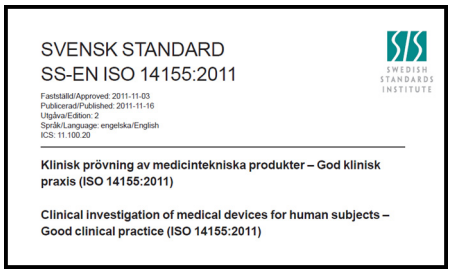
Harmonized standards

ISO 14155:2011
Clinical Investigation of medical devices for human subjects – Good clinical practice



SVENSK STANDARD
SS-EN ISO 14155:2011

SWEDISH STANDARDS INSTITUTE

Fastställd/Approved: 2011-11-03
Publicerad/Published: 2011-11-16
Utgåva/Edition: 2
Språk/Language: engelska/English
ICS: 11.100.20

Klinisk prövning av medicintekniska produkter – God klinisk praxis (ISO 14155:2011)

Clinical investigation of medical devices for human subjects – Good clinical practice (ISO 14155:2011)

ISO 14971
Medical devices – Application of risk management to medical devices

RISK ASSESSMENT MATRIX

SEVERITY / PROBABILITY	Catastrophic (1)	Critical (2)	Marginal (3)	Negligible (4)
Frequent (A)	High	High	Serious	Medium
Probable (B)	High	High	Serious	Medium
Occasional (C)	High	Serious	Medium	Low
Remote (D)	Serious	Medium	Medium	Low
Improbable (E)	Medium	Medium	Medium	Low
Eliminated (F)	Eliminated			

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Nordic Competent Authorities for Medical Devices

LÆGEMIDDELSTYRELSEN
DANISH MEDICINES AGENCY

Valvira
National Supervisory Authority
for Welfare and Health

Lyfjastofnun
Icelandic Medicines Agency

Helsedirektoratet
Norwegian Directorate of Health

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Are you ready?

Things to find out first:

- Product qualification – is it a medical device or something else?
- Product classification – what type of medical device is it?
- What essential requirements are applicable?
- Are the essential requirements met?
- Is the Clinical Evaluation outcome sufficient to support clinical performance and safety of the product?
- Do you need to make a clinical investigation?
- Has the product been CE-marked?

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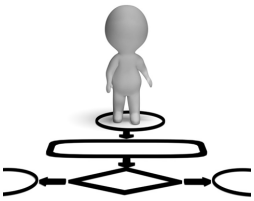
When is a notification required?



General Nordic requirement for

- Non-CE marked medical devices
- CE marked devices used outside the intended use covered by the CE- marking

Local requirements to notify CA in case of:

- IVD studies in Finland (if the above applies) and Norway.
- All AIMD studies in Finland
- In vitro diagnostic studies in Denmark - if the device is intended to come in direct or indirect contact with the human body

Note: A clinical investigation may also be a clinical trial of a medicinal product.

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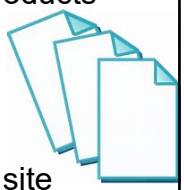
Critical components of the submission dossier

- Study documentation **compliant with ISO 14155:2011**
  - Clinical Investigation Plan
  - Investigator Brochure
  - Patient Information and Informed Consent
- **Declaration of Conformity** for investigational products
- Ethics Committee opinion
- Instructions for use
- Insurance
- Site list and CV of principal investigator for each site



NB! List above is not exhaustive

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Declaration of conformity for investigational devices

Manufacturer

Medical device under investigation

Clinical investigation plan title and ID no.

The manufacturer of the above investigational device hereby confirms that the device under investigation conforms to the essential requirements of the Medical Device Directive 93/42/EEC
apart from the aspects covered by the investigations and that, with regard to these aspects, every precaution has been taken to protect the health and safety of the patient

Dated & Signed by Manufacturer's representative

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Do you need to wait for an approval?

- Clinical investigations for products that are not
  - Class III
  - Implants
  - Invasive products for longterm use in class IIa and IIb

  may be initiated immediately after notification to the CA in Finland, Iceland and Norway, provided EC approval is in place.

- In all other cases wait for 60 days or an approval letter

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NB! Counting those 60 days …

- Note that the assessment of the application will not be initiated until the application is valid
- **60 work days** in Finland, **clockstop applies** when applicant is asked to provide additional information
- **60 calendar days** in Norway, **clockstop applies** when applicant is asked to provide additional information
- **60 calendar days** in Sweden and Denmark, **no clockstop**, normally 10 days for applicant to respond to CA questions

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CA Notification and approval requirements

Investigational device characteristics						
Device is not CE marked	*Class III *Implants *Invasive products for longterm use in class IIa and IIb	Notification + Approval	Notification + Approval	Notification + Approval	Notification + Approval	Notification + Approval
	Other	Notification + Approval	Notification only	Notification only	Notification only	Notification + Approval
Device used for another purpose than the purpose CE marked by the manufacturer	*Class III *Implants *Invasive products for longterm use in class IIa and IIb	Notification + Approval	Notification + Approval	Notification + Approval	Notification + Approval	Notification + Approval
	Other	Notification + Approval	Notification only	Notification only	Notification only	Notification + Approval
Device used for the purpose that has been CE marked by the manufacturer	AIMD		Notification + Approval			
	Other					
IVD products	In contact with body (direct or indirect)	Notification + Approval	Notification only		Notification only	
	Other		Notification only		Notification only	

Other national particulars

Review period					
60 Days	Calendar days	Work days		Calendar days	Calendar days
Clockstop	No	Yes		Yes	No
Response time	Approx 10 days	No deadline		No deadline	Approx 10 days
Study Report					
Requirement	Submit to CA	Make available	Make available	Submit to CA	Make available
Fees					
Initital submission	20.297 or 25.994 DKK depending on risk class	335 or 840 € depending on risk class	660.000 ISK	No fee	20.000 SEK
Amendment submission	5.036 DKK	No fee	150.000 ISK	No fee	No fee

DK: The submission package needs to include a Letter of authority to allow the Danish Health and Medicines Agency and the monitor to access the subject's medical records (separate from consent form)

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Tips and hints

- Review the CA website for instructions PRIOR to your submission
- Contact the CA if you have outstanding questions after the review
- Submit in format indicated by the CA. Submissions are accepted in local language or in English.
- Include all mandatory attachments with your submission
- Coordinate submissions in combination trials
- Provide additional material promptly, if requested by the CA
- If total handling time is important to you, ensure you have people available to handle the question round

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Clinical investigation of medical devices


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